

Reference Point

Separation anxiety syndrome in dogs and cats

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Distress associated with separation from a preferred companion or group has been identified among individuals from a variety of disparate species, including birds,^{1,2} dogs,^{3,4} cats,^{5,6} horses,⁷⁻¹² pigs,¹³ sheep,¹⁴ goats,^{14,15} cattle,^{16,17} cetaceans,^{18,19} and primates (including humans).²⁰⁻²³ In human medicine, the term **separation anxiety disorder (SAD)** is used to refer to excessive and recurrent distress associated with separation from home or major attachment figures.²³ It is seen primarily in children and adolescents,^{23,24} but is increasingly identified in adults of all ages.^{24,25}

Similarly, dogs may experience a negative emotional reaction when their preferred associate (favorite person or pet companion) is absent.^{3,4} Dogs that are sent to board at kennels, hospitalized at veterinary clinics, abandoned by the roadside, surrendered to shelters, or left alone at home and dogs that survive the illness or death of a pack member may suffer various degrees of anxiety.^{3,26} In dogs, the degree of distress is likely dependent, in part, on the degree of attachment to the absent figure.²⁷

Domestic cats have traditionally been viewed as asocial or even antisocial. However, a growing body of evidence suggests that cats do form social bonds²⁸⁻³² and may develop separation reactions similar to those in dogs.⁵

In people, SAD is clearly distinct from panic attacks and other severe emotional reactions.²³ In contrast, separation reactions in dogs and cats are often characterized by extreme emotional and behavioral reactions, and the degree of anxiety displayed by some dogs is consistent with the diagnostic criteria for panic attacks and other more serious psychiatric disorders in people, some of which are accompanied by intense physiologic and cognitive symptoms of fear and discomfort.²³ For example, in a person, self-mutilation would be considered a serious symptom and would prompt revision of a diagnosis of SAD to a diagnosis of an obsessive-compulsive disorder or another serious diagnostic consideration. In pets, on the other hand, separation reactions may be characterized by psychogenic grooming that results in serious self-inflicted injury.

It seems, therefore, that separation reactions in dogs and cats have much in common with phobic dis-

orders and panic attacks in people. Anxiety can escalate to panic, and coping mechanisms may fail when the apparent jeopardy of a context is inflated, as affected animals overreact to ambiguous or unpredictable situations. In 1 study,³³ the ambiguity of a situation was shown to exacerbate anxiety in children with separation issues, and this may be relevant to dogs and cats that exhibit dysfunctional separation behavior. In addition, there may be a clinical association between SAD in people and the development of anxiety disorders,²⁴ and there is some evidence that suggests that this may be true in dogs as well.³⁴

Some elements of separation reactions in dogs and cats may share similarities with aspects of bipolar affective disorders in people.²³ People with bipolar affective disorders experience cycles of manic-depressive illness that may last for days or months, during which their mood shifts dramatically between agitation and lethargy. Similarly, during a separation period, a dog or cat with a severe separation reaction may oscillate between relative manic (agitated) states (eg, episodes of vocalization and destructiveness) and depressive (subdued) states.

Clearly, the signs of separation anxiety in dogs and cats are unique and frequently more insidious, compared with the relatively milder psychiatric symptoms that define SAD in people. Therefore, a diagnostic label of separation anxiety in dogs and cats is inadequate, because it does not encompass the range and intensity of behaviors exhibited by emotionally reactive pets. Given the complexity and diversity of the clinical and subclinical signs related to separation reactions in veterinary patients, the term of **separation anxiety syndrome (SAS)** has been introduced.⁵

Separation anxiety syndrome in dogs and cats represents a group of emotional, behavioral, and physiologic responses that vary in intensity and clinical appearance. This redefinition highlights new insights into the condition, and the present review attempts to examine more closely how sociality and domestication in dogs and cats may be a precursor to the development of the dysfunctional social ties that characterize SAS. Parallels will be drawn between the overdependence of pets with SAS and opioid drug dependence. As pets live longer, the incidence of SAS in geriatric pets is predicted to rise, and this condition will have to be distinguished from other ailments. Finally, grief in pets is discussed for the first time as a unique form of SAS.

Sociality in Dogs and Cats

To the best of our knowledge, separation reactions are not a characteristic of asocial species, for without

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the proclivity toward formation of social bonds, separation-related anxiety would not exist.³⁵ By definition, individuals that belong to social species are sensitive to the absence of an attachment figure.^{2,3,5,7-9,12-17,19,20-22} Individual experiences, inborn emotionality, degree of attachment, and species-specific sociality determine the specific signs and the intensity of those signs in animals with separation reactions.

In essence, therefore, development of separation-related distress can be viewed as a measure of sociality (ie, the tendency of individuals to associate or assemble in groups). Sociality is a plastic phenomenon that is, in part, neurologically programmed in dogs and cats. Puppies are most susceptible to social exposure when they are between 6 and 13 weeks old.³⁶ A similar phenomenon is identified in kittens between 2 and 7 weeks old.³² By 8 weeks of age, puppies and kittens are increasingly defensive, and as the CNS matures, social interactions become more tentative.³⁶

Puppies and kittens can thrive only when raised with nurturing social interaction. Puppies raised in total isolation, for example, develop abnormal responses to new environments and significant increases in the threshold to pain response.³⁶ Kittens raised in isolation may become more aggressive. In fact, early separation from the queen can result in fear of, or aggressiveness toward, cats and people, learning disabilities, increased activity, and increased predatory behavior.^{6,36} Early human handling helps to produce more tame kittens and puppies that are more adaptive to change (stress) throughout their lifetimes. Just a few minutes a day is all that is necessary, although kittens or puppies < 2 weeks old should not be overly handled.^{32,36}

The social behavior of dogs reflects their descent from wolves, which have a highly structured social organization,³⁷⁻³⁹ and the domestication of dogs was likely facilitated by intrinsic social parallels with people. In both wolves and people, for instance, group cohesion and cooperation are essential to individual survival and species success. Among wolves and dogs, establishment of a dominance hierarchy defines the social structure of the pack. However, emotional attachments between pack members is the glue that cements and solidifies the social unit. A recent study⁴⁰ suggests that during the process of domestication, dogs may have developed particular social-cognitive abilities that enable them to be more skillful at reading human communicative signals. These skills surpass those of the great apes and wolves and may be apparent in puppies that are only a few weeks old and have had limited human contact.⁴⁰

Social behavior in domestic cats is quite complex. Hierarchies within a group of cats are formed with subtle and sometimes not so subtle displays of dominance and submission^{28,31,32} and with multifaceted social communication through facial expression,³² body posturing,³² and vocal communication.^{32,41} Affiliative behavior, which facilitates proximity and intimate social interaction, is common among domestic cats.²⁸ Preferential alliances (what humans might call friends) are consistently seen and reflected by a cat's choice of partners for sleeping; resting, sitting, and playing together; sharing food; allogrooming; and allorubbing. Members

of a stable social group of cats have been shown to touch each other as much as half the time. Females may cooperate in rearing their young.²⁸ Most pet cats engage in frequent social interactions with conspecific housemates, as well as with their owners. In fact, 1 study²⁹ showed that most cats slept with a family member, shared food with people, and greeted their owners at the door. Sociality toward people has been shown to be governed by learned and inherited mechanisms.³⁰

Perhaps one of the major differences between cats and dogs is that dogs generally require ongoing social interaction. Although cats are sociable by nature, as adults they do not require human or feline contact to survive. In addition, there is much wider individual variation in sociality among cats than among dogs, in that some cats are indeed solitary and others more socially dependent. This social variability may be explained in part by the fact that cats have not been systematically domesticated, compared with dogs.

Additional insight into the social differences between domestic cats and dogs stems from the observation that social systems may be flexible and frequently depend on environmental resources (eg, abundance of food) and predatory pressures.³⁵ Domestic cats have apparently developed the social skills necessary to cope with fluctuations in population density. For instance, when food is abundant in a restricted but desirable space, cats will tolerate crowding, and aggression may be inhibited.^{28,29} Social pressures on outdoor cats are driven by the availability of food, the population density of other outdoor cats, and the presence of various dangers. Adjustable group size and sociality are recognized in a variety of other highly social species, including apes⁴² and cetaceans.^{43,44} This facultative sociality of domestic cats may help explain the greater range in social affinity among adult cats, compared with the more consistent and, perhaps, obligate sociality of domestic dogs, which have a greater dependence on group cohesion. The inherent sociality of dogs and cats is fundamental to SAS and, ultimately, to the prevention and treatment of separation reactions.

Adaptive and Maladaptive Separation Reactions in Dogs and Cats

In puppies and kittens, some separation reactions are normal and desirable, as they are important to survival. Young puppies and kittens, for instance, normally emit distress calls when removed from the nest. In fact, a queen will not retrieve a straying kitten unless the lost kitten vocalizes.³²

Newborn puppies and kittens first experience separation when the bitch or queen temporarily leaves the nest or when they are removed from the litter and held by human hands. Soon after, these puppies and kittens must shift their normal social attachments from littermates and the dam to their new human owners. Mild signs of SAS, therefore, may be normal in newly adopted young pets as they adjust to separation from their littermates and dam.

When these separation reactions are retained beyond the puppy or kitten period, however, they may become maladaptive. In juvenile and adult dogs and cats with SAS, in particular, the distress expressed

often seems excessive in context with the actual danger of the situation. The process of domestication appears to have resulted in retention of juvenile behaviors in adults, and the dependence of the young on parental figures is a characteristic of juvenile behavior. Kneading behavior in adult cats that is directed toward a preferred human companion is one example. Although there may be some variability in canine attachment behavior to humans, the canine-human bond is thought to most resemble the mother-infant interaction in people.⁴⁵ This intimate and mutual bond is part of what makes pet ownership so gratifying and emotionally satisfying. It may also be what predisposes dogs and cats to the dysfunctional separation reactions that characterize SAS. Both cats and dogs may retain juvenile characteristics of separation distress in the form of a predisposition toward neurotic overdependence on their human owners as part of a genetic by-product of domestication.

Overdependence may be reinforced unintentionally by pet owners. For example, pets may become unused to separation if they are not given the opportunity to learn to cope with time away from their caretakers.⁴⁶ Pet owners may be reluctant to leave young pets and, thereby, unintentionally encourage neurotic social attachment.

Behavioral Aspects of SAS

Sociality is fundamental to the behaviors associated with SAS. The clinical signs of SAS often reflect a pet's frustration with and anxiety caused by social separation and, specifically, separation from an important attachment figure. Separation-related behavior problems in dogs and cats vary widely in form and degree, even in the same individual. Clinical signs are seen primarily during an owner's absence, although anxious behaviors any time during the perideparture period may be attributed to SAS. For instance, anticipatory distress is common in the period preceding the owner's actual departure.^{3,4} Dogs learn the habitual sequence of events leading to an impending departure, and anxiety often begins before the owner leaves. Persistent shadowing or following of the owners and extremely excited greeting of a returning owner are associated with SAS in dogs.

Signs of SAS in dogs include agitation (eg, restlessness, pacing, and jumping); physiologic abnormalities (eg, vomiting, salivating, trembling, hyperventilating, and tachycardia); elimination abnormalities (eg, urination and defecation, with or without diarrhea); vocalization (eg, whining, whimpering, barking, and howling); destruction (eg, escape attempts such as chewing, digging, scratching, and jumping through doors or windows); self-mutilation (eg, excessive grooming, hair pulling, and self-directed aggression); signs of depression (eg, social withdrawal, lethargy, inappetence, and submissive or fearful postures or facial expressions); and aggression (eg, mouthing, nipping, growling at, or biting the owner at departure). Signs are typically most intense within the first 15 minutes after the owner's departure, although they may persist much longer or occur intermittently during isolation.^{3,4} They may be seen alone or in any combination and may progress in intensity or change in appearance

over time. For example, agitation may be the only sign or may simply be the first or last in a sequence of other SAS-related signs, such as inappropriate elimination in the form of diarrhea or urination. Signs of depression (eg, inactivity and social withdrawal) may be the only indication of SAS in some dogs or may follow a period of agitated destructiveness and excessive vocalization in others. Signs of SAS must also be differentiated from similar signs in dogs and cats with other disorders.^{3,5,46}

There are 3 primary categories of misbehavior in cats and dogs with SAS^{3,5,46}: inappropriate elimination, vocalization, and destructiveness. Inappropriate elimination may be intentional (eg, territorial marking to release anxiety) or an involuntary physiologic effect of extreme distress (eg, stress-induced diarrhea or urination). In 1 study,⁵ house soiling was the most common problem in cats with SAS. In this study, inappropriate defecation was more common in neutered female cats than in neutered male cats. Seventy-five percent of cats that urinated inappropriately in association with the absence of an attachment figure did so exclusively on the owner's bed. Excessive or persistent vocalization, typically attention-seeking distress calls, is reported in both dogs and cats with SAS. Destructiveness among dogs with suspected SAS may include desperate escape attempts made during confinement to a crate or restricted space. In such cases, this barrier anxiety may be similar to claustrophobia (which can occur independently of separation issues) and should be distinguished from separation-triggered misbehavior. In cats with SAS, destructiveness was observed in neutered males but not in neutered females.⁵ However, not all destructive behavior that occurs in a owner's absence is due to a separation reaction. Accidental destruction of property by an unsupervised playful and inquisitive pet must be included in the differential diagnosis of destructiveness.^{3,5,46}

Other less common signs of SAS are also recognized.^{3,46} Self-mutilation, such as psychogenic licking or compulsive tail chewing, for instance, may be associated with separation reactions in both dogs and cats. In addition, dogs may injure themselves during attempts to push, chew, and scratch their way out of a confining enclosure. Puncture and laceration wounds may result from protruding metal wires or sharp edges of damaged crates. In these cases, injury is not self-inflicted but rather a consequence of destruction. Psychogenic overgrooming appears to be more common in female cats with SAS than in males and may be underreported.⁵

Another less frequent, but not unknown, sign of SAS is aggressiveness.³ Anticipatory anxiety in some dogs may be so intense that they begin to nip or bite, apparently to disrupt the owner's preparations to depart or prevent their approach to the door. Similar behavior associated with an owner's impending absence has not been identified in cats. Nonetheless, cats can learn to nip to gain attention; therefore, it would not be inconceivable that cats, like dogs, with SAS would act aggressively toward their owners as they exit the home.

The frequently progressive and often intense signs of separation distress displayed by some dogs and cats

may have much in common with drug dependence and acute withdrawal from opioids. Beyond the genetic and learned components of SAS, there may be an underlying biochemical mechanism that may have parallels to opioid drug dependence. This is supported by the finding that some signs of SAS may be alleviated with opiates. Low doses of oxymorphone and morphine were shown to profoundly reduce crying in briefly isolated puppies,⁴⁷ and physiologic measures of separation distress increased slightly in cattle treated with the opiate antagonist naloxone.¹⁷ Thus, it is possible that opiate agonists might be effective in the treatment of SAS in dogs and other species.

Predisposing Factors for SAS

Common risk factors associated with SAS in dogs include a history of traumatic separation, inexperience with being left alone, excessive greetings and prolonged departures by owners, changes in the owner's routine (dogs susceptible to SAS may develop clinical signs in response to even slight changes in routine), relocation to a new home, addition of a new pet sitter, and a death in the family (conspecific or other family member). In addition, dogs that shadow or follow their owners or that are neutered, purebred, or have a single owner are more likely to develop SAS.^{3,46,48,49} One study⁵⁰ suggested that purebred dogs may respond better to treatment than dogs of mixed breeding, although this deserves further investigation.

Risk factors for SAS in cats⁵¹ appear to be consistent with risk factors in dogs and include having an owner who works long hours, a change (new schedule or extension) in the owner's work schedule, frequent business or vacation travel by the owner, an increase in the time the owner spends with family or friends, development of a new romantic involvement for the owner, departure of a family member (eg, through divorce or death), and death or abrupt removal of another pet. In addition, cats with a history of shelter adoption after 3 months of age, cats that follow their owners around the house, neutered cats, and cats owned by a single individual are more likely to develop SAS. Additional investigation into SAS in cats is warranted.

Unique Manifestations of SAS in Dogs and Cats

SAS in geriatric patients—Older pets, like older people, may be less adaptive to change and stress in general. In particular, normal senile changes may reduce the ability of aging pets to cope with separation from attachment figures and removal from or changes in familiar surroundings. As pets live longer than ever before, SAS may become more common in geriatric patients. Underlying medical problems can reduce an aging pet's emotional adaptability and cognitive capacity; therefore, underlying medical conditions should be considered in older pets with behavioral disturbances, including SAS.^{52,53} In my experience, for example, a sudden onset of SAS in middle-aged dogs may be associated with hypothyroid disease. Metabolic diseases and other physical ailments (eg, deafness, blindness, and arthritis) must be considered when aging pets

show behavioral changes. One study⁵² reported that concurrent age-related physical ailments, including, but not limited to, degenerative joint disease and renal disease, were likely to exacerbate behavior problems. However, the same study also determined that behavior problems, including SAS, frequently occur in older pets in the absence of any health issues. Nevertheless, SAS can emerge in older dogs without any previous history of separation reactions.

Old dogs with behavior problems, including SAS, often benefit from behavioral therapy.⁵² In cats, SAS may be more common among aging females than males, possibly because of sex-based differences in feline social behavior.⁵ A thorough physical evaluation, including laboratory analysis of blood and urine, remains an important part of the workup of cats and dogs with signs of SAS, regardless of age.

Cognitive dysfunction must be ruled out in older dogs and cats suspected of having SAS because of the overlapping clinical features of these disorders. Hypervocalization, nonresponsiveness, disorientation, and changes in the sleep-wake cycle and feeding patterns may be suggestive of cognitive dysfunction syndrome in dogs and must be distinguished from signs associated with SAS.⁵⁴⁻⁵⁷ A clinical entity resembling cognitive dysfunction syndrome has been identified in geriatric cats,⁵⁸ and amyloid plaques similar to the senile neural plaques that characterize Alzheimer's disease in humans and cognitive dysfunction syndrome in dogs have been identified in cats with signs of cognitive dysfunction.⁵⁶ In cats and dogs, cognitive dysfunction syndrome has been distinguished from SAS and other clinical considerations and may respond temporarily to treatment with selegiline.

Grief as a form of SAS in dogs and cats—Grief can be viewed as the most extreme separation reaction among social animals. Public mourning and private expressions of grief are considered normal human responses following the death of a family member or friend,²³ and feelings of sadness, insomnia, weight loss, and decreased appetite are recognized symptoms of loss in people. The stages of grief in humans have been defined⁵⁹ and reflect the cognitive and emotional process of coping with death.

Unlike people, it is unclear whether animals are able to cognitively process the concept of death. Regardless, it is not strictly important whether dogs and cats cognitively understand that a preferred companion has died or been permanently or temporarily removed. They may develop signs of emotional distress simply as a result of the absence of this individual. Separation anxiety syndrome is a reaction to separation itself. Thus, dogs and cats may indeed exhibit behaviors that reflect the emotional impact of the death of a close companion.

Separation reactions in dogs and cats may be exacerbated by the moods of their human caretakers who may be mourning the death of the same person or pet. Signs of lethargy or agitation, social withdrawal or uncharacteristic overdependence, anorexia, and other changes may develop. In addition, stress can trigger latent or subclinical disease. Signs of profound or per-

sistent depression and anxiety may justify referral to a board-certified veterinary behaviorist and the use of psychoactive medications, but most dogs and cats adjust to the death of an attachment figure within several weeks.

Treatment of SAS

Treatment of SAS in dogs and cats should focus on managing clinical signs in the individual patient and should consider the pet's needs for social interaction, intellectual stimulation, and exercise.^{3,5,46} Pets whose emotional, psychological, and physical needs are met regularly and reliably will be less prone to develop the dysfunctional overdependence and anxious misbehavior that characterize SAS. Increased attention in the form of interactive play, petting, and grooming should be scheduled such that the caretaker's planned absence coincides with the pet's desire to rest. In newly introduced dogs, the progression of initial mild signs of SAS (due to separation from littermates, dam, or previous owners) or sensitization to SAS that may emerge later in life may be reduced by providing bedding near the new owners overnight (social animals naturally remain together for safety while sleeping).

Events that occur during the perideparture period (ie, the period just before, during, and after the departure of the preferred associate) can reinforce a pet's tendency toward SAS and, therefore, should be as calm as possible. For example, greeting rituals are normal social behaviors in both dogs and people. The exuberance of some human greetings, however, may create conflict and reinforce undesirably chaotic greetings by the dog. Overly enthusiastic greetings (eg, whining, jumping, or pacing) laden with redirected (eg, scratching) or displaced (eg, running to get a toy) activities are typical of dogs with SAS.^{3,48} In addition, long departure rituals characterized by a melodramatic or anxious attitude on the part of the owner may cause dogs and cats to view the owner's departure as an anxiety-provoking event, exacerbating the degree of anxiety during separation.

In dogs, departures should be preceded with a long leash walk appropriate to the individual's age and level of activity and to prevailing weather conditions. Presentation of an attractive chew toy, such as a rawhide bone or hollow toy filled with treats, promotes a positive association with departure and provides an appropriate oral outlet for anxiety. Disruption of a predictable departure sequence and progressive desensitization to the owner's habitual departure routine are also helpful.^{46,50} Some dogs with SAS may also have other anxiety-related issues, such as thunderstorm phobia.³⁴ Thus, potential causes of anxiety must be thoroughly explored, and the treatment plan must address all important causes.

Confined house cats have a restricted territory and, consequently, reduced options for physical and intellectual stimulation. Cat owners should compensate by providing additional distractions, such as interactive play with attractive toys, and many of the same activities suggested for dogs. Targets of SAS-associated elimination should be made inaccessible or unattractive. The treatment of house soiling in cats is described in detail elsewhere.⁶⁰⁻⁶³

The addition of another pet to keep a pet with SAS company is not necessarily a solution to the problem. It is thought that the pet's emotional response is caused by the absence of a particular individual and not social isolation as such. Tuber et al⁶⁴ measured the behavioral and glucocorticoid responses of adult domestic dogs in a shelter situation following removal of a kennel mate and placement in a novel environment for 4 hours. Measures of stress were unaffected by the kennel mate's presence or absence; however, stress was reduced by the presence of a human caretaker. This study suggests that human companionship may be more important to dogs and may point to a difference in the social relationship of dogs toward people.⁴⁰ However, these results should be interpreted cautiously, because glucocorticoid responses to separation may be more reflective of degree of attachment. The absence of an effect following separation from an affiliative conspecific may be attributable to differences in separation from partners that are merely affiliative, compared with separation from partners to which one is emotionally attached.²⁷

Anecdotal evidence suggests that in some cases, the addition of a companion animal may improve an anxious dog's ability to cope with separation from its owner. However, addition of a new pet may make the situation worse if it draws the owner's time and attention away from the already anxious resident pet. New housemates must be gradually introduced, despite the temptation to accelerate the process, particularly when the new pet is intended to ease anxiety in a resident pet. For both cats and dogs, conflicts between new housemates are not unusual. Although such conflicts typically resolve, some pets are incompatible. Thus, the benefits and risks should be carefully considered before adding a new pet to a household with a dog or cat with SAS.

The use of psychoactive medication may be necessary for treatment of SAS in dogs and cats if behavior modification alone is unsuccessful. In dogs in particular, psychoactive medication may be needed to control signs of SAS. Dog owners may be less likely to comply with techniques that require more time and effort, such as desensitizing affected dogs to departure cues, and may be more likely to follow simple behavior modifying techniques, such as providing a chew toy at departure.⁵⁰

In dogs and cats with acute or extreme SAS, such as a reaction to the death of an attachment figure or substantial destruction of property, use of a medication with predictable effects and a rapid onset is important. Alprazolam and other benzodiazepines are rapidly absorbed.⁶⁵ In cats, triazolam is an attractive alternative to diazepam, which has previously been associated with adverse effects, although no problems have been recently reported.⁶⁶ Tricyclic antidepressants may require several weeks to take effect⁶⁵ and may be more appropriate for long-term management. Clomipramine has been successful for the treatment of SAS in dogs.⁶⁷⁻⁶⁹ However, in 1 study,⁶⁹ administration of clomipramine without concomitant behavioral treatment was unsuccessful in controlling separation-related behavior problems, whereas behavior modification alone was successful in reducing the severity of clinical signs. Psychoactive medication may be an important comple-

ment to behavioral treatment, but is unlikely to replace the need for modification of the pet's lifestyle, which is fundamental to the problem.

A CBC, serum biochemical profile, urinalysis, and thyroid function tests are recommended prior to administration of psychoactive drugs, both to identify underlying problems and to provide baseline values in the case of long-term drug administration. Laboratory tests should be repeated periodically during prolonged drug treatment to detect adverse effects.

References

1. Sufka KJ, Hughes RA. Differential effects of handling on isolation-induced vocalizations, hypoalgesia, and hyperthermia in domestic fowl. *Physiol Behav* 1991;50:129-133.
2. Davis CS. Parrot psychology and behavior problems. *Vet Clin North Am Small Anim Pract* 1991;21:1281-1288.
3. Borchelt PL, Voith VL. Diagnosis and treatment of separation-related behavior problems in dogs. *Vet Clin North Am Small Anim Pract* 1982;12:625-635.
4. Lund JD, Jorgensen MC. Behaviour patterns and time course of activity in dogs with separation problems. *Appl Anim Behav Sci* 1999;63:219-236.
5. Schwartz S. Separation anxiety syndrome in cats: 136 cases (1991-2000). *J Am Vet Med Assoc* 2002;220:1028-1033.
6. Guyot GW, Bennett TL, Cross HA. The effects of social isolation on the behavior of juvenile domestic cats. *Dev Psychobiol* 1980;13:317-329.
7. Mal ME, Friend TH, Lay DC, et al. Behavioral responses of mares to short-term confinement and social isolation. *Appl Anim Behav Sci* 1991;31:13-24.
8. Mal ME, Friend TH, Lay DC, et al. Physiological responses of mares to short-term confinement and social isolation. *Equine Vet Sci* 1991;11:96-102.
9. Strand SC, Tiefenbacher S, Haskell M, et al. Behavior and physiologic responses of mares to short-term isolation. *Appl Anim Behav Sci* 2002;78:145-157.
10. McClure SR, Chaffin MK, Beaver BV. Nonpharmacologic management of stereotypic self-mutilative behavior in a stallion. *J Am Vet Med Assoc* 1992;200:1975-1977.
11. Alexander SL, Irvine CH, Livesey JH, et al. Effect of isolation stress on concentrations of arginine, vasopressin, alpha-melanocyte-stimulating hormone and ACTH in the pituitary venous effluent of the normal horse. *J Endocrinol* 1988;116:325-334.
12. Houpt KA, Houpt TR. Social and illumination preferences of mares. *J Anim Sci* 1989;66:2159-2164.
13. Schrader L, Ladewig J. Temporal differences in the response of the pituitary adrenocortical axis, the sympathoadrenomedullary axis, heart rate, and behaviour to a daily repeated stressor in domestic pigs. *Physiol Behav* 1999;66:775-783.
14. Lyons DM, Price EO, Moberg GP. Social grouping tendencies and separation-induced distress in juvenile sheep and goats. *Dev Psychobiol* 1993;26:251-259.
15. Carbonaro DA, Friend TH, Dellmeier GR, et al. Behavioral and physiological responses of dairy goats to isolation. *Physiol Behav* 1992;51:297-301.
16. Boissy A, Le Neindre P. Behavioral, cardiac and cortisol responses to brief peer separation and reunion in cattle. *Physiol Behav* 1997;61:693-699.
17. Rushen J, Boissy A, Terlouw EM, et al. Opioid peptides and behavioral and physiological responses of dairy cows to social isolation in unfamiliar surroundings. *J Anim Sci* 1999;77:2918-2924.
18. McCowan B, Reiss D. The fallacy of "signature whistles" in bottlenose dolphins: a comparative perspective of "signature information" in animal vocalizations. *Anim Behav* 2001;62:1151-1162.
19. Janik VM, Slater PJ. Context-specific use suggests that bottlenose dolphin signature whistles are cohesion calls. *Anim Behav* 1998;56:829-838.
20. Laudenslager ML, Held PE, Boccia ML, et al. Behavioral and immunological consequences of brief mother-infant separation: a species comparison. *Dev Psychobiol* 1990;23:247-264.
21. Gust DA, Gordon TP, Brodie AR, et al. Effect of a preferred companion in modulating stress in adult female rhesus monkeys. *Physiol Behav* 1994;55:681-684.
22. Norcross JL, Newman JD. Effects of separation and novelty on distress vocalizations and cortisol in the common marmoset (*Callithrix jacchus*). *Am J Primatol* 1999;47:209-222.
23. *DSM-IV: diagnostic and statistical manual of mental disorders*. 4th ed. Washington, DC: American Psychiatric Association, 1994;110-113, 328-362, 394-395, 684.
24. Manicavasagar V, Silove D, Franz CP, et al. Continuities of separation anxiety from early life into adulthood. *J Anxiety Disord* 2000;14:1-18.
25. Wijeratne C, Manicavasagar V. Separation anxiety in the elderly. *J Anxiety Disord* 2002;in press.
26. Hennessy MB, Davis HN, Williams MT, et al. Plasma cortisol levels of dogs at a county animal shelter. *Physiol Behav* 1997;62:485-490.
27. Hennessy MB. Hypothalamic-pituitary-adrenal responses to brief social separation. *Neurosci Biobehav Rev* 1997;21:11-29.
28. Crowell-Davis SL, Barry K, Wolfe R. Social behavior and aggressive problems of cats. *Vet Clin North Am Small Anim Pract* 1997;27:549-568.
29. Bernstein PL, Strack M. A game of cat and house: spatial patterns and behavior of 14 domestic cats in the home. *Anthrozoos* 1996;9:25-39.
30. Reisner IR, Houpt KA, Erb HN, et al. Friendliness to humans and defensive aggression in cats: the influence of handling and paternity. *Physiol Behav* 1994;55:1119-1124.
31. Voith VL, Borchelt PL. Social behavior of domestic cats. *Compend Contin Educ Pract Vet* 1986;8:637-645.
32. Bradshaw J. *The behavior of the domestic cat*. New York: CABI Publishing, 1998;68-69, 92-110, 140-162, 169-171.
33. Bogels SM, Zigterman D. Dysfunctional cognitions in children with social phobia, separation anxiety disorder, and generalized anxiety disorder. *J Abnorm Child Psychol* 2000;28:205-211.
34. Overall KL, Dunham AE, Frank D. Frequency of nonspecific clinical signs in dogs with separation anxiety, thunderstorm phobia, and noise phobia, alone or in combination. *J Am Vet Med Assoc* 2001;219:467-473.
35. Wilson EO. *Sociobiology*. Cambridge, Mass: Harvard University Press, 1980;20-31.
36. Houpt KA, Wolski TR. *Domestic animal behavior for veterinarians and animal scientists*. Ames, Iowa: The Iowa State University Press, 1982;189-201.
37. Vila C, Savolainen P, Maldonado JE, et al. Multiple and ancient origins of the domestic dog. *Science* 1997;276:1687-1689.
38. Savolainen P, Zhang YP, Luo J, et al. Genetic evidence for an East Asian origin of domestic dogs. *Science* 2002;298:1610-1613.
39. Simpson BS. Canine communication. *Vet Clin North Am Small Anim Pract* 1997;27:445-464.
40. Hare B, Williamson C, Tomasello M. The domestication of social cognition in dogs. *Science* 2002;298:1634-1636.
41. Moelk M. Vocalizing in the house cat; a phonetic and functional study. *Am J Psychol* 1944;57:184-205.
42. Goodall J. *The chimpanzees of Gombe. Patterns of behavior*. Cambridge, Mass: Harvard University Press, Belknap Press, 1986; 673.
43. Connor RC, Smolker RA, Richards AF. Aggressive herding of females by coalitions of male bottlenose dolphins (*Tursiops* sp) In: Harcourt AH, de Waal FBM, eds. *Coalition and alliances in humans and other animals*. Oxford, England: Oxford University Press, 1992;415-443.
44. Reiss D, McCowan B, Marino L. Communicative and other cognitive characteristics of bottlenose dolphins. *Trends Cogn Sci* 1997;1:140-145.
45. Topal J, Miklosi A, Csanyi V, et al. Attachment behavior in dogs (*Canis familiaris*): a new application of Ainsworth's (1969) strange situation test. *J Comp Psychol* 1998;112:219-229.
46. McCrave EA. Diagnostic criteria for separation anxiety in the dog. *Vet Clin North Am Small Anim Pract* 1991;21:247-255.
47. Panksepp J, Herman B, Conner R, et al. The biology of social attachments: opiates alleviate separation distress. *Biol Psychiatry* 1978;13:607-618.

48. Flannigan G, Dodman NH. Risk factors and behaviors associated with separation anxiety in dogs. *J Am Vet Med Assoc* 2001;219:460-466.

49. Patronek GJ, Glickman LT, Beck AM, et al. Risk factors for relinquishment of dogs to an animal shelter. *J Am Vet Med Assoc* 1996;209:572-581.

50. Takeuchi Y, Houpt KA, Scarlett JM. Evaluation of treatments for separation anxiety in dogs. *J Am Vet Med Assoc* 2000;217:342-345.

51. Patronek GJ, Glickman LT, Beck AM, et al. Risk factors for relinquishment of cats to an animal shelter. *J Am Vet Med Assoc* 1996;209:582-588.

52. Chapman BL, Voith VL. Behavioral problems in old dogs: 26 cases (1984-1987). *J Am Vet Med Assoc* 1990;196:944-946.

53. Reisner I. The pathophysiologic basis of behavior problems. *Vet Clin North Am Small Anim Pract* 1991;21:207-224.

54. Cummings BJ, Head E, Ruehl WW, et al. The canine as an animal model of human aging and dementia. *Neurobiol Aging* 1996;17:259-268.

55. Borras D, Ferrer I, Pumarola M. Age-related changes in the brain of the dog. *Vet Pathol* 1999;36:202-211.

56. Cummings BJ, Satou T, Head E, et al. Diffuse plaques contain C-terminal A β ₄₂ and not A β ₄₀: evidence from cats and dogs. *Neurobiol Aging* 1996;17:653-659.

57. Dimakopoulos AC, Mayer RJ. Aspects of neurodegeneration in the canine brain. *J Nutr* 2002;132:1579S-1582S.

58. Houpt KA. Cognitive dysfunction in geriatric cats. In: August JR, ed. *Consultations in feline internal medicine*. 4th ed. Philadelphia: WB Saunders Co, 2001;584-591.

59. Kubler-Ross E. *On death and dying*. New York: MacMillan Publishing Co, 1979;38-137.

60. Simpson BS. Feline house soiling. Part I. Inappropriate elimination. *Compend Contin Educ Pract Vet* 1998;20:1319-1329.

61. Simpson BS. Feline house soiling. Part II. Urine and fecal marking. *Compend Contin Educ Pract Vet* 1998;20:1331-1349.

62. Pryor PA, Hart BL, Bain MJ. Effects of a selective serotonin reuptake inhibitor on urine spraying behavior in cats. *J Am Vet Med Assoc* 2001;219:1557-1561.

63. Pryor PA, Hart BL, Bain MJ, et al. Causes of urine marking in cats and effects of environmental management on frequency of marking. *J Am Vet Med Assoc* 2001;219:1709-1713.

64. Tuber DS, Sanders S, Hennessy MB, et al. Behavioral and glucocorticoid responses of adult domestic dogs (*Canis familiaris*) to companionship and social separation. *J Comp Psychol* 1996;110:103-108.

65. Overall K. Pharmacologic treatments for behavior problems. *Vet Clin North Am Small Anim Pract* 1997;27:637-664.

66. Center SA, Elston TH, Rowland PH, et al. Fulminant hepatic failure associated with oral administration of diazepam in 11 cats. *J Am Vet Med Assoc* 1996;209:618-625.

67. King JN, Simpson BS, Overall KL, et al. Treatment of separation anxiety in dogs with clomipramine: results from a prospective, randomized, double-blind, placebo-controlled, parallel-group, multicenter clinical trial. *Appl Anim Behav Sci* 2000;67:255-275.

68. Seksel K, Lindeman MJ. Use of clomipramine in treatment of obsessive-compulsive disorder, separation anxiety and noise phobia in dogs: a preliminary clinical study. *Aust Vet J* 2001;79:252-256.

69. Podberscek AL, Hsu Y, Serpell JA. Evaluation of clomipramine as an adjunct to behavioural therapy in the treatment of separation-related problems in dogs. *Vet Rec* 1999;145:365-369.



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