

CANINE VACCINATION GUIDELINES

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There are two sets of canine vaccination guidelines available: those produced by the American Animal Hospital Association [1] and those from the WSAVA Vaccination Guidelines Group (VGG) [2]. The fundamental principle of both sets of guidelines, as encapsulated by the VGG, is that *'we should aim to vaccinate every animal with core vaccines and to vaccinate each individual less frequently by only giving non-core vaccines that are necessary for that animal'*. This presentation aims to explain how this statement relates to canine vaccination.

It is firstly necessary to understand the definitions of 'core' and 'non-core' vaccines. CORE vaccines are those that all animals should receive to protect them against diseases of global significance or where legislation may dictate [i.e. canine rabies]. The use of NON-CORE vaccines is dictated by geographical location, lifestyle and exposure risk. Some vaccines are NOT RECOMMENDED because there is little scientific justification for their use.

For dogs, the core vaccines are those that protect against canine distemper virus (CDV), canine adenovirus (CAV) and canine parvovirus-2 (CPV). In any country in which rabies is an endemic disease, then rabies vaccination is also considered core for dogs. Non-core canine vaccines include those that protect against leptospirosis, canine parainfluenza virus (CPI), *Bordetella bronchiseptica* and *Borrelia*. Canine coronavirus (CCV) vaccine is not recommended as there is little evidence that CCV is a primary enteric pathogen or that the vaccine can protect against such infection.

WSAVA guidelines provide generic advice to practitioners, but it is impossible to ensure that the guidelines are tailored to best fit the local situation in each of the 76 WSAVA member countries. The VGG encourages national associations to adapt and modify the guidelines for local use where appropriate. This process might involve altering the

classification of a vaccine. For example, in the UK, *Leptospira* vaccine is considered core for the dog and attempts are now being made to provide data that define disease prevalence^[3].

CORE VACCINATION

The vaccination of puppies is determined by the transfer of maternally-derived antibody (MDA) from the bitch in colostrum. This antibody is crucial for protection of the pup during early life, but simultaneously blocks the endogenous immune response of the puppy to vaccination. Canine immunoglobulin has a half life of around 11 days and there is progressive decline in MDA concentrations in pups over the first weeks of life. The 'window of susceptibility' occurs when there is no longer sufficient maternal antibody to provide full protection from infectious disease, but where sufficient antibody remains to block the ability of the pup to make its own immune response to modified live virus (MLV) vaccine. Traditionally, this window has been taken to occur at between 8 – 10 weeks of age, but new evidence shows that higher titre vaccines increase maternal antibody concentrations leading to persistence of MDA for longer periods of time. Studies have now shown that around 1 in every 10 puppies has 'blocking' levels of MDA at 12 weeks of age. For this reason, vaccination guidelines now recommend that puppy vaccination (with MLV core vaccines) starts at 8 – 9 weeks of age, with a second vaccine 3 – 4 weeks later and a third vaccine given between 14 – 16 weeks of age (preferably at 16 weeks), to ensure that all pups have received at least one dose of vaccine in the absence of interfering MDA. A 12 month booster vaccine (either at 12 months of age or 12 months after the 16 week vaccine) is given to ensure full immunity develops. Where rabies is endemic, pups should receive 1 dose of vaccine at 12 weeks of age, but the VGG suggests that in a high-risk situation, a second dose of vaccine may be given 2 - 4 weeks later.

Early socialization of puppies is regarded as highly important for appropriate behavioural development, but has often been seen to be contradictory to advice about vaccination. Although there is some risk involved in pups that have not undergone a complete early-life vaccination schedule attending a 'puppy party', this can be minimized by using an appropriate venue and ensuring that all dogs attending are vaccinated. A recent US study has



reinforced the minimal risk that is involved [4].

For adult dogs, MLV core vaccines should be given no more frequently than every 3 years. For CDV, CAV, CPV there is excellent correlation between the presence of serum antibody (virus neutralization test or haemagglutination inhibition test) and protection from challenge with infectious virus. There are extensive data showing that protective antibody persists in adult dogs, even when they have only been vaccinated as puppies up to 14 years previously [5, 6]. More importantly there are data that underpin the legal registration of canine core MLV vaccines for either 3 or 4 years, based on challenge studies that show that vaccinated dogs resist infection for that minimum period after vaccination (the minimum duration of immunity, DOI). Other experimental data show that dogs vaccinated as puppies only are protected from live virus challenge with CDV and CPV at 9 years of age [5]. On this basis, most of the internationally produced canine MLV core vaccines used in the USA, Europe and elsewhere now have a licensed minimum DOI of either 3 or 4 years.

This is also true for most of the internationally produced adjuvanted killed rabies vaccines that may legally be given every 3 years rather than annually. Where such products are available with a 3-year licensed DOI, but governmental legislation insists on annual rabies vaccination, it is beholden on the veterinary profession to lobby for changes to the law in order to prevent unnecessary revaccination of adult dogs. For example, in the USA, state laws changed gradually, such that now every US state stipulates triennial revaccination of dogs against rabies.

The ideal core revaccination schedule for adult dogs would therefore be revaccination every third year with CDV, CAV, CPV and rabies. Where rabies revaccination is still required annually, the schedule might be CDV, CAV and CPV triennially and rabies annually. Triennial revaccination reduces the number of unnecessary vaccines given to adult dogs and therefore reduces the chances of adverse reactions.

The WSAVA guidelines also suggest that we should aim to vaccinate MORE animals. This relates to the phenomenon of 'herd immunity'. Herd immunity suggests that where at least 75% of a herd of animals is vaccinated, it is difficult for an infectious

disease outbreak to occur. The significance of herd immunity has been clearly demonstrated in the UK human populations where the uptake of measles vaccination of children decreased from the late 1990s and even in 2013 there was a major measles outbreak in Wales. In small animal medicine, the 1995 outbreak of CDV infection in Finland was also attributed to reduced herd immunity in the canine population. The 'herd' for a small animal practitioner is the population of dogs and cats living within his or her practice area – and our aim should be to have as many of these animals vaccinated as possible, in order to reduce the chances of disease outbreak in the herd.

NON-CORE VACCINATION

Non-core vaccines should be selected for the individual dog based on assessment of that particular animal's risk of exposure to the disease and assessment of the benefits of vaccination to that pet versus the risk of adverse reaction. Decision making for non-core vaccines would be facilitated by having available good quality data and disease distribution maps related to small animal infectious diseases. Unfortunately, with the exception of rabies in the USA and Europe, such distribution maps are not widely available. Some national schemes have been developed by industry or academic groups which allow practitioners to input cases of particular infectious diseases into a database that presents the information as disease distribution maps.

Monitoring the distribution and evolution of infectious diseases is an important part of vaccinology. An excellent example is canine leptospirosis, which has recently attracted much research interest as the importance of particular serovars in causing canine disease in different geographical locations is determined. This new knowledge has led to the introduction of tetravalent canine leptospirosis vaccines in the US and Europe; the antigenic composition of which is related to the prevalence of serovars in each location. Similarly, in some countries (e.g. the USA and Korea) vaccines are available to protect against strains of canine influenza virus (CIV). This infection remains an issue for dogs that are intensively kennelled and transported (e.g. racing greyhounds), but the CIV vaccine would not be recommended for general use among pet dogs.



Non-core vaccines may be included into the puppy vaccination schedule if dictated by risk assessment. Intranasal vaccines protecting against some elements of the canine respiratory disease complex (i.e. CPi and *Bordetella bronchiseptica*) might be used as early as four weeks of age with a second vaccine given 4 weeks later. The VGG recommends that where *Leptospira* vaccines are used in puppies (particularly adjuvanted bacterins) that these be given after the MLV core vaccines (e.g. at 18 and 22 weeks of age).

A major difference between MLV core vaccines and all of the non-core vaccines is that non-core vaccines (where used) require annual boosters as their DOI is no greater than 12 – 18 months. Adult dogs given non-core vaccines must therefore receive these annually. In many situations therefore, adult dogs receive ‘annual revaccination’, but just with fewer components than might have been used in the past. There are particular challenges in adopting the WSAVA guidelines in countries where reduced component vaccine products (e.g. a vaccine containing only CDV, CAV and CPV rather than these being part of an eight-in-one combination) are not yet available. The veterinary profession should lobby industry and regulators to make such products more widely available.

SEROLOGICAL TESTING IN CANINE VACCINOLOGY

The recent availability of in-practice rapid test kits for determining seroprotection against canine core viral diseases has revolutionized our ability to deliver vaccines in a safer and more scientific fashion. There are currently two such test systems available. The Titerchek™ kit is marketed by Zoetis and determines whether a dog is protected from infection by CDV and CPV. The VacciCheck™ kit is produced by Biogal Laboratories and determines protection from CDV, CAV and CPV. Both are ELISA-based systems that use slightly different technology, but are well validated against the ‘gold standard’ VN and HAI tests. Although the tests may cost more than simply revaccinating the dog, there are an important tool in the annual health check and are greatly appreciated by owners who understand the benefit of not automatically revaccinating an adult dog where this might not be required.

Serology can be used to determine whether a puppy has responded to vaccination. Testing at 20 weeks demonstrates the presence of a pup immune response (not MDA at this time) and a positive test at this time removes the need for the 12 month booster vaccine. A pup that was seronegative at 20 weeks may respond to further vaccination, but if it does not, it may be one of the rare low-responder or non-responder dogs that are genetically incapable of mounting an immune response to a particular vaccine antigen.

Serology can inform decision making as to whether an adult dog that has suffered a suspected adverse event post vaccination requires core vaccine boosters in the future. Testing of newly adopted adult dogs of unknown vaccination history can also determine whether the animal is protected or requires vaccination. In the US and Europe, progressive practitioners are now offering serological testing INSTEAD of automatic triennial core revaccination for adult dogs. Seropositive dogs do not require revaccination as they are already protected.

Serology also has a major role to play in the management of infectious disease (CDV and CPV) outbreaks in canine shelters. All dogs within the shelter are rapidly tested for the presence of antibody. Seropositive dogs (which are protected) should be separated from seronegative dogs (which are susceptible). Seronegative dogs should be vaccinated, but should not be adopted out of the shelter until beyond the incubation period for the infectious agent concerned (e.g. 2 weeks for CPV, 6 weeks for CDV). Dogs that require entry to the shelter should also be tested. Seropositive dogs may enter the shelter and be mixed with other seropositive animals. Seronegative dogs should be fostered and not enter the shelter until they have seroconverted post vaccination.

THE ANNUAL HEALTH CHECK

All aspects of vaccination should fall under an annual health check programme that reduces the emphasis on vaccination as a reason for visiting the practice and considers holistically the overall health and wellbeing of the pet. A discussion about which vaccines (or serological tests) might be offered in any one year is just one part of the annual health check. The importance of vaccination can be reinforced by using the VGG fact sheets.



Vaccination (or serology) should be appropriately invoiced so emphasis is placed on the professional consultation.

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ADVANCES IN THE DIAGNOSIS AND TREATMENT OF CANINE HYPERADRENOCORTICISM: PAST, PRESENT AND FUTURE

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Hyperadrenocorticism is associated with excessive production or administration of glucocorticoids and is one of the most commonly diagnosed endocrinopathies in the dog.

AETIOLOGY

Hyperadrenocorticism can be spontaneous or iatrogenic. Spontaneously occurring hyperadrenocorticism may be associated with inappropriate secretion of ACTH by the pituitary (pituitary-dependent hyperadrenocorticism) or associated with a primary adrenal disorder (adrenal-dependent hyperadrenocorticism).

PATHOPHYSIOLOGY

Pituitary-dependent hyperadrenocorticism

accounts for over 80% of dogs with naturally occurring hyperadrenocorticism. Excessive ACTH secretion results in bilateral adrenocortical hyperplasia and increased cortisol secretion.

Adrenal-dependent hyperadrenocorticism

accounts for the remaining 15–20% of spontaneous cases of hyperadrenocorticism in dogs and may be caused by unilateral or bilateral adrenocortical tumours, which can be benign or malignant.

CLINICAL SIGNS

Any breed of dog can develop hyperadrenocorticism but Poodles, Dachshunds and small terriers appear more at risk at developing pituitary-dependent hyperadrenocorticism. Adrenocortical tumours occur more frequently in larger breeds of dog.

Pituitary-dependent hyperadrenocorticism is usually a disease of the middle-aged to older dog, with a median age of 7–9 years. Dogs with adrenal-

