¹ Back to the basics- let's talk vaccines

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3 Abstract

4 Vaccination is an important management tool utilized in all food animal production models. Multiple vaccines exist

5 for beef and dairy cattle that cover a myriad of diseases from respiratory viruses and bacteria to venereal infections.

6 The core vaccinations recognized both in beef and dairy cattle include vaccinations for respiratory disease,

7 leptospirosis, and clostridial disease. Vaccination schedules vary depending on the operation. This proceeding will

8 discuss some of the diseases of beef and dairy cattle that are protected against by vaccinations and common

9 vaccination schedules.

10 Keywords

11 Beef Cattle, Dairy Cattle, Vaccinations

12 Introduction to vaccines

There are several different types of vaccinations available depending on the pathogen that is to be protected against. 13 14 Bacterial vaccinations are generally considered killed, meaning there is no live component of the bacteria in the 15 vaccination. These killed vaccines can be either bacterins that target the actual bacteria itself that causes disease or 16 toxoids that target the toxins produced by overgrowth of infectious bacteria. Viral vaccinations can also be killed or 17 can be modified-live. Modified-live vaccinations have a live component of the pathogen that has been modified to 18 avoid causing clinical disease but to stimulate a strong immunological response. Finally, autogenous vaccinations 19 are killed vaccinations made from a specific strain of the pathogen that is affecting a specific animal or herd. These 20 vaccinations are made to protect a herd from a specific bacterial pathogen that is causing clinical disease within that 21 herd.

22 Clinically, there is a difference between modified-live and killed vaccinations. Modified-live vaccinations will 23 generally stimulate a stronger immune response due to the ability of the pathogen to replicate within the body, 24 requiring only one dose of vaccination. Killed vaccinations will not stimulate a strong enough immune response 25 with one dose for the body to create effective antibodies because the pathogen will not replicate within the body. 26 Therefore, a follow-up dose, or booster, is required for full immunity. Modified-live vaccinations are marketed as a 27 dry cake containing the pathogen that must be reconstituted with a specific volume of sterile water. This product has 28 a long shelf life when not reconstituted but once reconstituted it is recommended to use the entire vaccination within 29 hours or days. Killed vaccinations have a longer shelf-life and do not need to be reconstituted.

30 Pathogens protected by vaccinations in cattle

31 Respiratory Viruses

32 The bovine respiratory disease complex, also known as "shipping fever", is one of the costliest diseases that affects beef cattle. The cost of treatment, loss of condition and productivity for individual animals, and mortality associated 33 34 with respiratory disease can be devastating to a beef producer. The bovine respiratory disease complex includes both 35 respiratory viruses and bacteria. The four main viral diseases of concern are bovine respiratory syncytial virus 36 (BRSV), infectious bovine rhinotracheitis (IBR), bovine viral diarrhea virus (BVDV), and parainfluenza-2 (PI-3) 37 BRSV is a paramyxovirus that affects the lower respiratory tract. Respiratory epithelial cells become multi-38 nucleated giant cells (known syncytial cells) that compromise lung function. BRSV can be asymptomatic but will 39 clinically cause an interstitial pneumonia that can result in death. Clinical signs include fever, dyspnea, open-mouth 40 breathing, bullae formation in the caudal lung fields that may rupture and lead to atelectasis, and subcutaneous 41 emphysema over the thorax (often following atelectasis). On necropsy, affected animals will have inflated lungs that 42 do not collapse when the chest cavity is open. PI3 is also a paramyxovirus but will not cause severe clinical disease 43 as a sole infectious agent. Instead, this virus will exacerbate other viral and bacterial infections. Clinical signs may 44 include cough, fever, and malaise/lethargy.

45 IBR is caused by bovine herpesvirus-1 which causes an upper respiratory infection. Clinical signs include fever,

46 cough, nasal discharge, hyperemia of the nares, and diphtheritic membranes on the nasal mucosa that appear as

47 white plaques. No lower respiratory involvement is noted with this infection. Bovine herpesvirus-1 may also lead to

48 late gestation abortion and abortion storms. Like with herpesviruses of other species, the animal will not fully clear
49 the infection and instead will go through periods of remission. Recrudescence of the virus occurs during times of
50 stress.

51 BVDV had two phenotypic types- type I and type II. BVDV does not classically cause primary respiratory disease. 52 Instead, it is an extremely immunosuppressive virus. Transient infection is possible in young and adult animals 53 where animals will remain viremic for 2 weeks and may show little to clinical signs (although immunosuppression is 54 evident). Infection in pregnant animals will, however, lead to more obvious clinical presentation. Cattle affected 55 within the first 45 days of gestation will experience early embryonic loss. Cattle affected between day 45 and 125 56 (30-150) will give birth to a persistently infected calf. This calf can be born appearing healthy or be weak and 57 unthrifty. The calf will shed the virus in all bodily fluids including mucus, blood, tears, urine, and feces and help to 58 spread the disease amongst the herd. Only about 25% of persistently infected calves will live to adulthood and 59 produce offspring, which will be persistently infected. The other 75% either succumb to other disease (the virus is 60 still severely immunosuppressive in persistently infected calves) or develop the fatal mucosal disease which 61 involves either mutation from the non-cytopathic form of BVDV that classically creates persistently infected 62 animals to the cytopathic form or superinfection with the cytopathic form. This is often a rare presentation. Cattle 63 affected between day 75-150 may have calves that develop birth defects including the most common, cerebellar 64 hypoplasia. Cattle affected after 180 days gestation will either have late term abortion, a stillborn calf, or a calf that 65 is born without the virus (the calf was immunocompetent enough to avoid becoming persistently infected). BVDV 66 type II is known to also cause thrombocytopenia along with immunosuppression.

67 Many companies market vaccinations for respiratory viruses. Both killed and modified-live options exist.

68 Respiratory virus vaccinations are considered a core vaccination for cattle and are utilized in both beef and dairy

69 production models. See below for a vaccination schedule example.

70 Respiratory Bacteria

71 The respiratory bacteria that affect cattle include Mannheimiahemolytica, Pasteurella multocida, Histophilussomni,

72 and *Mycoplasma bovis*. These bacteria are secondary invaders often following viral respiratory infection.

73 Mannheimiahemolytica and Pasteurella multocida are normal inhabitants of the nasal passage and will invade the

74 lower respiratory tract to establish disease when the mucociliary apparatus is not functioning well and the

respiratory immune functions are overwhelmed. *M. hemolytica* will cause fibrinous bronchopneumonia with extensive damage to the lung cranioventrally. An exotoxin, specifically a leukotoxin, leads to *M. hemolytica* being one of the most destructive pathogens. On necropsy, extensive fibrin covers the lungs and affected lung tissue is plum to gray, heavy, and sinks in formalin. *Pastuerellamultocida* can cause a similar clinical picture with less destruction and bronchopneumonia is characterized as purulent.

*Histophilussomni*causes a generalized vasculitis that can lead to several clinical diseases. Pneumonia from this
pathogen is characterized as purulent bronchopneumonia, but this vasculitis is also known for leading to
thromboembolic meningoencephalitis. This causes neurologic signs in cattle where they develop a fever, become
depressed then recumbent. As the disease progresses, they may become more neurologic with opisthotonos and
seizures and will ultimately die. *H. somni* may also cause arthritis and otitis interna/media. *Mycoplasma bovis* causes
a caseonecrotic bronchopneumonia with micro abscess formation in the lungs. Other infections caused by *M. bovis*include mastitis, arthritis, otitis interna/media, and keratoconjunctivitis.

87 Commercial vaccinations are available for *M. hemolytica*, *P. multocida*, and *H. somni*. *M. hemolytica* is a killed

88 toxoid, *H. somni* is a killed bacterin, and *P. multocida* is a killed bacterial extract. These pathogens are often given in

89 combination with respiratory viral vaccination although a stand-alone product with a *M. hemolytica* and *P. multocida*

90 combination does exist. These vaccinations may be useful prior to shipment of animals. Respiratory viral pathogen

91 combination products contain modified-live versions of the respiratory viruses and therefore boosters are not needed

92 with these products. The *M. hemolytica* and *P. multocida* product recommends a single 2-milliliter dose

93 intramuscularly followed by a booster only if stressful scenarios are expected. These vaccinations are not

94 considered core vaccinations across the cattle industry.

95 Clostridial Pathogens

96 There are many clostridial pathogens that can affect cattle. There are vaccinations that typically protect against 7 or 8

97 different clostridial pathogens labeled for use in cattle. These vaccinations are killed vaccines that are given as a 5-

98 milliter dose under the skin. These vaccinations may cause reaction site lesions at the injection location.

99 The 8-way clostridial vaccination protects against *Clostridium chauvoei*, *C. novyi*, *C. sordelli*, *C. septicum*, *C.*

100 *haemolyticum*, *C. perfringens* type C and D and *C. tetani* (7-way clostridial vaccinations do not have a *C.*

101 tetanicomponent). "Blackleg" caused by C. chauvoei is considered a major concern in cattle and is often the most 102 common clostridial disease recognized by cattle producers. It is important to remember that almost all clostridial 103 pathogens (except for C. perfringens) are found in the soil. These pathogens are opportunistic and cause disease in 104 two ways- either the bacteria invade the body through external wounds or through micro-abrasions in the mouth and 105 travel via the bloodstream to a target organ or the bacteria will cause infection after soil or fecal material 106 contaminate a wound. Clostridial pathogens are also anaerobic and do require an anaerobic environment to be 107 created to multiply. Finally, it is important to remember that clostridial pathogens cause disease by producing toxins 108 that affect the body. The following table may be helpful to organize the clostridial pathogens:

Bacteria	Pathogenesis	Target Organ	Clinical Disease
Clostridium chauvoei	Dormant spores activated by tissue damage and/or contaminated wounds	Muscle	Blackleg- subcutaneous emphysema and necrotizing myositis
Clostridium novyi type B	Dormant spores activated by tissue damage	Liver	Black Disease- necrotic hepatitis
Clostridium haemolyticum	Dormant spores activated by tissue damage	Liver	Bacillary Hemoglobinuria (Red Water)- hemolytic anemia, hemoglobinuria
Clostridium septicum	Dormant spores activated by tissue damage and/or contaminated wounds	Muscle	Malignant Edema- local/regional pain, swelling, signs of shock
Clostridium sordelli	Contaminated wounds	Subcutaneous tissue	Big Head- swelling, edema, necrosis of tissue

109

- 111 the Great Lakes regions of the country to protect from more than just the classic "Blackleg". This is due to the
- 112 presence of liver flukes. Liver flukes will migrate through the liver, causing damage to the liver tissue and allowing

¹¹⁰ Geographically, herds may consider use of the 8-way clostridial vaccination in the Pacific Northwest, Southeast, and

for dormant spores of *C. novyi* type B and *C. haemolyticum* to activate. Both conditions are fatal, making prevention
paramount.

115 Reproductive Pathogens

116 Leptospirosis is an important pathogen of cattle that can lead to liver and kidney damage as well as abortion. A 117 commercially available vaccination is labeled for cattle. This vaccination is a 5-way leptospirosis vaccination that 118 includes the serovars Pomona, Hardjo-Bovis, Grippotyphosa, Icterohemorrhagiae, and Canicola. Hardjo-Bovis is 119 considered host-adapted to cattle. Abortion can be caused by many of the serovars listed above. Leptospirosis is 120 transmitted via urine from affected animals, and vaccination may be most appropriate for animals that live in areas 121 with standing water (ponds, areas of poor drainage). Unfortunately, the vaccination for leptospirosis (a killed 122 bacterin with a 2-milliliter dose) has a short duration of immunity often only lasting several months. Leptospirosis is 123 considered a core vaccination in most dairy production models and is given at multiple points in the cow's

124 production life due to this short duration of immunity.

125 The venereal diseases of cattle, found mostly in beef cattle, are caused by *Campylobacter fetus* subspecies *fetus*. 126 Campylobacter fetus subspecies venerealis, and Tritrichomonasfoetis. These pathogens are spread during coitus. 127 Bulls are asymptomatic and will infect cows leading to early pregnancy loss, infertility, prolonged calving season, 128 and pyometra (especially with T. foetis). The newest vaccination available for cows claims to reduce the spread of T. 129 foetis and protect against C. fetus. This vaccination is a killed bacterin and comes as a combination product with 130 both venereal pathogens or as a singular T. foetis vaccination. It is recommended to give 2-milliliters of the single 131 pathogen product or 5-milliliters of the combination product under the skin then booster 2-4 weeks later. The final 132 booster should be done at least 4 weeks prior to the breeding season. In the past, many vaccinations for venereal 133 diseases were not considered very reliable, but this new product shows promise. This is not a core vaccination across 134 the cattle industry but is most useful in cow-calf operations utilizing natural cover.

Brucella abortus is the final reproductive vaccination that may be recommended for cattle. Brucella abortus is a
reportable disease that can lead to abortion in cattle. This vaccination is a modified-live vaccine that can only be
administered by licensed and class II accredited veterinarians. It is a zoonotic pathogen, and disease transmission
can occur via the vaccination. This vaccination is only approved for use in heifers between the age of 4 and 12
months of age. No booster is required.

140 Pinkeye

141 Infectious bovine keratoconjunctivitis, also known as "pinkeye" is caused by multiple bacteria including Moraxella

142 bovis, Moraxella bovoculi, Mycoplasma bovis, and Mycoplasma bovoculi. These bacteria will lead to epiphora,

143 blepharospasm, central corneal ulceration, corneal edema, and neovascularization of the cornea. A commercial

144 vaccination is available for *Moraxella bovis* which is a killed bacterin. The efficacy is questionable especially with

145 the variety of pathogens that can cause clinical infectious bovine keratoconjunctivitis. Autogenous vaccination may

be more effective at preventing pinkeye in herds that routinely deal with this problem.

147 Foot Rot

Foot rot is caused by *Fusobacterium necrophorum*. This condition is caused by an environmental pathogen that is ubiquitous in the soil and will cause interdigital dermatitis that does not extend beyond the interdigital space. Both beef and dairy cattle can be affected by this disease. Extensive environmental and animal management is necessary to control infection and reduce clinical cases including the use of foot baths and flushing to avoid standing manure in barns, management tasks that are easier to accomplish in the dairy industry.

A commercialvaccination does exist for *F. necrophorum* but the efficacy of this product is questionable. Other
 management practices such as removal of animals from wet, muddy, or manure heavy pastures and the

155 implementation of foot baths may be better alternatives to vaccination.

156 Rabies

Rabies is a fatal neurologic infection caused by lyssaviruses of the family Rhabdoviridae. Rabies is zoonotic and fatal to humans. The rabies vaccination is a killed vaccine. The rabies vaccine for large animals is only labeled for sheep, cattle, and horses. Rabies vaccination should be done after calves are over 3 months of age and should be boostered annually. The most common large animal specific rabies vaccination requires a 2-milliliter dose administered under the skin. It is highly recommended that any bovid with regular human contact (pet, petting zoo, educational animal) should be vaccinated for rabies in endemic areas.

163 Mastitis

164 There are several commercial products available for vaccination against coliform mastitis. The main pathogen

165 included in this vaccination is *E. coli*. Dairy farms may utilize this vaccination in their vaccine protocol, especially if

166 coliform mastitis is a common occurrence on the farm. Other management techniques such as clean and dry calving

167 pens, proper flushing of free-stall alleys, and adequate bedding turn-over in free-stalls can aid in prevention of

168 coliform mastitis.

169 Scours

Vaccination against common pathogens that lead to calf scours are commercially available. It is recommended that these vaccinations be administered to the dam prior to parturition, limiting the shedding of pathogens from the dam and into the environment while adding antibodies to the colostrum for the newborn calf. Healthy pregnant cows and heifers can be administered 2-milliliters of killed vaccination intramuscularly followed by a booster 3 weeks later. Ideally this vaccination will be boostered 3-6 weeks before calving. This killed vaccine for the dam contains E. coli,

175 rotavirus, coronavirus, and *Clostridium perfringens* type C.

176 A modified-live viral vaccination is also available for calves. This vaccination contains coronavirus and rotavirus

and is administered as a 3-milliliter dose orally to newborn calves. Dams can also be vaccinated with this product

178 and will receive two 3-milliliter doses 3 weeks apart. Ideally, the second booster dose should be given 30 days prior

to parturition to allow for proper antibody deposition into the colostrum.

180 Scour vaccinations may be helpful on dairy and beef operations, but cleanliness and proper management of

181 newborns can be just as helpful. For bottle raised calves, making sure that all equipment is properly cleaned and

182 sterilized after use is essential. Removing fecal material from the calf's environment will also decrease exposure to

183 pathogens classically found in the manure.

184 Vaccination schedule

185 The following vaccination schedules are suggestions. Different operations may elect to provide vaccinations at

186 different times during the life of the animal or elect to add other "core" vaccinations depending on diseases found

187 within that herd. The recommended core vaccinations for cattle in almost all management systems are respiratory

188 virus vaccination, leptospirosis vaccination, and clostridial vaccination.

189 Dairy Cattle

190 This protocol is primarily for heifers and cows due to the short tenure of bulls on most dairy farms. At birth it is

191 recommended to give a modified-live intranasal respiratory virus vaccination to all calves, although bull calves may

192 not be vaccinated due to their short tenure on the farm. This vaccination contains BRSV, PI3, and IBR. Injectable

193 products are not recommended until the animal is at least 6 weeks of age at which time injectable respiratory

194 viruses/leptospirosis vaccinations (either killed or modified-live) and clostridial vaccination (killed) are

195 recommended. At 9-10 weeks of ageany killed respiratory virus/leptospirosis vaccinationshould be boostered and

196 the clostridial vaccination will require a booster.

197 Depending on the farm, producers may elect to administer the first injectable vaccinations at 4-6 months of age.

198 Often called "Calfhood" vaccines, this set of vaccination includes the respiratory virus/leptospirosis vaccination and

199 the clostridial vaccination but also the *Brucella abortus* vaccination. Appropriate boosters for killed vaccinations

200 would follow 3-4 weeks later but *B. abortus* does not require a booster. Pre-breeding vaccinations will be

administered at 10 months of age or 3 weeks prior to breeding and include boosters to the respiratory

viruses/leptospirosis vaccine and clostridial vaccine. Finally, at pregnancy diagnosis it is recommended to boosterthe leptospirosisvaccination.

204 Dry off for dairy cattle is an important time to provide protection from pathogens that may lead to complications in 205 the future. For far-off cows (cows in the first 30 days of dry off), leptospirosisvaccination and vaccination for 206 coliform mastitis (if the producer is interested in this) are recommended. Thirty days later, when cows are considered 207 in the close-up dry off stage, leptospirosisvaccination, clostridial vaccination, and (if elected) vaccination for 208 coliform mastitis pathogens and vaccination against agents that cause scours in calves are recommended. In the 3-4-209 week post-partum period, it is recommended that cows receive the respiratory virus and leptospirosis vaccinations, 210 clostridial vaccination, and the final coliform mastitis pathogen vaccination. Pre-breeding vaccinations will then 211 start the cycle over again for multiparous females.

212 Beef Cattle

Although dairy cattle vaccination protocols may vary greatly from farm to farm, there is more consistently than inthe beef production model. Vaccination in the beef cattle industry depends on how often cattle are handled. Ideally,

calves will be given an initial dose of respiratory virus/leptospirosis and clostridial vaccinations two-three months prior to weaning and a booster one month later. This would allow immunity to be established prior to weaning the calf, an extremely stressful time in the calf's life. However, many beef producers may elect to wean and vaccinate calves at the same time which is often counter-productive. The stress from weaning will interfere with immunity development stimulated from the vaccination.

220 For cows being bred with live cover, pre-breeding vaccination for *Campylobacter* spp. and *Tritrichomonasfoetis* is 221 recommended. Pre-breeding vaccination, like dairy cattle, involving respiratory virus/leptospirosis vaccination and 222 clostridial vaccination may also be recommended. Pre-calving vaccination with respiratory virus/leptospirosis and 223 clostridial vaccination as well as scour pathogens vaccination is recommended with the booster being given 3 to 4 224 weeks before calving. Depending on where producers are geographically, it is more challenging to plan a structured 225 beef cattle vaccination program. For example, in the southeastern United States it is very common for bulls to be 226 with cows year-round. This means there is no structured calving season and it is challenging (if not impossible) to 227 have a structured vaccination program. Many cow-calf producers will provide a once-a-year booster vaccination for 228 respiratory virus/leptospirosis and clostridial disease to cows while vaccinating calves close to weaning or at 229 weaning.

230 Conclusion

There are a myriad of vaccinations available for beef and dairy cattle, and vaccination strategies may differ within
each industry. It is important that the veterinarian recognize what vaccinations should be considered core vaccines
(respiratory virus, leptospirosis, and clostridium) and what vaccinations can be added on a herd-by-herd basis.
Veterinarians benefit producers by working with them to establish vaccination protocols that best fit their operations
and by making producers aware of the disease threats that vaccinations can protect against.