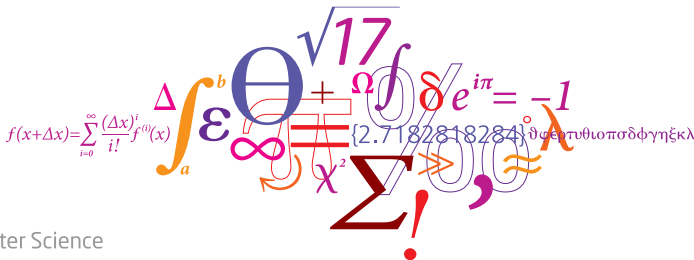


Automated and Early Detection of Disease Outbreaks

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Section for Dynamical Systems



DTU Compute

Department of Applied Mathematics and Computer Science

Outline

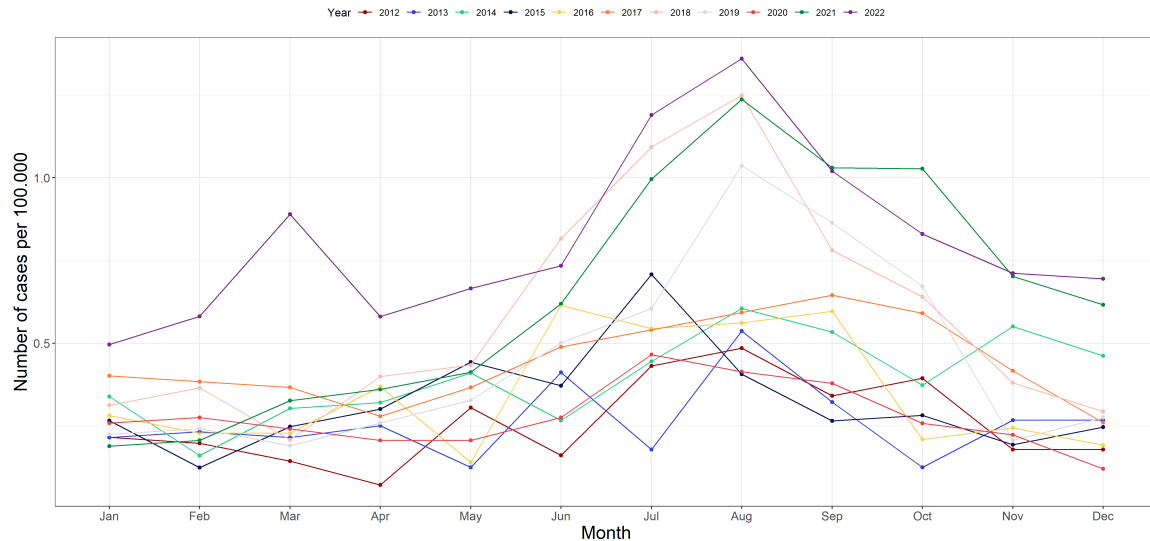
- Data exploration
 - Shiga- and verotoxin producing E. coli.
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Shiga- and verotoxin producing E. coli.

Date	ageGroup	y_{it}	x_{it}
2008-01-01	<1 year	2	64137
2008-01-01	1-4 years	2	259910
2008-01-01	5-14 years	2	680529
2008-01-01	15-24 years	1	635838
...
2022-12-01	5-14 years	5	634139
2022-12-01	15-24 years	1	721286
2022-12-01	25-64 years	10	3031374
2022-12-01	65+ years	12	1204892

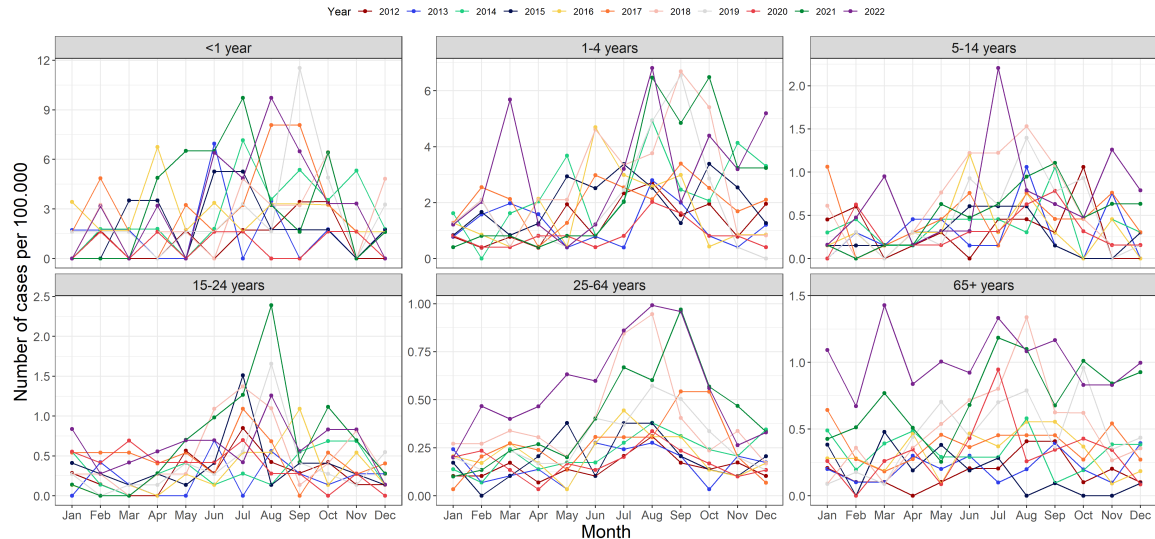
Shiga- and verotoxin producing E. coli.

Shiga- og verotoxin producerende E. coli.



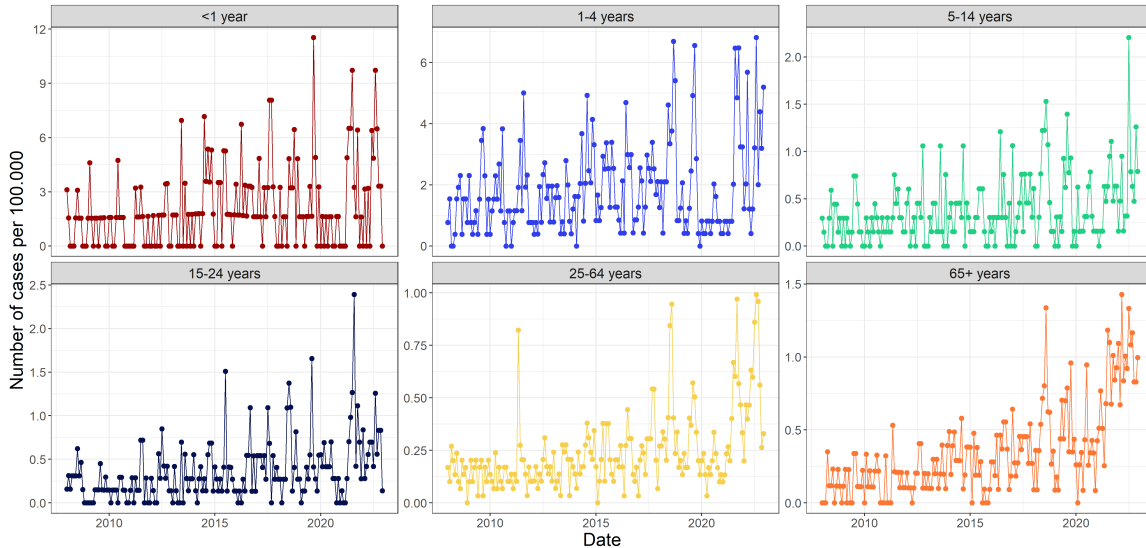
Shiga- and verotoxin producing E. coli.

Shiga- og verotoxin producerende E. coli.



Shiga- and verotoxin producing E. coli.

Shiga- og verotoxin producerende E. coli.

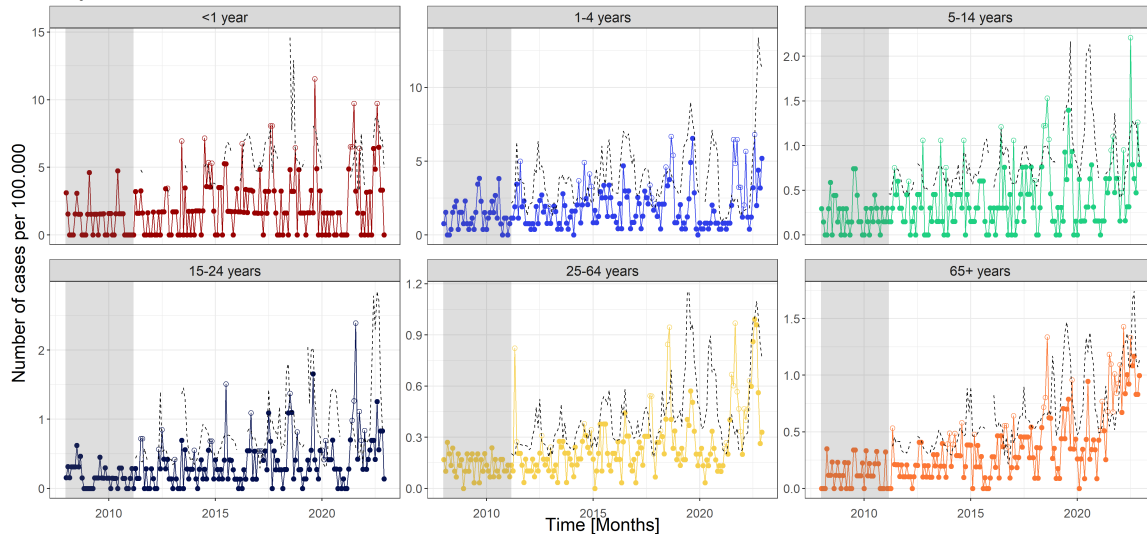


State-of-the-art methods for aberration detection is presented in Salmon, Schumacher, and Höhle 2016 and implemented in the R package **surveillance**. The R package includes methods such as the Farrington method introduced by Farrington et al. 1996 together with the improvements proposed by Noufaily et al. 2013.

State-of-the-art Farrington

Shiga- og veratoxin producerende E. coli.

Farrington method

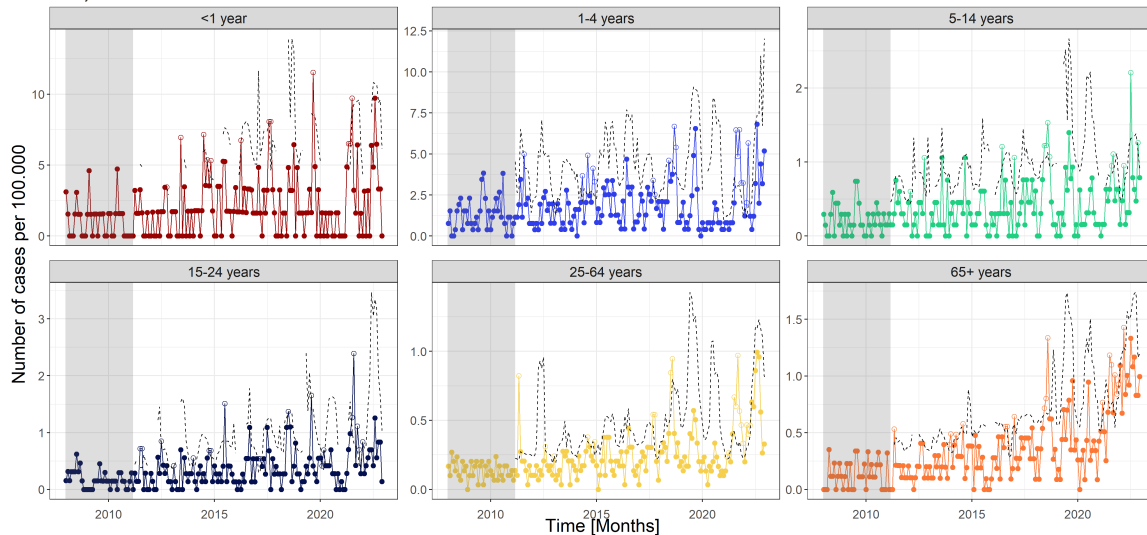


State-of-the-art

Noufailly

Shiga- og veratoxin producerende E. coli.

Noufailly method



Hierarchical Poisson-Normal model

The conditional distribution, $Y|u$, of the count observations are assumed to be a Poisson distribution with intensities λ_{it} . Also, we shall assume that the count is proportional to the population size, x_{it} , within each age group, i , at a given time point, t . Hence, in terms of the canonical link for the Poisson distribution the model for the fixed effect is

$$\log(\lambda_{it}) = \mathbf{X}_i^T \beta_{it} + \log(x_{it}) \quad (1)$$

Here \mathbf{X}_i is $T \times 6$ -dimensional, and β_{it} contains the corresponding fixed effect parameter. The random effects u_{it} are assumed to be Gaussian.

$$u_{it} = \epsilon_{it} \quad (2)$$

where $\epsilon_{it} \sim N(0, \sigma^2)$ is a white noise process, and σ is a model parameter.

Henceforth, the model can be formulated as a two-level hierarchical model

$$Y_{it}|u_{it} \sim \text{Pois}(\lambda_{it}e^{u_{it}}) \quad (3a)$$

$$u_{it} \sim N(0, \sigma^2) \quad (3b)$$

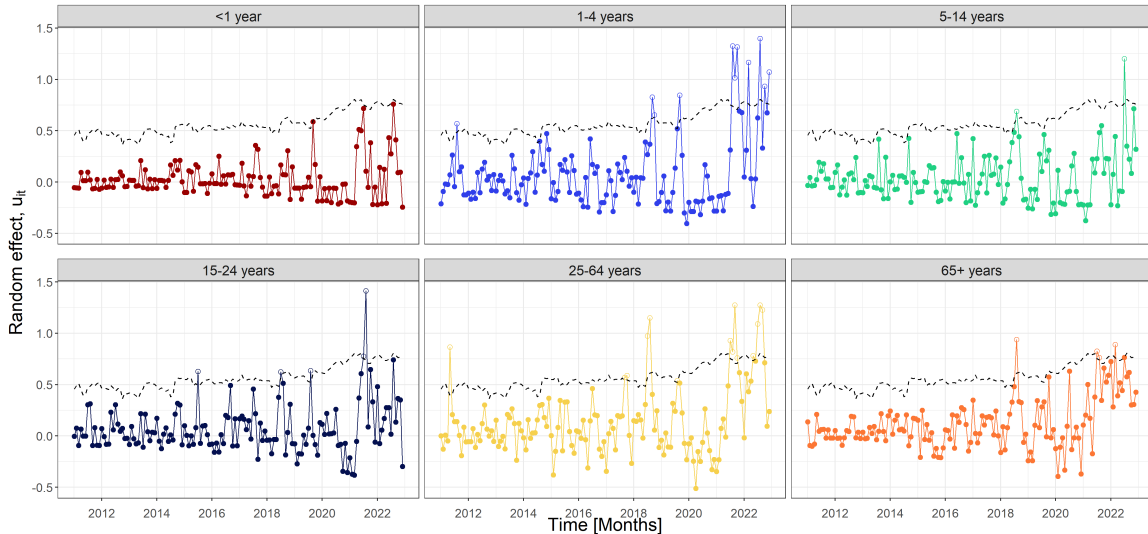
Here should be something about threshold calculation

Parameter	Estimate	Std. Error
$\beta_{<1year}$	10.83	0.11
$\beta_{1-4years}$	13.64	0.09
$\beta_{5-14years}$	13.94	0.10
$\beta_{15-24years}$	14.05	0.09
$\beta_{25-64years}$	16.61	0.08
$\beta_{65+years}$	14.83	0.09
σ	1.01	0.04

Results

Shiga- og veratoxin producerende *E. coli*.

Hierarchical Poisson Normal model



Hierachical Poisson-Gamma model

Likewise, in the compound Poisson-Gamma model the conditional distribution, $Y|u$, of the count observations are assumed to be a Poisson distribution, but this time the intensities, λ_{it} , are defined as

$$\log(\lambda_{it}) = \mathbf{X}_i^T \beta_{it} + \log(x_{it}) \quad (4)$$

Here \mathbf{X}_i is $T \times 6$ -dimensional, and β_{it} contains the corresponding fixed effect parameter. Additionally, the random effects u_{it} are assumed to be Gamma distributed.

Subsequently, the model can be formulated as a two-level hierarchical model

$$Y_{it}|u_{it} \sim \text{Pois}(\lambda_{it}u_{it}) \quad (5a)$$

$$u_{it} \sim G(1/\phi_i, \phi_i) \quad (5b)$$

$$\begin{aligned} P[Y = y_i] &= g_Y(y; \lambda, \phi) \\ &= \frac{\lambda^y}{y! \Gamma(1/\phi) \phi^{1/\phi}} \frac{\phi^{y+1/\phi} \Gamma(y + 1/\phi)}{(\lambda\phi + 1)^{y+1/\phi}} \\ &= \frac{\Gamma(y + 1/\phi)}{\Gamma(1/\phi) y!} \frac{1}{(\lambda\phi + 1)^{1/\phi}} \left(\frac{\lambda\phi}{\lambda\phi + 1} \right)^y \\ &= \binom{y + 1/\phi - 1}{y} \frac{1}{(\lambda\phi + 1)^{1/\phi}} \left(\frac{\lambda\phi}{\lambda\phi + 1} \right)^y, \text{ for } y = 0, 1, 2, \dots \end{aligned} \tag{6}$$

where we have used the convention

$$\binom{z}{y} = \frac{\Gamma(z + 1)}{\Gamma(z + 1 - y) y!} \tag{7}$$

The marginal distribution of Y is a negative binomial distribution, $Y \sim \text{NB}(1/\phi, 1/(\lambda\phi + 1))$

Inference on individual group means

Consider the hierarchical Poisson-Gamma model in (5), and assume that a value $Y = y$ has been observed. Then the conditional distribution of u for given $Y = y$ is a Gamma distribution,

$$u|Y = y \sim G(y + 1/\phi, \phi/(\lambda\phi + 1)) \quad (8)$$

with mean

$$E[u|Y = y] = \frac{y\phi + 1}{\lambda\phi + 1} \quad (9)$$

and variance

$$V[u|Y = y] = \frac{(y\phi^2 + \phi)}{(\lambda\phi + 1)^2} \quad (10)$$

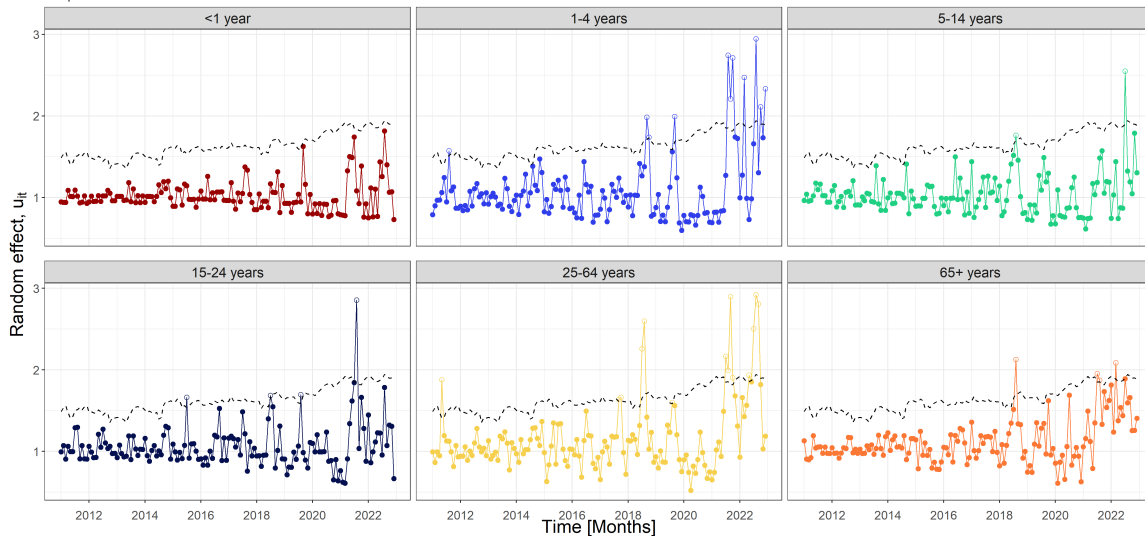
Here should be something about threshold calculation

Parameter	Estimate	Std. Error
$\beta_{<1year}$	11.23	0.08
$\beta_{1-4years}$	13.94	0.06
$\beta_{5-14years}$	14.30	0.07
$\beta_{15-24years}$	14.42	0.07
$\beta_{25-64years}$	16.87	0.06
$\beta_{65+years}$	15.24	0.06
ϕ	0.45	0.03

Results

Shiga- og veratoxin producerende E. coli.

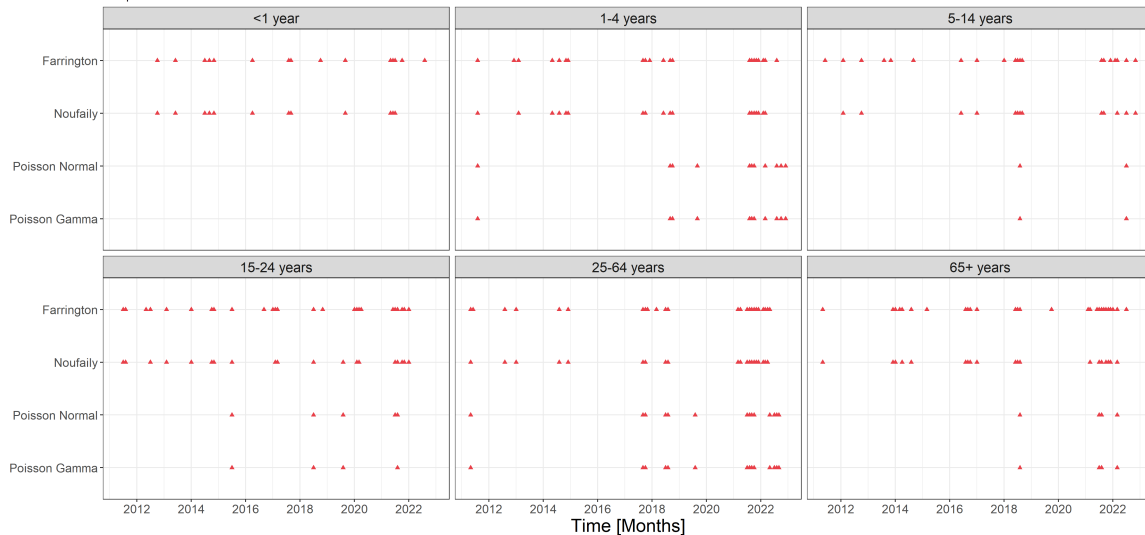
Compound Poisson Gamma model



Comparison of methods

Shiga- og veratoxin producerende E. coli.

Compare methods



- Farrington, C. P. et al. (1996). "A Statistical Algorithm for the Early Detection of Outbreaks of Infectious Disease". In: *Journal of the Royal Statistical Society. Series A (Statistics in Society)* 159.3, pp. 547–563. ISSN: 09641998, 1467985X. URL: <http://www.jstor.org/stable/2983331> (visited on 01/27/2023).
- Noufaily, Angela et al. (2013). "An Improved Algorithm for Outbreak Detection in Multiple Surveillance Systems". en. In: *Online Journal of Public Health Informatics* 32.7, pp. 1206–1222.
- Salmon, Maëlle, Dirk Schumacher, and Michael Höhle (2016). "Monitoring Count Time Series in R: Aberration Detection in Public Health Surveillance". In: *Journal of Statistical Software* 70.10, pp. 1–35. DOI: 10.18637/jss.v070.i10. URL: <https://www.jstatsoft.org/index.php/jss/article/view/v070i10>.

Proof - Probability function for Y

The probability function for the conditional distribution of Y for given u

$$f_{Y|u}(y; \lambda, u) = \frac{(\lambda u)^y}{y!} \exp(-\lambda u) \quad (11)$$

and the probability density function for the distribution of u is

$$f_u(u; \phi) = \frac{1}{\phi \Gamma(1/\phi)} \left(\frac{u}{\phi}\right)^{1/\phi-1} \exp(-u/\phi) \quad (12)$$

Proof - Probability function for Y

Given (11) and (12), the probability function for the marginal distribution of Y is determined from

$$\begin{aligned}
 g_Y(y; \lambda, \phi) &= \int_{u=0}^{\infty} f_{Y|u}(y; \lambda, u) f_u(u; \phi) du \\
 &= \int_{u=0}^{\infty} \frac{(\lambda u)^y}{y!} \exp(-\lambda u) \frac{1}{\phi \Gamma(1/\phi)} \left(\frac{u}{\phi}\right)^{1/\phi-1} \exp(-u/\phi) du \\
 &= \frac{\lambda^y}{y! \Gamma(1/\phi) \phi^{1/\phi}} \int_{u=0}^{\infty} u^{y+1/\phi-1} \exp(-u(\lambda\phi + 1)/\phi) du
 \end{aligned} \tag{13}$$

Proof - Probability function for Y

In (13) it is noted that the integrand is the *kernel* in the probability density function for a Gamma distribution, $G(y + 1/\phi, \phi/(\lambda\phi + 1))$. As the integral of the density shall equal one, we find by adjusting the norming constant that

$$\int_{u=0}^{\infty} u^{y+1/\phi-1} \exp\left(-u/\left(\phi/(\lambda\phi + 1)\right)\right) du = \frac{\phi^{y+1/\phi} \Gamma(y + 1/\phi)}{(\lambda\phi + 1)^{y+1/\phi}} \quad (14)$$

and then (6) follows

Appendix

Proof - Conditional distribution of Y

The conditional distribution is found using Bayes Theorem

$$\begin{aligned}
 g_u(u|Y=y) &= \frac{f_{y,u}(y,u)}{g_Y(y;\lambda,\phi)} \\
 &= \frac{f_{y|u}(y;u)g_u(u)}{g_Y(y;\lambda,\phi)} \\
 &= \frac{1}{g_Y(y;\lambda,\phi)} \left(\frac{(\lambda u)^y}{y!} \exp(-\lambda u) \frac{1}{\phi \Gamma(1/\phi)} \left(\frac{u}{\phi}\right)^{1/\phi-1} \exp(-u/\phi) \right) \\
 &\propto u^{y+1/\phi-1} \exp(-u(\lambda\phi+1)/\phi)
 \end{aligned} \tag{15}$$

We identify the *kernel* of the probability density function

$$u^{y+1/\phi-1} \exp(-u(\lambda\phi+1)/\phi) \tag{16}$$

as the kernel of a Gamma distribution, $G(y+1/\phi, \phi/(\lambda\phi+1))$