

NDVtrailPresentation

Egil Fischer

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Introduction

Trial set-up

NDV trial Presentation

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Introduction

Material and
methods

Results

Conclusion
and discussion

NDV trial:

- Transmission in vaccinated and unvaccinated groups
- Four separate groups (i.e. 2 replicates per treatment)
- Challenge in 10 out of 20 birds
- ON- and CLO-swab at each day post challenge (DPI) for 14 days
- ELISA at the end of the trial

Transmission parameteres

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$$R = \beta \cdot T_{inf}$$

- Infectious period T_{inf} = number of days a bird is infectious
- Transmission coefficient β = number of new infections by one infectious bird per day
- Reproduction ratio R = number of new infectious birds per infectious bird during its entire infectious period

If $R < 1$ an infection will go extinct.

Research questions

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- What are the values of the transmission parameters?
- Do the transmission parameters differ between vaccinated and unvaccinated groups?
- Is the R in vaccinated groups below 1?

Material and methods

Visualization of descriptive data

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- Survival plot
- Epidemic curves (number of infectious animals divided by the total) were produced.
- Histogram of infectious period

Statistical analyses

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The infectious period T_{inf} : Parametric survival analyses assuming Normal distribution

The transmission parameter β : Generalized linear model with a complementary loglog-link function:

```
glm(cbind(C, S-C) ~ 1, offset = log(I/N), family =  
binomial(link = "cloglog"), data = data.set,  
na.action = na.omit)
```

The basic reproduction number $R_0 = \beta \cdot T_{inf}$ The confidence interval is calculated by

$$\log(R) = \log(\beta) + \log(T_{inf}) \pm Z_{0.05} \cdot (\log(SE(\beta)) + \log(SE(T_{inf})))$$

Assumes independence between the transmission parameter and the infectious period.

Transmission model and further assumptions

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Transmission model

- SIR: Susceptible → Infectious → Recovered and immune

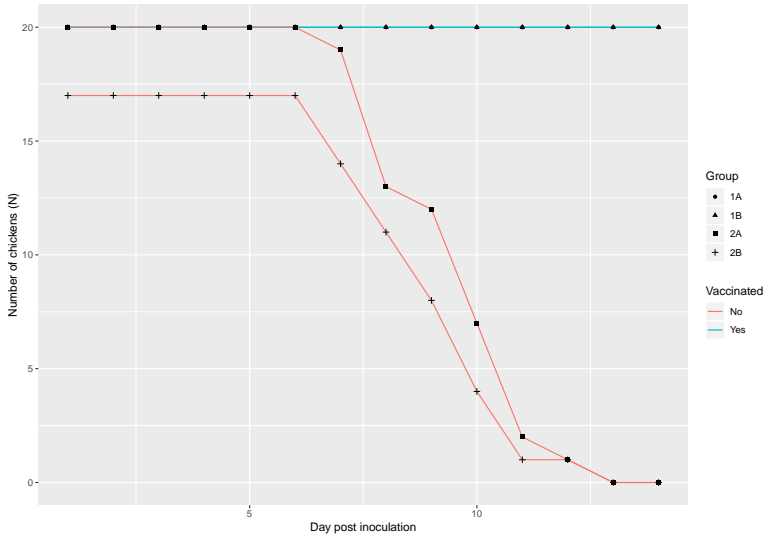
The determination of a single sample to be positive depends on two assumptions

- Detection threshold (default set to $C_t < 36$)
- Sample is positive if at least one consecutive sample is positive: Pos, Neg, Pos, Pos = first positive is assumed to be negative, 3rd and 4th positive (same for negative but the opposite)

Results

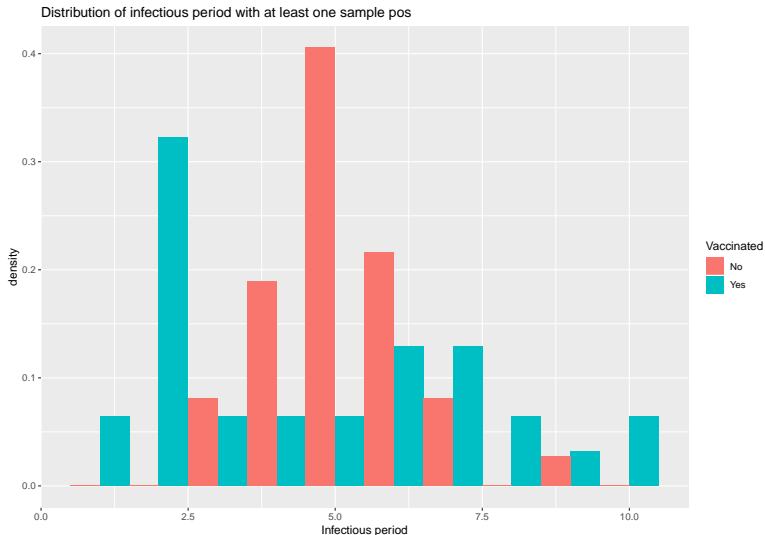
Survival curve

survivalplot



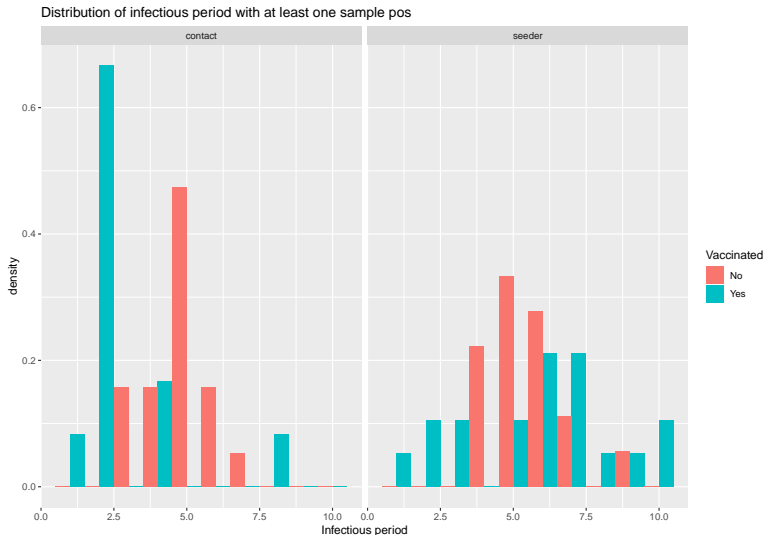
T_{inf} at least one (CLO or ON) positive sample

hist.one



T_{inf} at least one (CLO or ON) positive sample

hist.one.chal



Statistical analyses ~ using parametric survival analyses

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Survival analyses assuming a Gaussian distributed T_{Inf} and only using the contact birds.

```
summary(surv.fit.one.SIR.gaussian)
```

```
##
```

```
## Call:
```

```
## survival::survreg(formula = survival::Surv(infer,  
##      data = out.indiv.one.SIR[out.indiv.one.SIR$Cha  
##      ], na.action = na.omit, dist = "gaussian")
```

```
##              Value Std. Error      z      p  
## (Intercept)   4.789      0.323 14.81 <2e-16  
## VaccinatedYes -1.622      0.547 -2.96 0.0030  
## Log(scale)    0.343      0.132  2.60 0.0093
```

```
##
```

```
## Scale= 1.41
```

```
##
```

Transmission parameter β

```
summary(fit.one.SIR.fixed)
```

```
##
```

```
## Call:
```

```
## glm(formula = cbind(C, S - C) ~ Vaccinated, family =
```

```
##       data = out.nona.one.SIR, na.action = na.omit,
```

```
##
```

```
## Deviance Residuals:
```

```
##      Min       1Q   Median       3Q      Max
```

```
## -2.66807  -1.18662   0.06918   1.31868   2.82900
```

```
##
```

```
## Coefficients:
```

```
##              Estimate Std. Error z value Pr(>|z|)
```

```
## (Intercept)    0.1069    0.1730   0.618   0.537
```

```
## VaccinatedYes -0.3525    0.2231  -1.580   0.114
```

```
##
```

```
## (Dispersion parameter for binomial family taken to
```

β estimates

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```
beta.table
```

```
##      Treatment  beta  2.5%  97.5
## 1      Unvac 1.113 0.777 1.537
## 2       Vac 0.782 0.507 1.221
```

Vaccination does not lower the infection parameter β .

Summarize ~ infectious period

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```
mean.beta.inf.R[, c(1,5,6,7)]
```

```
##      Vaccinated      infper llinfper ulinfper
## 1                No 4.789474 4.155548 5.423399
## 2                Yes 3.167164 1.460171 4.874158
```

Vaccine reduces length of infectious period ($p = 0.004$)

Summarize $\sim \beta$

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```
mean.beta.inf.R[, c(1,2,3,4)]
```

##	Vaccinated	beta	llbeta	ulbeta
## 1	No	1.1128227	0.7766340	1.536829
## 2	Yes	0.7822649	0.3540352	1.686715

No difference in transmission rate

Summarize ~ R

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```
mean.beta.inf.R[, c(1,11,12,13)]
```

##	Vaccinated	R	llR	ulR
## 1	No	5.329835	3.5914571	9.275196
## 2	Yes	2.477561	0.4043934	6.431245

No difference in reproduction number

Conclusion and discussion

Conclusion

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Based on the assumption of one of both samples required to be positive, we find that the transmission parameter β is not lower, but the mean infectious period T_{inf} in the contact birds is lower, which results in a reproduction number R which is smaller (although not statistically significant), but is not lower than the threshold $R \leq 1$.

Optional additional analyses

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Preliminary results (models not checked)

- Different Ct as cut-off: e.g if $Ct = 30$, β lower values for vaccinated, T_{inf} cannot be estimated
- One positive sample means that animal is positive and first negative is negative: β lower values for vaccinated, and T_{inf} lower for vaccinated

Following no preliminary results

- Only base results on ON swabs
- SI model
- Use ELISA to determine infection yes/no.