### Estimate RSV onset and peak timing

Tutorial 2 for transition

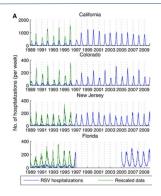
Gigi (Zhe Zheng) $^1$  Dan (Daniel Weinberger) Ginny (Virginia Pitzer)

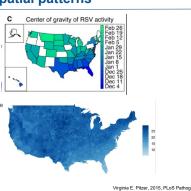
2023-01-04

 $<sup>^1</sup>z he. z heng@yale.edu; \ zhe. z heng@aya.yale.edu; \ gigi.z he. z heng@gmail.com$ 

### Background

#### Before the COVID-19 pandemic, the timing of RSV seasonal epidemics exhibited notable spatial patterns





Daniel Weinberger, 2015, CID

#### Outline

- ▶ In section 1, we will first introduce how to find the peak timing of RSV epidemics using harmonic regression (given regular annual/biennial seasonality). We will then talk about identifying the onset of RSV epidemics using second derivative method, regardless of the seasonality of RSV.
- ▶ In section 2, we will learn how to use R to identify peak timing and onset of RSV epidemics.
- ▶ In section 3, we will incorporate the spatial component of RSV epidemics. We will first introduce the concept of spatial autocorrelation and then lean to use R to account for spatial autocorrelation.

#### References

#### Relevant readings:

- RSV onset timing at county level
- RSV peak timing on state level
- Comparing RSV onset timing before and during the COVID-19 pandemic
- RSV peak timing at ZIP code level and the drivers of RSV spread

# Harmonic regression to estimate the peak timing of RSV epidemics

Note: Most of the following materials came from Dan and Ginny's Lecture notes abd Harmonic Regression by NCSS<sup>2</sup>.

Please check out:

- ▶ Dan's class: Public Health Surveillance
- ► Ginny's class: Quantitative Methods in Infectious Diseases

$$X_t = \mu + R\cos(2\pi f t + d) + e_t$$

Where

 $X_t$  is the time-series contains a periodic (cyclic) component.

 $\mu$  is mean of the series.

R is the amplitude of seasonality.

 $f = \frac{1}{\text{period}}$  is the frequency of the periodic.

d is the phase or horizontal offset.

 $e_t$  is the random error (noise) of the series.

#### t is the time step

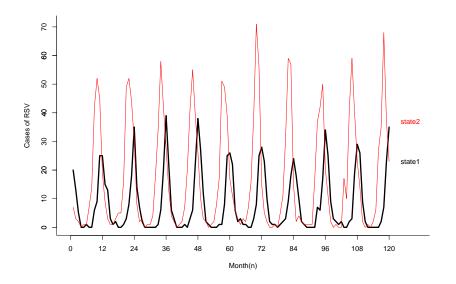
 $<sup>^2</sup> https://www.ncss.com/wp-content/themes/ncss/pdf/Procedures/NCSS/Harmonic\_Regression.pdf$ 

## Pseudo-RSV data: Simulate time series with a 12 month period

Imagine this is RSV case data from 2 states, and we want to investigate the epidemic characteristics in these states and the lag between states.

```
set.seed(123)
n=120 # 10 years
t \leftarrow seq(1,n)
amp1=2.5 # high amplitude
freq=1/12 # frequency = 1/period
amp2=2 # low amplitude
xt1a=amp1*cos(2*3.14159*t*freq)
#other series shifted by 2 months
xt2a=amp2*cos(2*3.14159*t*freq+1)
#Simulate some poisson count data
xt1=rpois(n,exp((1+xt1a)))
xt2=rpois(n,exp(2+xt2a))
```

### The observed pseudo-RSV cases over time in two states



## Investigate the epidemic characteristics of the pseudo-RSV time-serieses

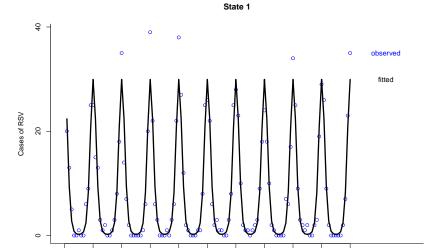
This is based on the prior knowledge that RSV has annual cycle in temperate regions and biannual cycle in high latitude regions. For other viruses or RSV circulation in other climate, you should consider using wavelet analysis to identify the periodicity first.

```
# Create the needed harmonic variables with
# 6 month, 12 month, and 24 month periodicities
t<-1:120
#Create harmonic variables
sin6=sin(2*pi*t/6)
cos6=cos(2*pi*t/6)
sin12=sin(2*pi*t/12)
cos12=cos(2*pi*t/12)
sin24=sin(2*pi*t/24)
cos24=cos(2*pi*t/24)</pre>
```

When you write the equations with sin and cos terms, you will need to include them both in an equation. For example, if the coefficient of cos12 is significant, you will need to include sin12 even though the coefficient is not significant in the summary.

## Fit a simple poisson regression with 12 month period for state 1

```
fit1a <- glm(xt1~sin12+cos12, family='poisson')
pred1a<- fitted(fit1a)
#summary(fit1a)</pre>
```



### Add in 24 month periodicity

## Number of Fisher Scoring iterations: 5

```
fit2a <- glm(xt1~sin12+cos12+sin24+cos24.family='poisson')
pred2a<- fitted(fit2a)
summary(fit2a)
##
## Call:
## glm(formula = xt1 ~ sin12 + cos12 + sin24 + cos24, family = "poisson")
##
## Deviance Residuals:
               10 Median
##
      Min
                               30
                                       Max
## -2.1406 -0.7825 -0.2894 0.3672 2.3956
##
## Coefficients:
             Estimate Std. Error z value Pr(>|z|)
##
## (Intercept) 0.94989 0.07873 12.065 <2e-16 ***
## sin12
              0.07283 0.05737 1.269 0.204
## cos12 2.44820 0.09474 25.840 <2e-16 ***
## sin24 0.04088 0.09204 0.444 0.657
## cos24 0.04591 0.03407 1.348 0.178
## ---
## Signif. codes: 0 '***' 0.001 '**' 0.05 '.' 0.1 ' ' 1
##
## (Dispersion parameter for poisson family taken to be 1)
##
##
      Null deviance: 1495.23 on 119 degrees of freedom
## Residual deviance: 101.72 on 115 degrees of freedom
## AIC: 436.78
##
```

#### Add in 6 month periodicity

## Number of Fisher Scoring iterations: 5

```
fit3a<-glm(xt1~sin12+cos12+sin24+cos24+sin6+cos6,family='poisson')
pred3a<-fitted(fit3a)
summary(fit3a)
## Call ·
## glm(formula = xt1 ~ sin12 + cos12 + sin24 + cos24 + sin6 + cos6.
      family = "poisson")
## Deviance Residuals:
               10 Median
                                        May
## -2.0585 -0.8416 -0.2304 0.4924 2.5479
## Coefficients:
              Estimate Std. Error z value Pr(>|z|)
## (Intercept) 1.00529
                         0.09101 11.045 <2e-16 ***
## sin12
              0 11920
                       0.09601 1.242
                                         0 214
## cos12
              2.32944
                       0.13971 16.673
                                         <2e-16 ***
## sin24
             0.04088 0.09204 0.444
                                         0.657
## cos24
             0.04591 0.03407 1.348
                                         0 178
             -0.04481 0.07659 -0.585
                                         0.558
## sin6
             0.08943
## cos6
                         0.07949 1.125
                                         0.261
## ---
## Signif. codes: 0 '***' 0.001 '**' 0.01 '*' 0.05 '.' 0.1 ' ' 1
##
## (Dispersion parameter for poisson family taken to be 1)
##
      Null deviance: 1495.23 on 119 degrees of freedom
## Residual deviance: 100.13 on 113 degrees of freedom
## ATC: 439.19
##
```

### Determine best model with AIC (smaller=better)

Winner is Model 1 (12 period). Also, when models have similar AIC score (within 2 points), we prefer a simpler model.

```
AIC(fit1a)

## [1] 434.8372

AIC(fit2a)

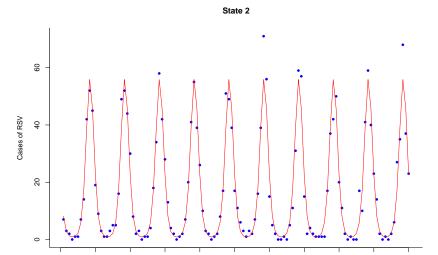
## [1] 436.7778
```

```
## [1] 439.1851
```

AIC(fit3a)

## Fit a simple poisson regression with 12 month period for state 2

```
fit1b <- glm(xt2~sin12+cos12, family='poisson')
pred1b<- fitted(fit1b)
#summary(fit1a)</pre>
```



## 

```
sin12
##
## 2,449281
beta_sin12_xt2<-coef(fit1b)['sin12']
beta cos12 xt2<-coef(fit1b)['cos12']
amp12 xt2<-sqrt(beta sin12 xt2^2+
                   beta cos12 xt2<sup>2</sup>)
amp12 xt2 #True value=2
##
      sin12
```

```
## 2.078447
```