



Teaming up for animal health

In the interest of animals, their owners and society at large





Poultry Health Seminar

Preventive measures & Gut health

Robert Jan Molenaar, DVM, FRCPath

Seeing more by looking more closely





Ask the following questions:

What am I seeing, smelling of feeling Why is this happening?

What should I do?



DOC – before arrival



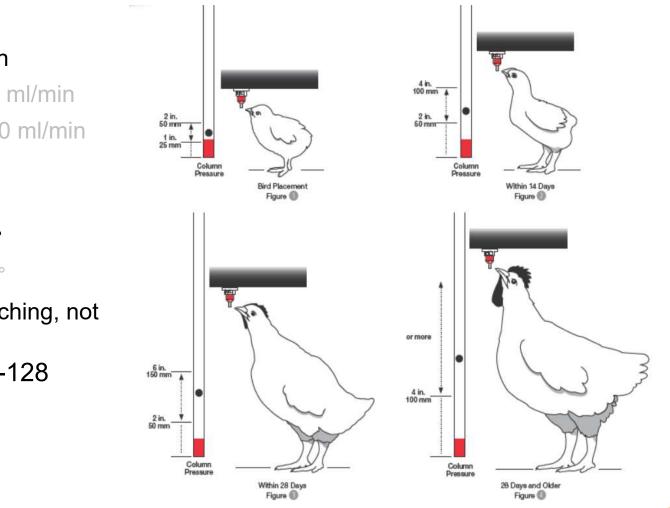
Water temperature	Effect
< 5 °C	Reduced intake of water
18-21 °C	Perfect
> 30 °C	Reduced intake of water
> 44 °C	Won't drink

Note: during the **application of live vaccine through the drinking water**, do **NOT** adhere to these temperatures



water





- Flow rate nipples
 - 0-7 days 20 ml/min
 - 7 -21 days 50 70 ml/min
 - > 21 days 70 100 ml/min
- Nipple height
 - Angle of chicken:
 - Chick: 35 to 45°
 - Growing: 75-85°
 - Straight head: reaching, not stretching
- Number of visits: 65-128







- Score 0 = closed, clean navel
- Score 1 = not completely closed, small button (<2mm) or lint
- Score 2 = open navel or large button (>2mm), wet, soiled, discoloured navel and/or soft, moist, mushy abdomen



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Crop filling



>80% after 4 hours, 85% after 8 hours, 95% after 24 hours



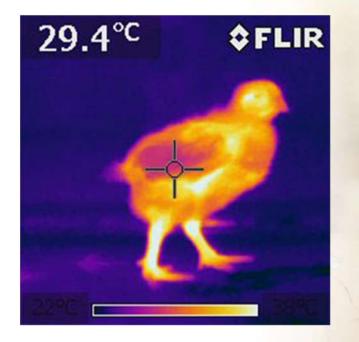


Determine filling and consistency
 (→ feed and water intake)

Cloacal temperature

- Adults: regulate their body temperature
- Baby chick: has to rely completely on environment (first 3 days)
- Measure temperature:
 - Outside temperature; changes rapidly
 - Internal temperature; useful!







Cloacal temperature

- Measure temperature
 - Upon arrival = indication of transport conditions
 - At 4 hours after arrival
 - At 8 hours after arrival
 - Daily (upto 7 days)

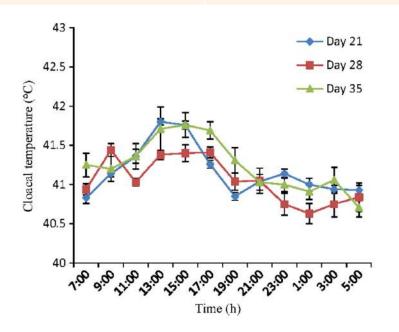




measure

Cloacal temperature and comfort zone

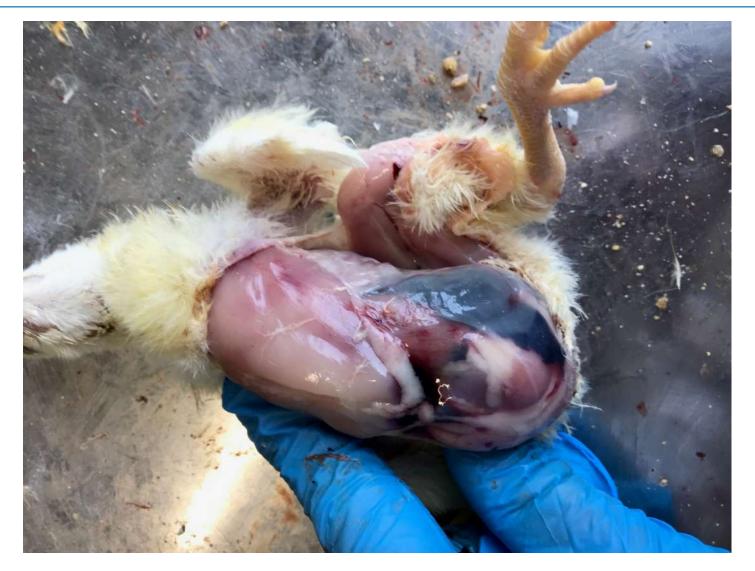
Age	Cloacal temperature
Day old chick- arrival	39,5 - 40,5
Day old chick – 4 hours	40,0 - 40,5
10 days	41,0 - 41,5





During all of this: be gentle







DOC arrival

Abnormal behaviour / sign of disease

• Example: Vit B2 deficiency





DOC arrival



- Open beak with no additional noises
 - High ambient temperature
 - Gasping for air found in chicks with an Aspergillus infection





- Natural chicken sounds
 - Clucking: lay, expectation of feed, frustration
 - Cackle: danger
 - Screech: fear, pain
 - Respiratory sounds
- Sound of the feeders
- Sound of the fans



Respiratory sounds





- Expectoration
 - irritation of the nasal passage and the beginning of the trachea, with excessive mucus development,
 - poor climatic conditions and a secondary *E. coli* complication. If occurring acutely, then it is most likely a viral infection, such as infectious bronchitis (IB), AMPV or NCD.

Coryza





Avian influenza







Production

Health

Behavior

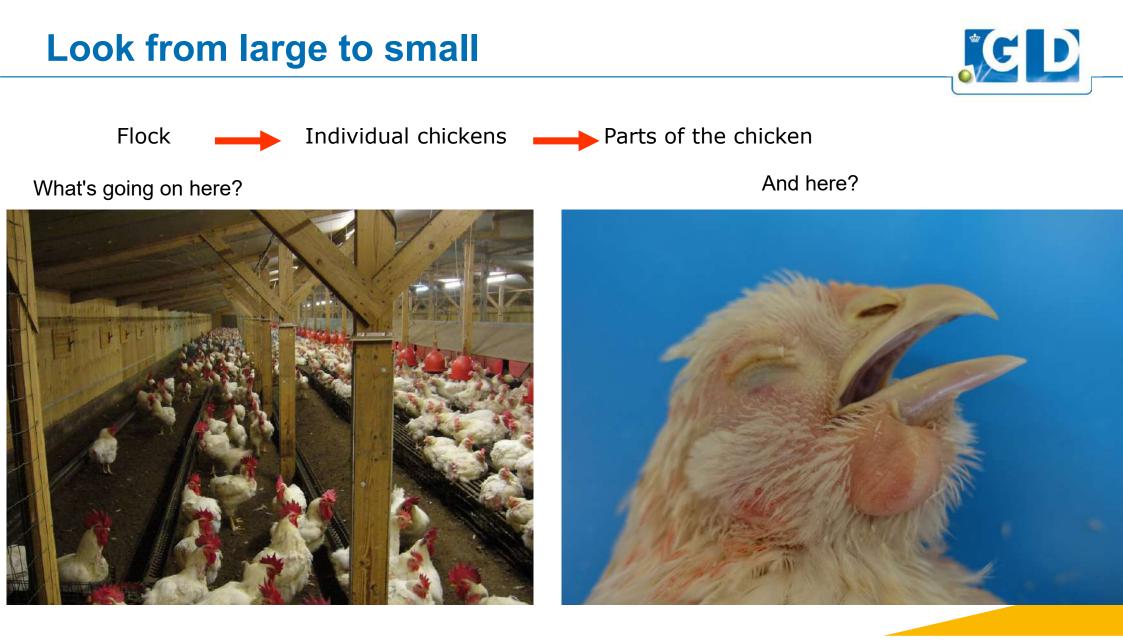












Walking around



- Take a different route as the care taker.
- Walk across the flock
- Scare them:
 - make a noise, see how they react:
 - Walk away, and observe when stop
 - Turning around?
 - Sitting down?
 - Look behind you
 - Birds should fill the empty space directly



Which one will take more feed











Bij het oppakken bledt een gezonde kip enige weerstand.



Een scherp uitstekend bot en te weinig bevlezing, wijst op een te lage voeropname.



Hoor je afwijkende geluiden, kijk dan naar eventuele natte neuzen en in de keelholte of je slijm of andere tekenen van een ontsteking ziet.



Zweilingen van of korstjes op de voetzolen zijn een teken van nat of scherp strooisel of scherpe uitsteeksels.



Stijve of warme gewrichten zijn vaak ontstoken.



Is de ruimte tussen de legbotjes smaller dan twee vingers, dan legt de kip niet.

Feathering



'Helicopter disease'

Runting and Stunting Syndrome

or

Malabsorption Syndrome

-Symptoms:

- -Poor digestion feed
- -Poor growth
- -Abnormal feathering

-Cause:

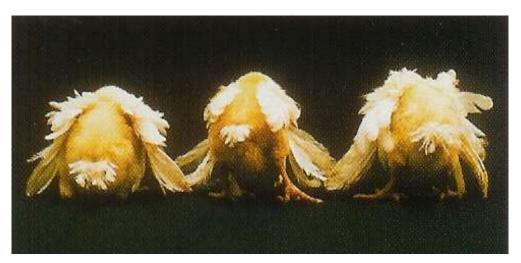
-Multifactorial

-Infections (Reo, Astro, Rota, ??).

-DDx

-Mycotoxins

-Deficiencies



Feathers





Gut health and development

From approx. week 3 onwards:

- Attention for the intestines; feed intake is rapidly increasing
- Passive and/or birds that are not alert, may have gut problems
- Should be well spread in the unit
 - If not, find the reason
- Daily gain accelerates to form skeleton, organs and muscle mass.
- At 7 days, a broiler may eat 20% of its bodyweight daily
- Too fast gain may lead to sudden death and susceptibility to diseases
 - Control by increasing dark hours and/or diluting feed.
- Vent pasting may occur due to change in feed
- Dark hours could be increased to 4 hours





Gut health and development



- Signs from Droppings
 - Undigested feed residues



Gut health and development

GD

- Signs from Droppings
- Manure consistency





- Intestinal dropping
 - solid and has a grayish-brown color. It is in parts surrounded by a thin white layer of urates







Caecal dropping



- Ceacal dropping
 - brownish-black



Caecal dropping





Right

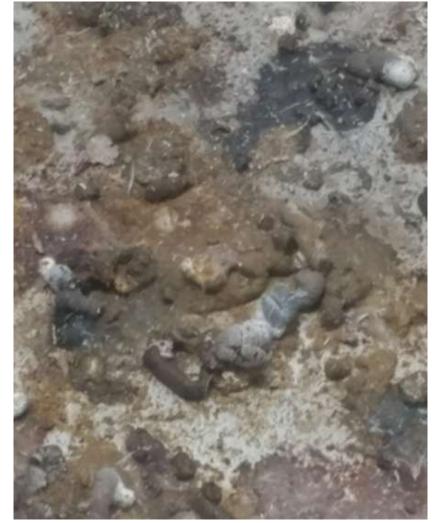
Reasonable

Not right

Diarrhoea

- Watery dropping
- No correct shape
- Highly accelerated passage
- Necrotic enteritis
- Coccidiosis





Bloody droppings

- Intestinal dropping
 - Blood around the dropping
 - Epithelium (orange)?
- Coccidiosis
 - E. tenella
 - E. necatrix
 - E maxima
- Invagination







Foamy cecale dropping

- Yellow like dropping
- Foamy
- Gas formation during fermentation in the ceaca
- Accelerated passage
 - Dysbacteriosis
 - Brachyspira



GD

Source: Broiler Signals

Bile / urates





Source: Broiler Signals

- Green dropping
- thin
- Highly accelerated passage
- Reduced feed intake
 - Newcastle disease
 - Avian Influenza
 - Gumboro/IBD
 - Acute septicaemia

Kidney disease







Source: Broiler Signals



Dark





Dark droppings usually indicate bleeding in the first part of the intestinal tract. The blood is also digested and turns an almost black colour. But an excreted piece of the intestine can also look like this (a piece that has become trapped and dies off).

Undigested feed grains





Lots of droppings and feed grains under a cage system. Digestion is very poor: feed and droppings are almost indistinguishable. The droppings are gel-like and greasy with clearly visible feed components. You can even see maize in this very poorly digested dropping; normally this is the most easily digestible ingredient of chicken feed.

43

Week 3 Gut health and development



- Signs from Droppings
- Assessing the moisture content



1. Manually

Pick up a dropping and squeeze it. In this photo you can see that the pellet contains water. When you squeeze it, it drips: this is not right. 2. Paper

Lay out paper in the broiler house on which to collect fresh droppings. If large water rings appear round the droppings, as shown on the photo, there is a problem with the manure. It could be disbacteriosis.

Gut health and development



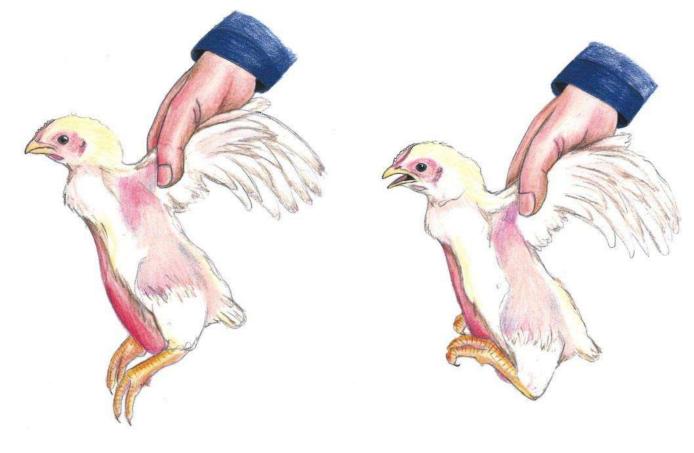
- Risk at feed transitions
- Problems may appear a few days after as gut has to re-adjust
- Optimum management reduces risks
- Chicken prefer larger particles
- In mash feeding, vitamins, trace minerals are especially in smaller particles



Week 3 Gut health and development



- Detecting stomach pain
- When lifting a chicken, feet should point forward
- Broiler at right lifts feet; could be early sign of stomach pain



Feathers on the floor







"Dysbacteriosis" evaluation

• Signs of decreased intestinal health

Avian Pathology

Publication details, including instructions for authors and subscription information: <u>http://www.tandfonline.com/loi/cavp20</u>

Morphometric evaluation of "dysbacteriosis" in broilers

E. Teirlynck ^a , M. D. E. Gussem ^b , J. Dewulf ^c , F. Haesebrouck ^d , R. Ducatelle ^d & F. Van Immerseel ^d

Ballooning



Dysbacteriosis?Loss of tonus

Artefact? Leakage tight junctions?



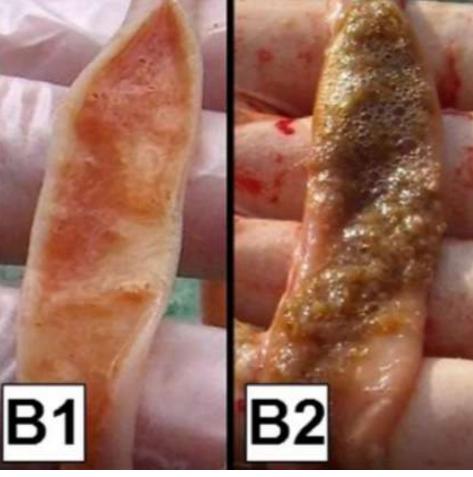




Abnormal contents

Foam (gas) Carotene Slime

Maldigestion Malsecretion Malabsorbtion



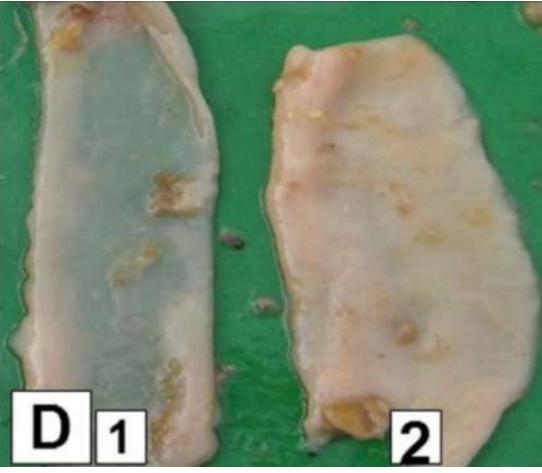


Thin gut wall



Compromised gut wall Compromised muscular layer

Infection Immune response Leakage tight-junctions?



Loss of tonus



Loss function tunica muscularis

Leakage tight-junctions?





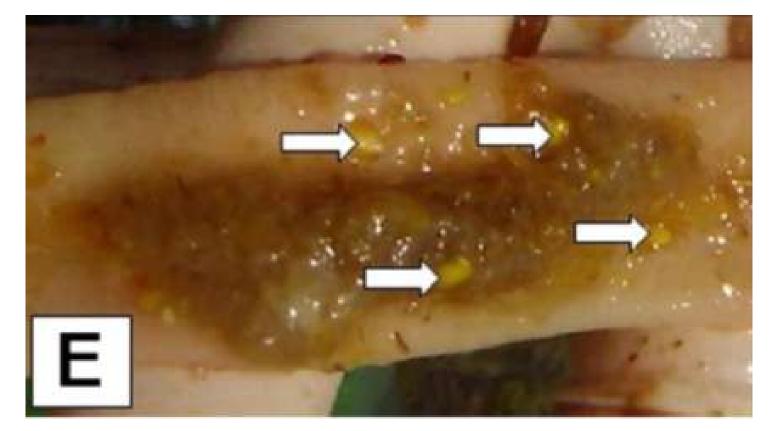
Undigested particles (colon)



Grain

Roots

Gizzard function Maldigestion





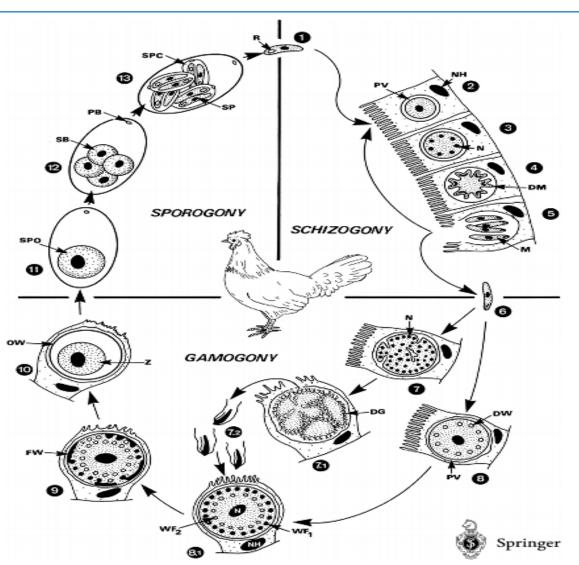
Inflammation / irritation







Coccidiosis





• Not in DOC, but potentially very early



E. acervulina



Very high reproductive potential

Duodenum

Epithelium, top of villus

Relatively mild

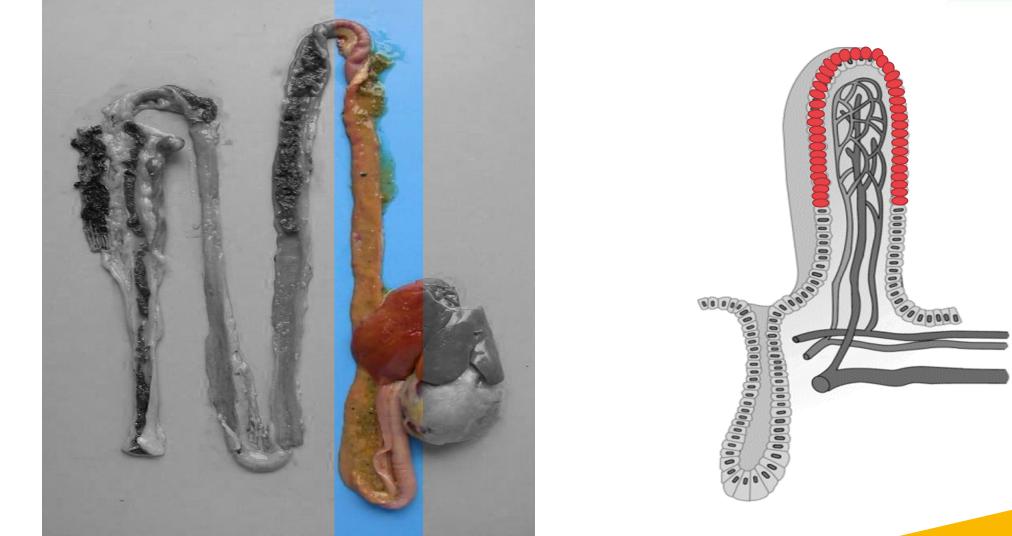
- Maldigestion
- Malabsorbtion

White streaks – coalescent white plaque



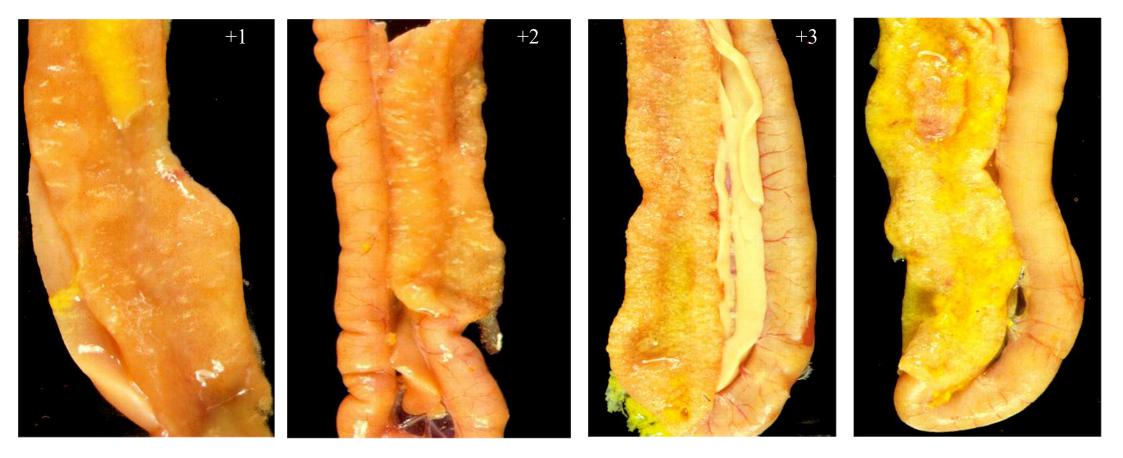






E. acervulina lesions





D.P. Conway & M.E. McKenzie, 1991. Poultry Coccidiosis







Low reproductive potential

Jejunum

Sub-epithelium, disruption of mucosa

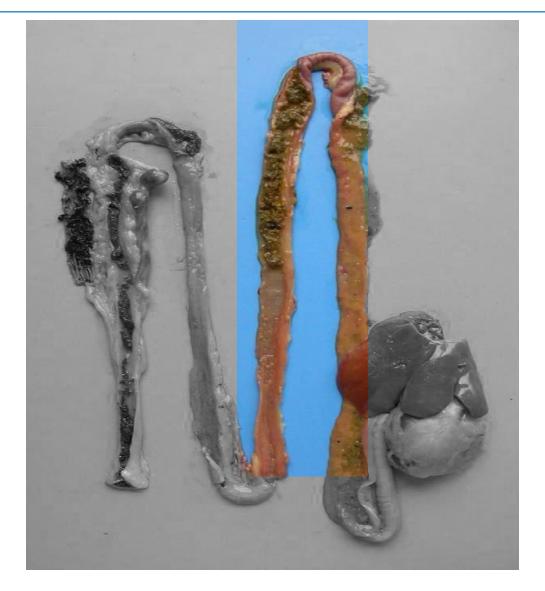
Mild - Severe

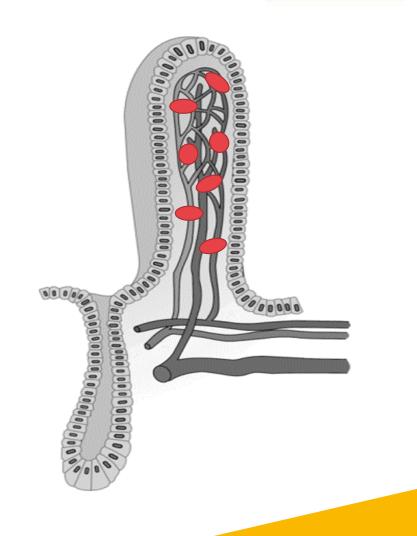
- Maldigestion
- Malabsorbtion
- Haemorage

Petechiae – bloody contents



E. maxima







E. maxima lesions





D.P. Conway & M.E. McKenzie, 1991. Poultry Coccidiosis







Medium reproductive potential

Caeca

Sub-epithelium, destruction of mucosa & muscularis

Severe

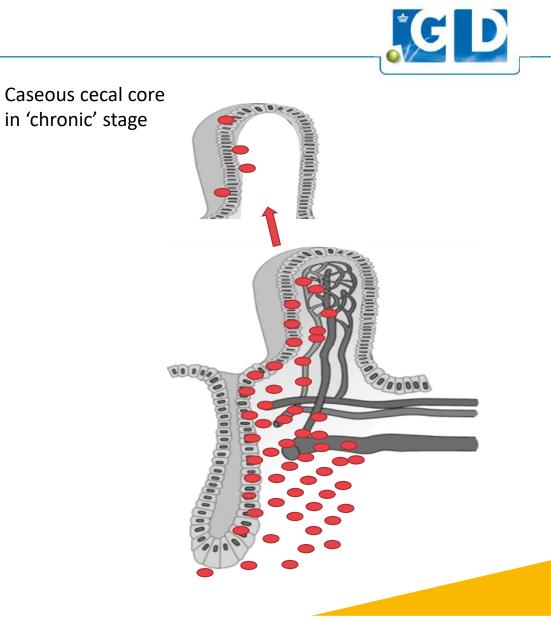
- Maldigestion
- Malabsorbtion
- Haemorrhage
- Death

Petechiae – blood filled caecum



E. tenella





E. tenella lesions





D.P. Conway & M.E. McKenzie, 1991. Poultry Coccidiosis







Thank you for your attention! Time for a short break



66



Poultry Health Seminar, continued

Biosecurity, Vaccination & Importance of monitoring

Robert Jan Molenaar, DVM, FRCPath

Vaccination

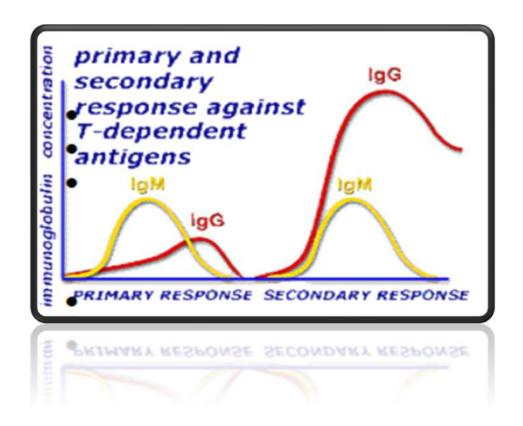




Aims of vaccination

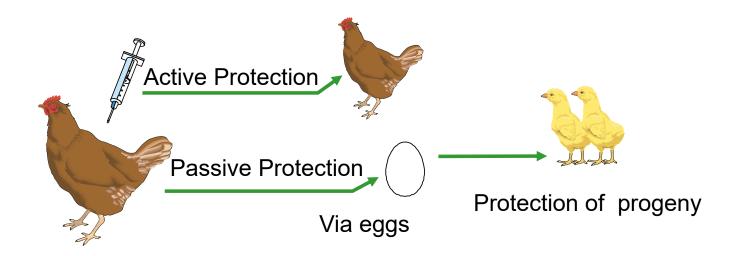
GD

- Formation of memory cells
 - Without the bird getting seriously ill (minimum of harm to the bird)
 - Inducing antibodies so that during a second contact the bird is protected.



Aims of vaccination







Live versus inactivated

Inactivated

• Application by injection

• Live (attenuated)

- Application by coarse spray
- Application by fine spray
- Application by eye drop
- Application by drinking water
- Application by wing-web
- Application by injection

• Which one can cause a vaccine reaction?





Live versus inactivated

All of them!

A vaccine reaction is required for an immune response





Inactivated



Inactivated vaccines contain formalin or oil to cause tissue irritation

Contains no live components, no risk of mutation or spreading

Safe and stable for storage

Immune respons to certain antigens of the pathogen (only 'exogenic')

Often more expensive

Best after priming with live vaccine

Systemic immunity (humeral response is high)

Less local immunity (cellular response is limited)

High protection after 2-4 weeks

No inactivation by local immunity

A lot of antigen: low risk of neutralisation by Ig



Live vaccines



Live (attenuated) vaccines cause tissue damage by replication

Early protection (1-2 weeks)

Protection does not last very long (depends on disease!)

Replication and infection of host cell simulate original pathogen

Better immune response due to simulation of natural infection

Local and systemic protection

Directed against different parts of the (total) virus: 'endogenic & exogenic antigens'

Mutations to original form might occur

Spreading and mass application possible

Often cheap per dose (compared to inactivated)

Fragile to warmth and to detergents

High level of MDA / Antibodies lower humeral response



Vector vaccines



Live (attenuated) vaccines cause tissue damage by replication

Replication stimulates the immune response of all kind of cells (cytotoxic T cells, all T helper cells, B cells)

Systemic immunity

Local immunity (low, variable)

High protection after 3-5 weeks

Against the inserted gene (protein)

HVT-vector: continuous replication, long lasting protection

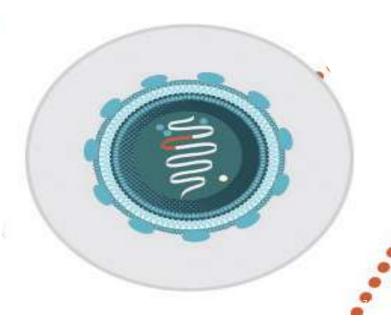
MDA has very little influence

Low spreading, individual application required

Sensitive for inactivation (cell bound)

Provides also protection against vector

VIRAL VECTOR



Boosting



- Revaccination with live vaccine (first also live):
 - Use secondary response for production of better immunity
 - Reactivate mucosal immunity (IgA)
 - Too soon after first vaccination often not effective as the protection of the previous vaccination can prevent the replication of the vaccine
- Revaccination with inactivated vaccine
 - No interference of the protection



General information for vaccination



- Use vaccines prior to their expiry date
- Follow vaccine manufacturers' instructions
- Vaccines are very sensitive to incorrect temperatures during delivery, in storage and on transport to farms
- Know what temperatures must be maintained; keep the cold chain.





- Check vaccines immediately on arrival to ensure correct vaccines have been delivered, that vaccine vials or containers are intact and that the correct temperature conditions exist
- Store live vaccines in a fridge at 4-8C
- Have max-min. thermometers in fridges. Don't open the fridge during power cut!





Take the vaccine out of the fridge just prior to vaccination
Transport in insulated containers. <u>Always!</u>



- Double-check that you have the correct vaccine
- Note the number of doses in the vaccine vial
- Check manufacturer's label advice
- Calculate the number of vials of vaccine required for the house.
 Overestimate a little.
- Set up
- Maintain vaccine at recommended temperature and other conditions



Application

- Injection
- Drinking water
- Coarse spray
- Fine spray











Live vaccination: the gold standard

Live Vaccines

- Cold chain!!
- Freeze dried
 - Stored at 4 8 °C
 - Out of the cooler: < 2 hours
 - Standard refrigerator



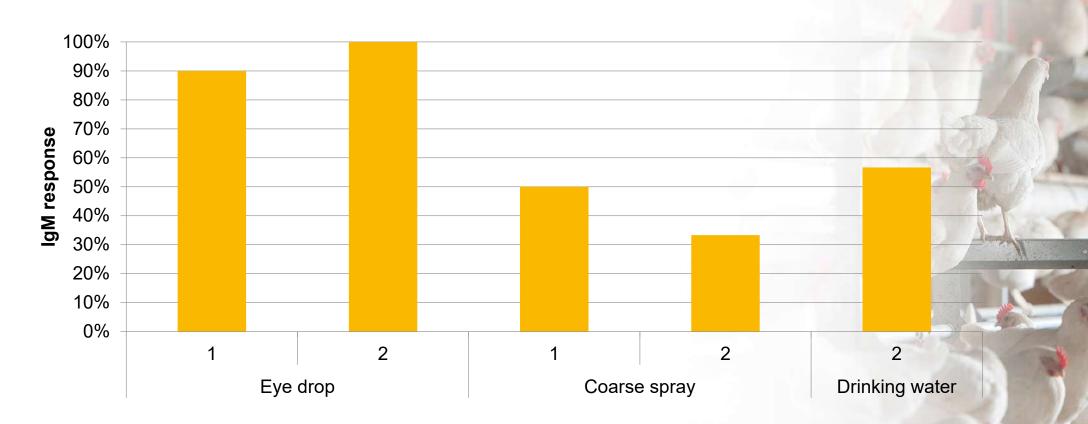


VETERINARY COLD CHAIN MANUAL Ensuring effective vaccines



Application methods





IBV H120 vaccination, IgM response SPF BroilersExperiment 1: 7 d.p.v. 10 birdsExperiment 2: 10 d.p.v. 30 birds

Spray vaccination

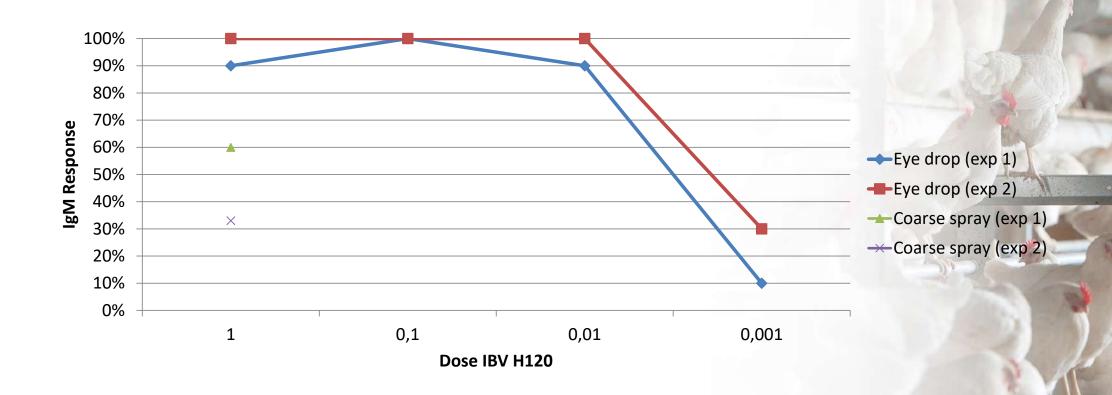
- 1. Preparation of the vaccine
- 2. Preparation of the flock
- 3. Application
- 4. Check





Spray vs eye drop





Water temperature & spray

3.2% improvement per degree Celcius lower

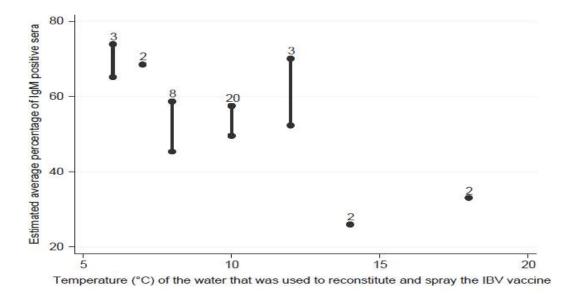


Figure 6. Association between the temperature of the water that was used to reconstitute and spray the vaccine and the average IgM response per broiler breeder flock after IBV spray vaccination at approximately 2 weeks of age. Bar: 95% confidence interval of the estimated average percentage of IgM-positive sera for each temperature of the water. Number on top of bar: number of flocks with that temperature of the water that was used to reconstitute and spray the vaccine.

De Wit et al. Av. Path (2010) 39(2) pg 123-131

2. Preparation of the flock

- Herding / rounding up
- Dimming lights
 - Less activity, eyes still open
 - 41,1% higher IgM response when light turned on
- Preparation vaccine
- Ventilation off
 - 15.5% higher IgM response when ventilation was turned off
- Heater off

De Wit et al. Av. Path (2010) 39(2) pg 123-131



3. Application of the vaccine

- Coarse spray or aerosol
- Distance to birds
- Walking speed \rightarrow hit each bird evenly
- Ventilation on



Application method

Coarse spray

- Large particles
 - 250 350 µm
- Short distances
- Contact conjunctiva
- Inhalation
- Ingestion

Aerosol

- Small particles
 - 20 100 µm
- Large distances
- Primarily inhalation



4. Check

- Manually check whether birds are wet
 - 100 birds per quarter
 - 90% wet
- Serology



IBV field trial (360 spray vaccinated flocks), practical

translation *De Wit et al*, Avian Pathology, 2010, pp 123-132

- Better results when:
 - Ventilation off during spray (15.5%, P=0,037)
 - (sufficient) Light on during spray (41%, P=0,009)
 - Second IBV vaccination not within 2 weeks (2,5% per extra day, P=0,005)
 - cold water was used (3,2% per °C, P=0,021)
 - Flock size/housing type: unclear
 - Cages significantly lower responses than floor housing (31% vs 53%, P=0,01)
 - Bigger flocks significantly lower results (1% per 1000 extra birds, P=0,04)





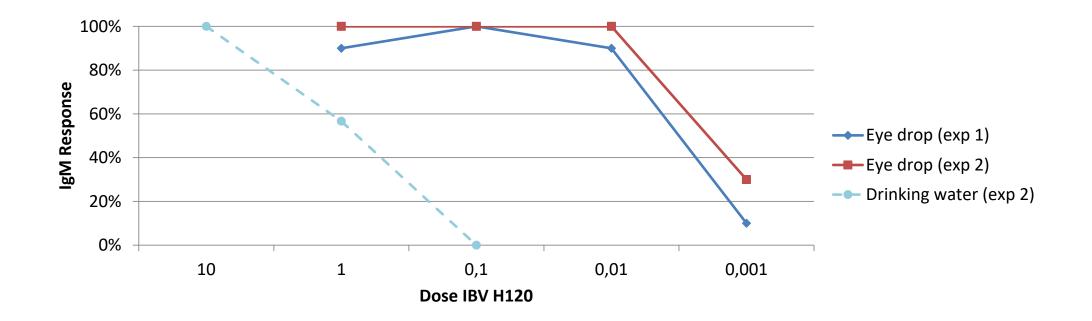
Water vaccination

- 1. Preparation of water lines
- 2. Preparation of the flock
- 3. Preparation of the vaccine
- 4. Supply vaccine
- 5. Check



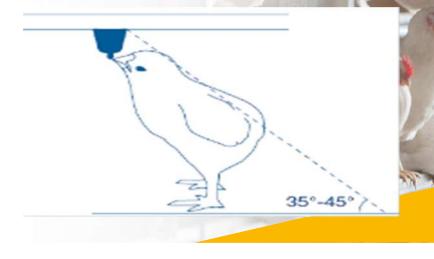
Drinking water vs eye drop





1. Prepare water lines

- Amount
 - drinkers (1 for 100)
 - nipples (1 for 10 to 12)
 - cups (1 for 12 to 15)
- Correct height
- Correct pressure (bell 0,4 0,8 bar)
- Correct flow (cup; 80-90 ml/min., nipple; 50-60 ml/min.)





1. Prepare water lines

- Best to vaccinate in early morning, when temperature is cool
- Stop treatments 24 hours before vaccination
 - Incl. necessary cleaning & disinfection
 - In less intense operated farms: stop treatments / detergents / disinfectants 3 days prior
- Flush
- Clean the bell drinker, medicator bucket and header tank with hot water (no detergent).
- Bleed & retract 1,5 2 hours
- Remove filter prior to vaccination
- Prepare vaccine
 - Add stabilizer?
 - Add coloring agent
- Provide in 2 stages of 1 hour each



1. Prepare water lines

Provide in 2 stages of 1 hour each

 The titer of reconstituted vaccines generally starts to decline after two hours

Alternative (only if 2x 1hr is not feasible):

- Vaccinate in 2 periods of 2 hours with 2/3 and 1/3 of the vaccine amount.
- Calculate the amount of water that will be drunk by the birds to be vaccinated using water consumption tables or prior morning consumption over 2 x 1 or 2 x 2 hours.

• General rule: age is the total liters of water per 1000 birds.



water consumption tables

- Tables based on feed intake x 1.85
- Ratio depends on temperature, feed, birds, etc.
- Always check the weight not the age
- Preferably use REAL data (if available) instead of general tables







Age in days	Weight	Feed intake	Water intake/ day
7	182	33	61
8	212	38	70
9	246	42	78
10	281	47	87
11	320	51	94
12	362	57	105
13	407	61	113
14	455	66	122



Age in days	Weight	Feed intake	Water intake/ day
15	500	70	105
15	506	73	135
16	561	78	144
17	618	83	154
18	678	89	165
19	741	95	176
20	806	101	187
21	874	107	198
22	944	114	211

Example flock 6000 birds



- Vaccination at 10 days
- Weight 240 gram
- Waterintake per day: 75 cc
- Waterintake 2 hours:

75/24 x 2 x 6000 = 37500 cc = 37,5 liter

Vaccination 2 periods of 2 hours of 35 liter

• Age: 10 x 6(000) = 60 liter (2 x 30)

Stabilizers

- Stability higher by addition of sterilized defatted or skimmed milk (1:50) or
- Skimmed milk powder (0.2%) or
- Coloring stabilizer (AviBlue, High Light or BlueDye)
- Visibility
- Add with high temperature
- Stabilizers are no compensation for inadequate water quality control



Milk proteins in the water bind chlorine and metallic ions, helping the survival of the vaccine virus in the water. **G**D

2. Preparation of flock

- Calculate expected water uptake 2 hours
 - Depends on age, breed, feed, temperature, length of water withdrawal
- Withdraw water for 1,5 2 hours
 - Synchronized drinking
- Do not adapt lighting scheme



3. Preparation of the vaccine

- Tank or dosator?
- Use cold water
- Use gloves
- Take required amount of vaccine out of the coolbox
- Add the stabilizer, wait 15 minutes (the chlorine has to be neutralized)
- Stir well
- Open the vial under water / use tablets
- Stir well
- Dissolved vaccine should be used within 2 hours

When using skimmed powder or skimmed milk: Add skim milk powder at the rate of 2 grams to each liter of total water or 1 liter of skimmed milk per 50 liters of total water to be used.









 Pour the diluted vaccine into the header tank or a larger bucket only used for vaccination = stock.

Oh and....

Stir well



Example



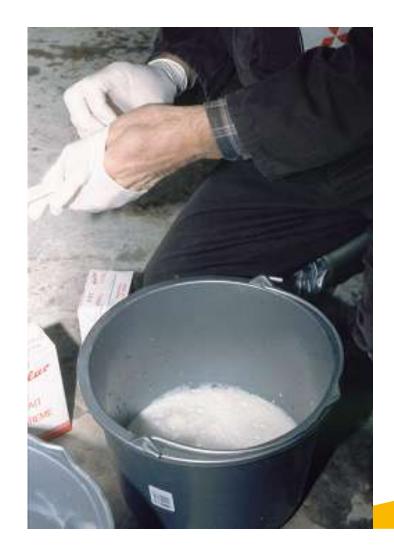
- 35 liter of water
 - = $\frac{3}{4}$ liter skimmed milk
 - = 70 gram skimmed milk powder
- Bucket: 5 liter of water + milk(powder)
- Divide the stock in the 30 liters of water to be used. (30 water + 5 stock)



Ingredients (source: Lohmann AH)







Protection: add skimmed milk

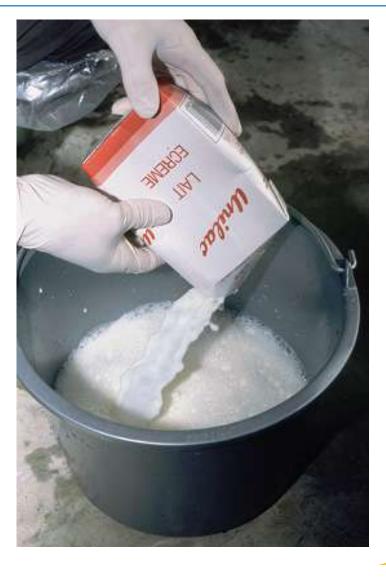


•Skimmed milk:

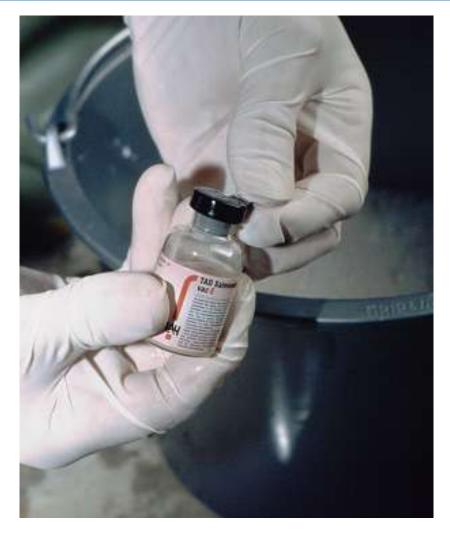
•protein protects the virus.

Fat has a negative effect on the virusPure skimmed milk of milkpowder.

•Take care dissolving (T)









Source: Lohmann AH











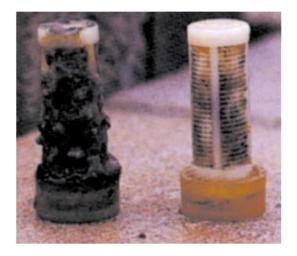




4. Supply vaccine (pipe lines)

Before preparation of the vaccine:

- Clean the lines by flushing clean water. Collect the water at the end of the line.
- Remove any water filters to the lines







4. Supply vaccine (pipe lines)



- Open valve
 - Check water flow + presence vaccine at the end of the line
 - Check flock activity
- Walk though the flock to chase away the birds from the drinker to allow other birds to drink.









- Check that all watering points are working reguarly.
- Replace the cleaned filters and turn on the water to the shed immediately after all the water containing vaccine has been drunk. Lines may need to be flushed to remove any air locks or blockages.



Water vaccination: bell drinkers



- Turn off the main water supply.
- Period of water restriction
- Dim the lights and clean the drinkers. No detergent.
- Prepare the vaccine and mix the stock in the total liters of drinking water.



Water vaccination: bell drinkers



- Dim the lights
- Bring the vaccine water to birds quickly with a watering can.
- Manpower !
- Increase light intensity and activate feeders.
- Walk around to house to stimulate the birds



Water vaccination 2: bell drinkers



- Repeat the vaccination procedure with the second batch of the vaccine
- When the water is finished turn on the main water supply.
- Check the water supply for any leakage or blockage.



Also possible: automatic bell drinkers

5. Check



- Check color stabilizer uptake
 - Check 100 birds per quarter of the house
 - 90% positive
- Serology



Problems during vaccination

- Birds are too thirsty
 - Over and under consumption
 of vaccine
 - Spilling of vaccine
 - Bad uniformity titers: protection level ↓
- Birds are not thirsty
 - Water intake more than 2 hours
 - Under consumption
 - Low titers: protection level \downarrow





Alternative Individual Vaccination



- Alternative in areas with low labour costs
- Oculo-nasal (eye drop) vaccination
 - Droplet in the eye and on the nostrils
- Oral vaccination
 - Dip the beak in vaccine water



Environmental temperature

- Correct body temperature: 41,0 41,5 °C
 - Set environmental temperature from this
- Cold: huddling, low activity
- Hot: panting, wings wide, low activity
 - Stress on respiratory tract
 - Particles deep in airways
- Temperature stress
 - Reduced immune response





Hot weather vaccination

- Vaccinate at coldest part of the day?
- Postpone vaccination a few days?
- Add stabilizer?
- Drinking water vaccination:
 - Water restriction: how long? (Dutch circumstances: 2 hours)
 - Per period: No more vaccine water than intake for 2.0 hours.



Monitoring



- Monitor for diseases on your farm: essential for informed decision on vaccinations / treatments
- Monitor for vaccine-take on your farm: can help to improve vaccination-results
- Next step: sharing findings or centralised monitoring

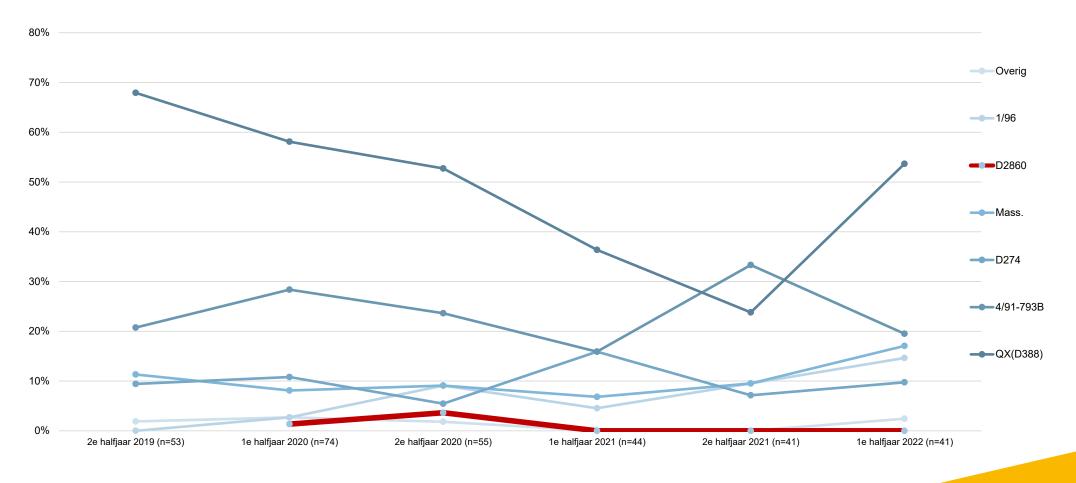
We'll shortly discuss two examples of the added value of a wider monitoring programme



D2860 & D181 – Dutch monitoring of IBV



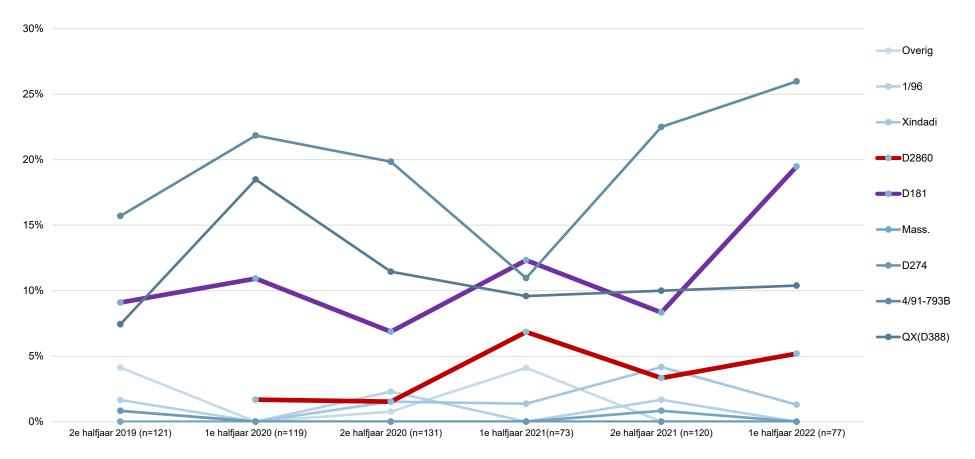
Broilers: % IB-types at farm level



D2860 & D181 – Dutch monitoring of IBV

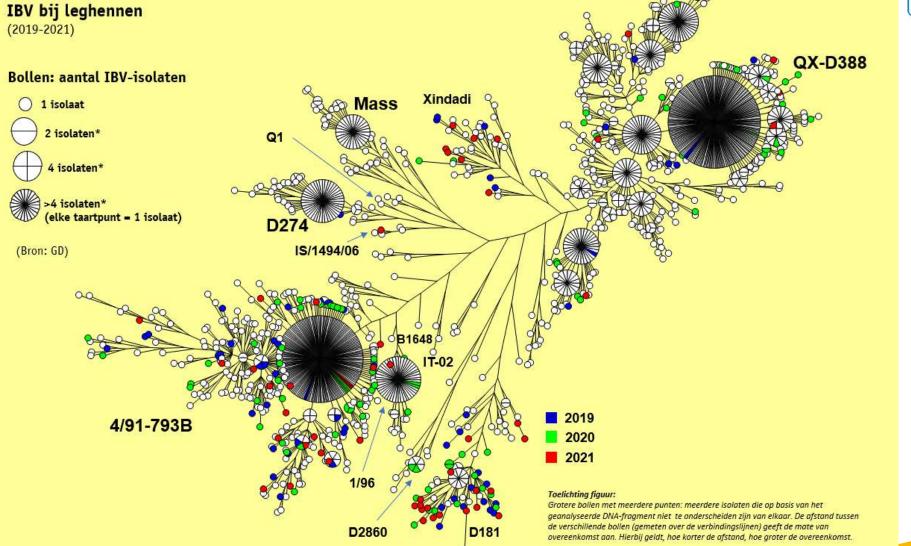




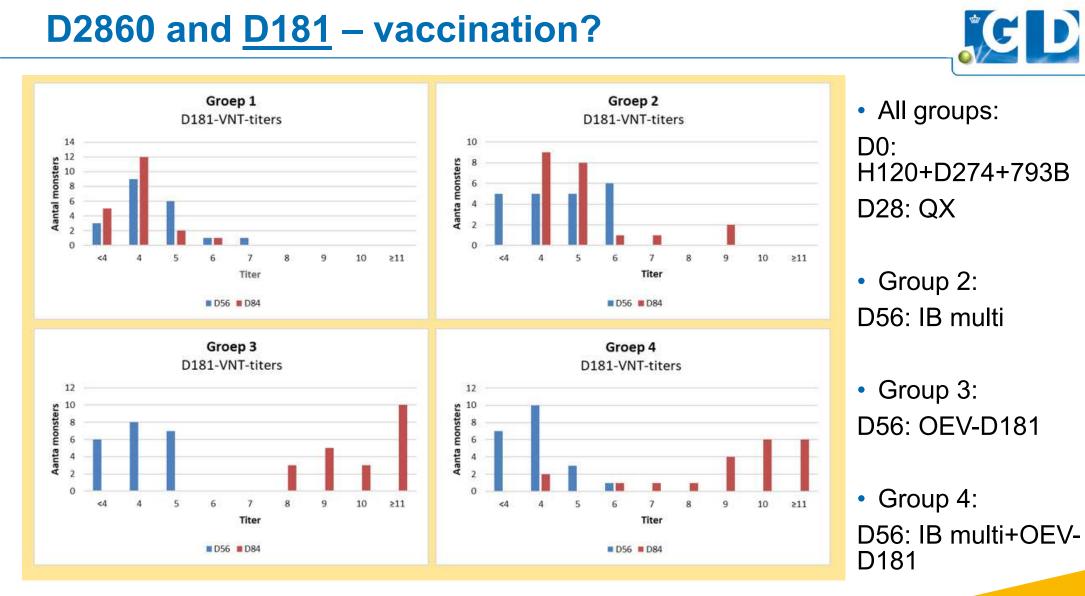


D2860 and D181 - genetics



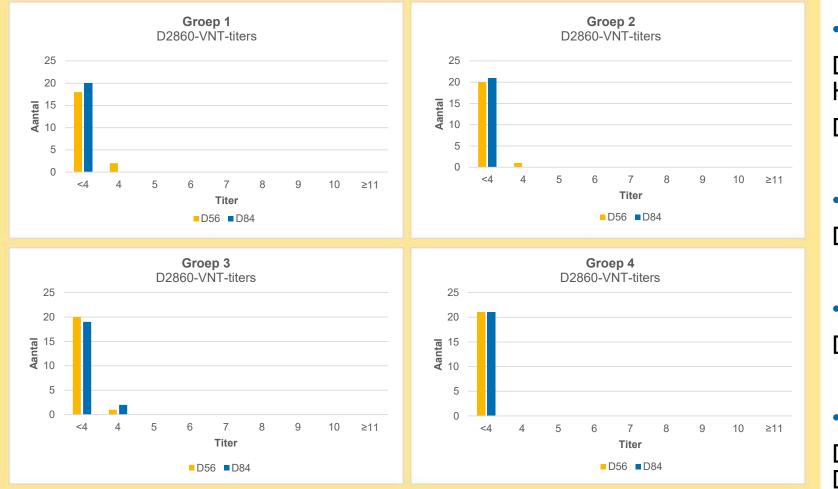


D2860 and D181 – vaccination?



D2860 and **D181 – vaccination?**





- All groups:
 D0:
 H120+D274+793B
 D28: QX
- Group 2:
 D56: IB multi
- Group 3: D56: OEV-D181
- Group 4:
 D56: IB multi+OEV-D181

Example: Marek's disease in the Netherlands

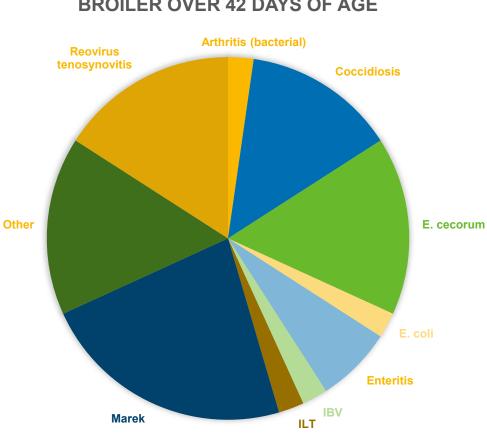


- Problems due to Marek's disease have changed in recent years, with increased problems in slower growing broiler breeds
- Initially the diagnosis was made incorrectly in the field



Monitoring GD, 2021



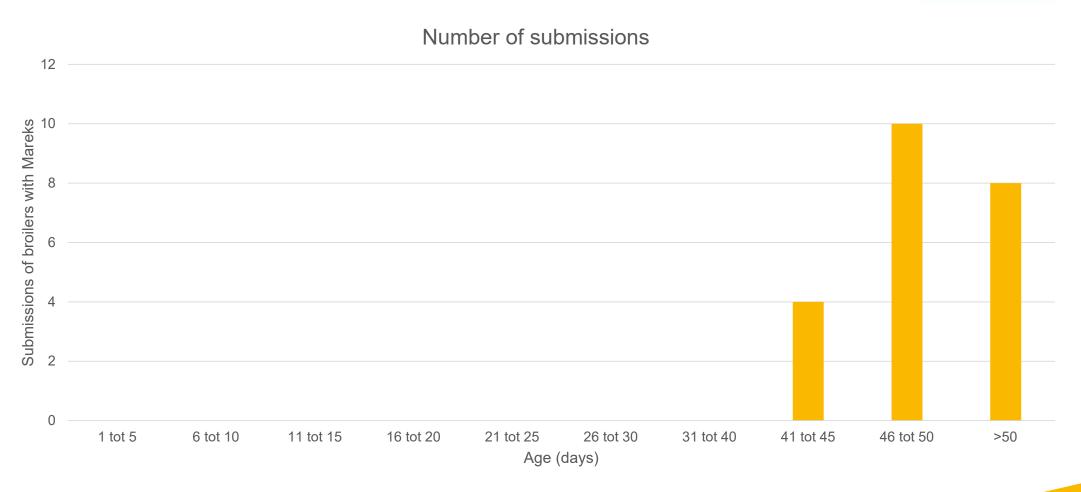


BROILER OVER 42 DAYS OF AGE



Age of submissions with Marek





When I started working at GD....

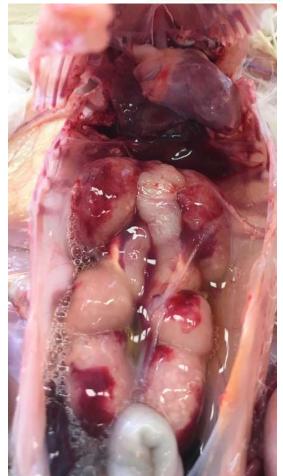




When I started working at GD....







Laying hens with tumors

When I started working at GD....



- Laying hens with tumors, sometimes nervous signs
- Maternal protection is insufficient
- Solution: Vaccinate all newly hatched laying hens at the hatchery







The classical nervous sign: leg





In the barn



Watch out! This doesn't have to be Mareks disease! (idiopathic polyneuritis, deficiencies, trauma,..)



Nowadays, it often looks different (neck)





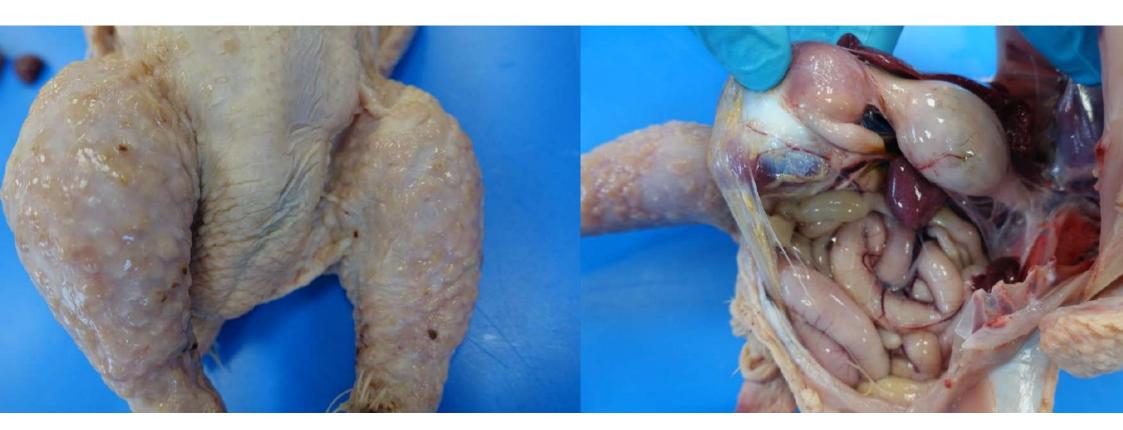
Originally confused with E. cecorum cases





Mareks during slaughter

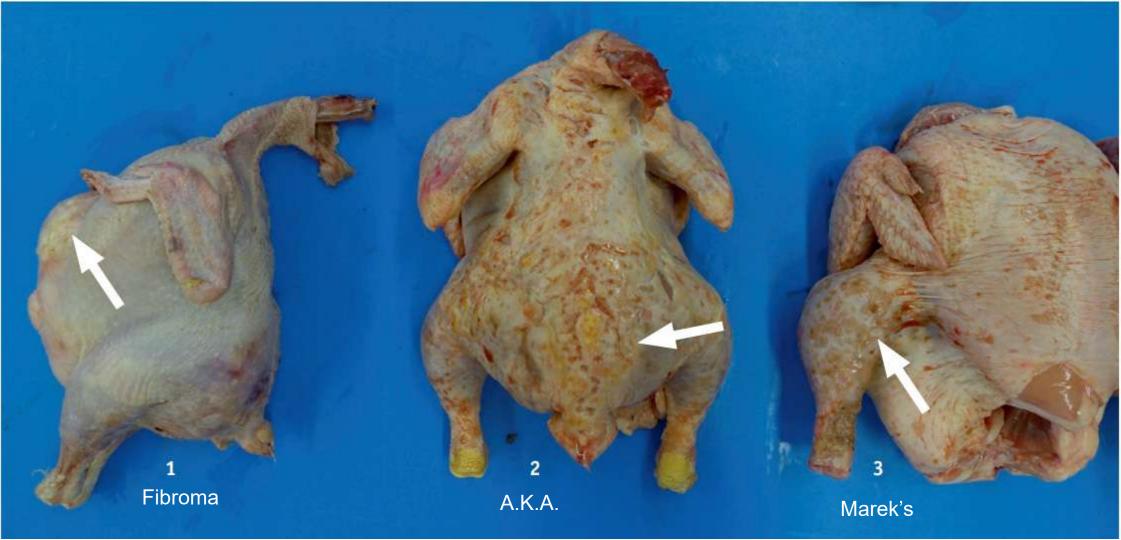






Multiple tumors during slaughter







- About 1 or 2 weeks after infection virus excretion starts
 - Fast growing chicken: late infection / low numbers of infected birds and slow spreading \rightarrow slaughtered before they have put a lot of virus in the house
 - Slower growing broiler: longer period of excredition → more virus in house → next flock is
 exposed to more virus → loner and more excretion → next flock even more exposed → …



Bron: https://karch10k.wordpress.com/2011/03/30/vicious-circle/

Example: Marek's disease in the Netherlands



- Problems due to Marek's disease have changed in recent years, with increased problems in slower growing broiler breeds
- Initially the diagnosis was made incorrectly in the field
- Monitoring & sharing of monitoringsdata has allowed to quickly and effectively prevent the disease







Thank you for your attention!