#### **Introduction to GLM**

**Generalized Linear Model** 

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#### Definition

The generalized linear models and linear models, allow to study the relation between the response variable (Y) and a sets of explanatory variables  $(X_1...X_k)$ 



#### The linear models are composed of:

- ► A response variable (Y) Variable of interest
  - Let's say that  $(Y_1...Y_n)$  is a sample of size n of Y.  $Y_1...Y_n$  are independant.
    - Y<sub>i</sub> is normally distributed
- **Explanatory variable(s)**  $(X_1...X_k)$  Variable(s) used to explain the variability in the response variable
- ► Explanatory variables can be expressed as :  $\beta_0 + \beta_1 X_1 + ... + \beta_k X_k$
- Sometimes, an explanatory variable X<sub>j</sub> can be deduced by elementary variables.
  - $X_3 = X_1 * X_2$



#### Linear models

More precisely, linear models can be expressed as :

$$E(Y) = \beta_0 + \beta_1 X_1 + \dots + \beta_k X_k + \epsilon$$

#### Where:

- ► E(Y) is the expected value of Y
- ε is the error parameter (must follow a normal distribution and homoscedastic)

We want to find the equation that best suits our data  $(Y_1...Y_n)$ . The parameters  $\beta_0, \beta_1...\beta_n$  can be estimated by the least-square method. Their estimations are those which minimize:

$$\sum_{i=1}^{n} (Y_i - (\beta_0 + \beta_1 X_{1i} + ... + \beta_k X_{ki}))^2$$



#### Limits of linear models

- Can't explain a response variable that don't follow a normal distribution
- Can't explain a response variable that takes value in a particular interval
- Explanatory variables must have a linear effect on the response variable

To overcome those issues, we can use a generalized linear model

## Introduction Generalized linear models



Distribution	Interval	Uses	link function
Normal	$]-\infty,+\infty[$	Linear response data	$E(Y) = \beta X$
Poisson	$[0,+\infty[$	Count data	$log(E(Y)) = \beta X$
Bernoulli	{0,1}	outcome of an event	$log\left(\frac{E(Y)}{1-E(Y)}\right) = \beta X$
Binomial	{0,, <b>N</b> }	outcome of $N$ events	$\log\left(\frac{E(Y)}{1-E(Y)}\right) = \beta X$
Exponential/Gamma	$]-\infty,+\infty[$	Exponential response data	$E(Y)^{-1} = \beta X$

#### Poisson distribution



#### Count data

Y is a categorial continuous data. Let's note  $E(Y) = \mu$ . Y  $\hookrightarrow Pois(\mu)$  with  $P(Y = k) = \frac{\mu^k e^{-\mu}}{k!}$ 

- $\blacktriangleright$   $E(Y) = Var(Y) = \mu$

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- $Y \hookrightarrow Pois(\mu)$  with  $P(Y = k) = \frac{\mu^k e^{-\mu}}{k!}$
- E(Y) = Var(Y) = μ
- ► The link function is the log, the model is:

$$log(\mu) = \beta_0 + \beta_1 X_1 + \dots + \beta_k X_k \in ]-\infty, +\infty[$$

#### Poisson distribution

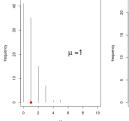


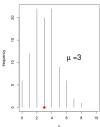
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- ► The link function is the log, the model is:

$$log(\mu) = \beta_0 + \beta_1 X_1 + \dots + \beta_k X_k \in ]-\infty, +\infty[$$
 on the count scale: 
$$\mu = \mathrm{e}^{\beta_0 + \beta_1 X_1 + \dots + \beta_k X_k} \in [0, +\infty[$$

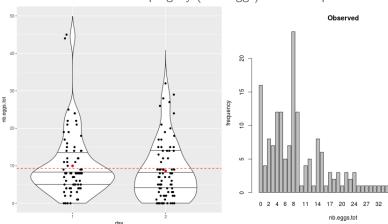




# Example: number of worm eggs against time roughly from Manon Grosmaire experiments



n measurements of size of progeny (=nb eggs) on 2 time points.





#### Is there eggs production?

glm0 <- glm(nb.eggs.tot  $\sim$  1, data, family = "poisson") The null model is:  $\log(\mu) = \beta_0$ 



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$$\log(\mu) = \beta_0$$

```
> summary(glm0)
Call:
glm(formula = nb.eggs.tot ~ 1, family = "poisson", data = data)
Deviance Residuals:
Min 1Q Median 3Q Max
-4.3166 -1.9670 -0.4421 1.3548 8.3888
Coefficients:
           Estimate Std. Error z value Pr(>|z|)
(Intercept) 2.23178 0.02606 85.63 <2e-16 ***
Signif. codes: 0 '***' 0.001 '**' 0.01 '*' 0.05 '.' 0.1 ' ' 1
(Dispersion parameter for poisson family taken to be 1)
    Null deviance: 1038.3 on 157 degrees of freedom
Residual deviance: 1038.3 on 157 degrees of freedom
AIC: 1598.9
Number of Fisher Scoring iterations: 5
```



If the model is true, asymptotically, estimators are gaussian.

```
> summary(qlm0)
Call:
glm(formula = nb.eggs.tot ~ 1. family = "poisson". data = data)
Deviance Residuals:
                  Median
                                       Max
-4.3166 -1.9670 -0.4421 1.3548
Coefficients:
           Estimate Std. Error z value Pr(>|z|)
(Intercept) 2.23178 0.02606 85.63 <2e-16 ***
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- Recalling,  $log(\mu) = \beta_0$ , thus  $\mu = e^{\beta_0}$ .
- $IC_{95\%}(\beta_0) = \beta_0 \pm 1.96 \times \sigma_{\beta_0}$



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- Recalling,  $log(\mu) = \beta_0$ , thus  $\mu = e^{\beta_0}$ .
- $IC_{95\%}(\beta_0) = \beta_0 \pm 1.96 \times \sigma_{\beta_0}$
- The mean number of eggs predicted is  $e^{\beta_0} \simeq 9.3[8.9, 9.8]$  which is significant (Wald test, p-value <5%).

# Example: number of worm eggs against time Day effect model



#### Is there a change of eggs production according to day?

 $glm1 \leftarrow glm(nb.eggs.tot \sim day, data, family = "poisson")$ The model is:

$$log(\mu) = \beta_0 + \beta_1 \times day$$

 $\triangle day$  is a factor: day = 0 for day 1, day = 1 for day 2).

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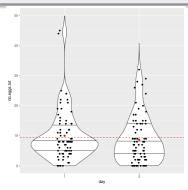
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```
> summary(glm1)
Call:
glm(formula = nb.eggs.tot ~ day, family = "poisson", data = data)
Deviance Residuals:
             10 Median
-4.4693 -2.1574 -0.6517 1.1964 8.0905
Coefficients:
           Estimate Std. Error z value Pr(>|z|)
(Intercept) 2.30132 0.03560 64.64 < 2e-16 ***
                      0.05226 -2.76 0.00577 **
dav2
           -0.14427
Signif. codes: 0 '***' 0.001 '**' 0.01 '*' 0.05 '.' 0.1 ' ' 1
(Dispersion parameter for poisson family taken to be 1)
   Null deviance: 1038.3 on 157 degrees of freedom
Residual deviance: 1030.6 on 156 degrees of freedom
AIC: 1593.3
Number of Fisher Scoring iterations: 5
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  - ► Effect of  $x_1$ :  $L_{x_1} = \prod_{i=1}^{n} \frac{\exp(\beta_0 + \beta_1 x_{1i})^{y_i} e^{-\exp(\beta_0 + \beta_1 x_{1i})}}{y_i!}$



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$$D_{effect x_1} = D_{null} - D_{x_1} \qquad \hookrightarrow \chi^2((n - p_{null}) - (n - p_{x_1}))$$



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If  $\chi^2_{obs} < \chi^2_{th}$ , the 2 models are not statistically different and you should choose the more parsimonious.



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  - $e^{\beta_1}=rac{\mu_{day_2}}{\mu_{day_1}}=0.9[0.78,0.96]$  indicates effect of ageing is significantly deleterious for eggs production (Wald test, p-value <5%).



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 $e^{\beta_1} = \frac{\mu_{day_2}}{\mu_{day_1}} = 0.9 [0.78, 0.96] \text{ indicates effect of ageing is significantly deleterious for eggs production (Wald test, p-value <5%)}.$ 

Is it though relevant to add a day effect?

```
> anova(glm0, glm1, test = "Chisq")
Analysis of Deviance Table

Model 1: nb.eggs.tot ~ 1

Model 2: nb.eggs.tot ~ day
Resid. Df Resid. Dev Df Deviance Pr(>Chi)

1 157
2 156 1030.6 1 7.6398 0.00571 **
---
Signif. codes: 0 '***' 0.001 '**' 0.01 '*' 0.05 '.' 0.1 ' ' 1
```



- ▶ On linear predictor scale,  $log(\mu) = \beta_0 + \beta_1 \times day$
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 $e^{\beta_1} = \frac{\mu_{day_2}}{\mu_{day_1}} = 0.9 [0.78, 0.96] \text{ indicates effect of ageing is significantly deleterious for eggs production (Wald test, p-value <5%)}.$ 

Is it though relevant to add a day effect?

 $\chi^2_{obs} < \chi^2_{th}$ , the day effect model explains significantly better the variability of the data (1-pchisq(7.6398, df = 1)).

# Example: number of worm eggs against time Zoom on model assumption



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Residual variance is estimated by  $\frac{1}{n-p} \sum_{i=1}^{n} (y_i - \mu_i)^2$ .

If Poisson law and model are adapted,  $\frac{1}{n-p}\sum\limits_{i=1}^{n}\frac{(y_i-\mu_i)^2}{\mu_i}\sim 1$ 

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If Poisson law and model are adapted,  $\frac{1}{n-p}\sum\limits_{i=1}^{n}\frac{(y_i-\mu_i)^2}{\mu_i}\sim 1$ 

or here it equals 6.98, indicating E(Y) < Var(Y).

Tests are not reliable and this is not the best model; it causes sd error to be deflated which could lead to significant predictor whereas it is not.



#### To deal with over-dispersion, knowing $E(Y) = \mu$

```
• quasiPoisson: Var(Y) = \phi \mu glm1b <- glm(nb.eggs.tot ~ day, data, family = "quasipoisson")
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Negative binomial:  $Y \hookrightarrow Pois(\Theta \times \mu)$  and  $\Theta \hookrightarrow Gamma(\alpha, \alpha)$  with  $E(\Theta) = 1$   $Var(Y) = \mu + \alpha \times \mu^2$  The link function becomes:  $log(\frac{\alpha\mu}{1+\alpha\mu})$ . glm2 <- MASS::glm.nb(nb.eggs.tot  $\sim$  day, data)

## Example: number of worm eggs against time Alternatives



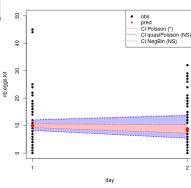
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▶ Negative binomial:  $Y \hookrightarrow Pois(\Theta \times \mu)$  and  $\Theta \hookrightarrow Gamma(\alpha, \alpha)$  with  $E(\Theta) = 1$ 

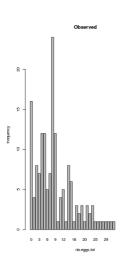
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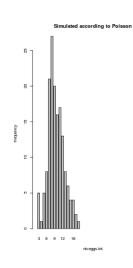
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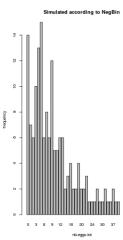


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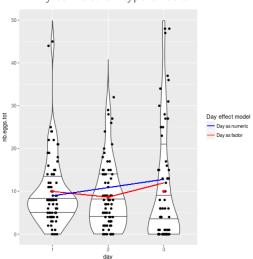








Why to insist on type of data?





Quick display of interaction: add the strain effect.

#### Without interaction:

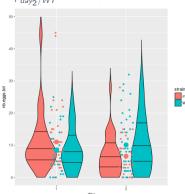
glm1ds <- glm(nb.eggs.tot ~ day+strain, data, family = "poisson")  $Im(\mu) = \beta_0 + \beta_1 \times day + \beta_2 \times strain$  $\mu_{day_2/WT} = e^{\bar{\beta}} 0 e^{\beta} 1 e^{\beta} 2$ 10-



Quick display of interaction: add the strain effect.

#### Without interaction:

# glmids <- glm(nb.eggs.tot $\sim$ day+strain, data, family = "poisson") $Im(\mu) = \beta_0 + \beta_1 \times day + \beta_2 \times strain$ $\mu_{day_2}/WT = e^{\beta_0} e^{\beta_1} e^{\beta_2}$



#### With interaction:

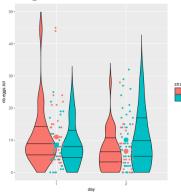
$${\tt glm1dsI} \, \leftarrow \, {\tt glm(nb.eggs.tot} \, \sim \, {\tt day*strain, \, data, \, family \, = \, "poisson")}$$



► Quick display of interaction: add the *strain* effect.

#### Without interaction:

glmids <- glm(nb.eggs.tot 
$$\sim$$
 day+strain, data, family = "poisson") 
$$Im(\mu) = \beta_0 + \beta_1 \times day + \beta_2 \times strain$$
 
$$\mu_{day_2/WT} = e^{\beta_0} e^{\beta_1} e^{\beta_2}$$



#### With interaction:

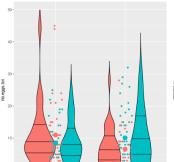
glmidsI <- glm(nb.eggs.tot 
$$\sim$$
 day\*strain, data, family = "poisson") 
$$Im(\mu) = \beta_0 + \beta_1 \times day + \beta_2 \times strain + \beta_3 \times day \times strain$$
 
$$\mu_{day_2/WT} = e^{\beta_0} e^{\beta_1} e^{\beta_2} e^{\beta_3}$$



Quick display of interaction: add the strain effect.

#### Without interaction:

glmids <- glm(nb.eggs.tot 
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 day+strain, data, family = "poisson") 
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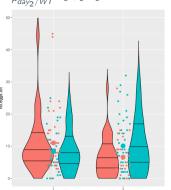
```
glmldsI <- glm(nb.eggs.tot \sim day*strain, data, family = "poisson") Im(\mu) = \beta_0 + \beta_1 \times day + \beta_2 \times strain + \beta_3 \times day \times strain \mu_{day_2/WT} = e^{\beta_0} e^{\beta_1} e^{\beta_2} e^{\beta_3}
```



Quick display of interaction: add the strain effect.

#### Without interaction:

# glmlds <- glm(nb.eggs.tot $\sim$ day+strain, data, family = "poisson") $Im(\mu) = \beta_0 + \beta_1 \times day + \beta_2 \times strain$ $\mu_{day_2/WT} = e^{\beta_0} e^{\beta_1} e^{\beta_2}$



#### With interaction:

```
glmidsI <- glm(nb.eggs.tot \sim day*strain, data, family = "poisson") 
 Im(\mu) = \beta_0 + \beta_1 \times day + \beta_2 \times strain + \beta_3 \times day \times strain 
 \mu_{day_2/WT} = e^{\beta_0} e^{\beta_1} e^{\beta_2} e^{\beta_2}
```

```
summary(qlm1dsI)
Call:
glm(formula = nb.eggs.tot ~ dav * strain. family = "poisson".
    data = data)
Deviance Residuals:
-4.6950 -1.9715 -0.6709
                            1.1131
Coefficients:
              Estimate Std. Error z value Pr(>|z|)
(Intercept)
              2.39987
dav2
              -0.51188
              -0.25447
 av2:strainWT 0.67988
                          0.11071
```

#### Models comparisons:

```
> anova(glmids, glmidsI, test = "Chisq")
Analysis of Deviance Table

Model 1: nb.eggs.tot ~ day + strain
Model 2: nb.eggs.tot ~ day * strain
Resid. Df Resid. Dev Df Deviance Pr(>Chi)
1 155 1029.53
2 154 990.53 1 39.001 4.236e-10 ***
```



### Proportion/% data

► 
$$Y \hookrightarrow Binom(n,p)$$
 with  $P(Y = k) = C_n^k (1-p)^{(n-k)}$ 



### Proportion/% data

- ►  $Y \hookrightarrow Binom(n,p)$  with  $P(Y=k) = C_n^k (1-p)^{(n-k)}$
- ▶ We are more interested on the frequency of the event  $\mu = \frac{Y}{n}$
- $\blacktriangleright$   $E(\frac{Y}{n}) = p$  and  $var(\frac{Y}{n}) = \frac{p(1-p)}{n}$



### Proportion/% data

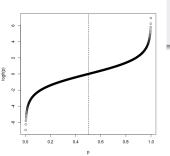
- ►  $Y \hookrightarrow Binom(n,p)$  with  $P(Y=k) = C_n^k (1-p)^{(n-k)}$
- We are more interested on the frequency of the event  $\mu = \frac{Y}{n}$
- $E(\frac{Y}{n}) = p$  and  $var(\frac{Y}{n}) = \frac{p(1-p)}{n}$
- ► The link function is the logit function, the model is:

$$logit(\mu) = log(\frac{\mu}{1-\mu}) = \beta_0 + \beta_1 X_1 + \dots + \beta_k X_k \in ] - \infty, + \infty[$$



### Proportion/% data

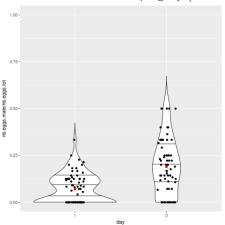
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- ►  $E(\frac{Y}{n}) = p$  and  $var(\frac{Y}{n}) = \frac{p(1-p)}{n}$
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$$\begin{split} \log\!it(\mu) &= \log\big(\frac{\mu}{1-\mu}\big) = \beta_0 + \beta_1 X_1 + \ldots + \beta_k X_k \in ] - \infty, + \infty[ \\ \text{on the proportion scale: } \mu &= \frac{1}{1+e^{-(\beta_0 + \beta_1 X_1 + \ldots + \beta_k X_k)}} \in [0,1] \end{split}$$

# Example: proportion of male worm eggs against time from Manon Grosmaire experiments

n measurements of male progeny (=nb eggs) on 2 time points.



### Is there male eggs production?

bglm0 <- glm(cbind(nb.eggs.male, nb.eggs.tot-nb.eggs.male) ~ 1,
data = data, family = "binomial")</pre>

The null model is:

$$log(\frac{\mu}{1-\mu}) = log(\frac{p_{male}}{1-p_{male}}) = log(odd) = \beta_0 \Leftrightarrow odd = e^{\beta_0}$$

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On 4 eggs, an odds equal to 3 indicates that 3 eggs against 1 will be male. The more the odds, the more the probability.

If the model is true, asymptotically, estimators are gaussian.

```
> summary(bglm0)
Call:
glm(formula = cbind(nb.eggs.male, nb.eggs.tot - nb.eggs.male) ~
   1, family = "binomial", data = data)
Deviance Residuals:
    Min
                     Median
                                   30
-2.04765 -1.01609 -0.04592 0.43340
                                        2.20961
Coefficients:
           Estimate Std. Error z value Pr(>|z|)
(Intercept) -1.89712 0.07739 -24.51 <2e-16 ***
Signif. codes: 0 '***' 0.001 '**' 0.01 '*' 0.05 '.' 0.1 ' ' 1
(Dispersion parameter for binomial family taken to be 1)
   Null deviance: 159.62 on 141 degrees of freedom
Residual deviance: 159.62 on 141 degrees of freedom
AIC: 369.72
Number of Fisher Scoring iterations: 4
```

- Recalling,  $odd = e^{\beta_0}$ .
- $IC_{95\%}(\beta_0) = \beta_0 \pm 1.96 \times \sigma_{\beta_0}$

LBI

If the model is true, asymptotically, estimators are gaussian.

```
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- Recalling,  $odd = e^{\beta_0}$ .
- $IC_{95\%}(\beta_0) = \beta_0 \pm 1.96 \times \sigma_{\beta_0}$
- ► The odd of male birth predicted is  $e^{\beta_0} \simeq 0.15[0.13, 0.17]$  which is significant (Wald test, p-value <5%).

# Example: proportion of male worm eggs against time Day effect model

### Is there a change of male eggs production according to day?

bglm1 <- glm(cbind(nb.eggs.male, nb.eggs.tot-nb.eggs.male)
~ day, data = data, family = "binomial")</pre>

The model is:

 $log(odd) = \beta_0 + \beta_1 \times day$ 

 $\triangle day$  is a factor: day = 0 for day 1, day = 1 for day 2.

- ► On linear predictor scale,  $log(odd) = \beta_0 + \beta_1 \times day$ 
  - ► For day 1: day = 0, thus  $log(odd_{day_1}) = \beta_0$

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- ► On linear predictor scale,  $log(odd) = \beta_0 + \beta_1 \times day$ 
  - ► For day 1: day = 0, thus  $log(odd_{day_1}) = \beta_0$
  - For day 2: day = 1, thus  $log(odd_{day_2}) = \beta_0 + \beta_1$
- ► Towards Odds Ratio:

$$\log\left(\frac{odd_{day_2}}{odd_{day_1}}\right) = \log(OR_{days}) = \beta_1$$

- ▶ On linear predictor scale,  $log(odd) = \beta_0 + \beta_1 \times day$ 
  - For day 1: day = 0, thus  $log(odd_{day_1}) = \beta_0$
  - For day 2: day = 1, thus  $log(odd_{day_2}) = \beta_0 + \beta_1$
- ► Towards Odds Ratio:
  - ►  $log(\frac{odd_{day_2}}{odd_{day_1}}) = log(OR_{days}) = \beta_1$
  - $OR_{days} = e^{\beta_1} = \times$  odd from day 1 to 2

- ▶ On linear predictor scale,  $log(odd) = \beta_0 + \beta_1 \times day$ 
  - For day 1: day = 0, thus  $log(odd_{day_1}) = \beta_0$
  - ► For day 2: day = 1, thus  $log(odd_{day_2}) = \beta_0 + \beta_1$
- ► Towards Odds Ratio:
  - $\log\left(\frac{odd_{day_2}}{odd_{day_1}}\right) = \log(OR_{days}) = \beta_1$
  - $OR_{days} = e^{\beta_1} = \times$  odd from day 1 to 2
  - OR<sub>days</sub> = 2.2[1.6,3.0] > 1 suggests ageing tends to favor chances to get males (Wald test, p-value <5%).</li>

```
> summary(bglm1)
glm(formula = cbind(nb.eggs.male, nb.eggs.tot - nb.eggs.male) ~
    day, family = "binomial", data = data)
Deviance Residuals:
Min 10 Median 30 Max
-2.4297 -0.8809 0.0000 0.3562 2.0473
Coefficients:
            Estimate Std. Error z value Pr(>|z|)
(Intercept) -2.3294
                         0.1252 -18.605 < 2e-16 ***
                         0.1602 5.017 5.23e-07 ***
Signif. codes: 0 '***' 0.001 '**' 0.01 '*' 0.05 '.' 0.1 ' ' 1
(Dispersion parameter for binomial family taken to be 1)
    Null deviance: 159.62 on 141 degrees of freedom
Residual deviance: 133.45 on 140 degrees of freedom
AIC: 345.55
Number of Fisher Scoring iterations: 4
```

► Is it though relevant to add a day effect?

▶ Is it though relevant to add a day effect?

Is it though relevant to add a day effect?

```
> anova(glm0, glm1, test = "Chisq")
Analysis of Deviance Table

Model 1: nb.eggs.tot ~ 1
Model 2: nb.eggs.tot ~ day
Resid. Df Resid. Dev Df Deviance Pr(>Chi)
1 157 1038.3
2 156 1030.6 1 7.6398 0.00571 **
...
Signif. codes: 0 '***' 0.001 '**' 0.01 '*' 0.05 '.' 0.1 ' ' 1
```

 $\chi^2_{obs} < \chi^2_{th}$ , the day effect model explains significantly better the variability of the data (1-pchisq(7.6398, df = 1)).

### The session is finished!



#### Thanks for coming!

- ► The next session will be on Machine Learning (co-clustering analysis) by Margot Selosse, PhD student, from Lyon II university, Foodle to come.
- ► A quick reminder about our Slack tchat.
- Any topics you want to discuss?