Outbreak detection

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Outbreak/epidemic activity detection

What defines the period of epidemic outbreak/activity? When does it start e how long it lasts? - Exponential growth? - Re > 1? - Number of cases? On a noise time series, how to separate fluctuations and isolated clusters from sustained outbreak?

H1N1pdm09 in Brazil



Not knowing the operational details leading to the "first burst" in mid-June, should we classify it as season's starting date/outbreak? What if it had happened in May?

Activity thresholds

Paradigm change: Increase rate \rightarrow incidence volume

The epidemic threshold

Paradigm change: Increase rate \rightarrow incidence volume Incidence level above which there is a clear separation between baseline and sustained increase of case counts (*threshold*). This turning point can be referred to as (*pre-*)epidemic threshold.

The epidemic threshold

How to estimate it?

• Sensitivity vs specificity

The epidemic threshold

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• Sensitivity vs specificity

Do we want it to be triggered at any burst ou only when there is a clear pattern of sustained increase?

The epidemic threshold

Methodology used by the USA CDC for Influenza-like illness (ILI): - The baseline is developed by calculating the mean percentage of patient visits for ILI during non-influenza weeks for the most recent two seasons and adding two standard deviations (2021-2022 and 2022-2023). A non-influenza time period (e.g., a "non-influenza week") is defined as two or more consecutive weeks in which each week accounted for less than 2% of the season's

total number of specimens that tested positive for influenza in public health laboratories. Region-specific baselines are calculated using the same methodology. Due to the wide variability in regional level data, it is not appropriate to apply the national baseline to regional data.

http://www.cdc.gov/flu/weekly/overview.htm

The epidemic threshold – CDC

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The epidemic threshold – CDC

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Non-influenza weeks' incidence histogram



The epidemic threshold – CDC



Moving Epidemics Method (MEM)

The MEM r-package (*Moving Epidemics Method*), uses a similar idea, but with a different methodology. The package offers several methods, all of them based on the Maximum **Accumulated Percentage** (MAP) to define "active phase". This package also offers methods for other activity thresholds and seasonal trends (endemic channels).

Bibliography: - Vega T., Lozano J.E. et al. (2004) Modelling influenza epidemic - can we detect the beginning and predict the intensity and duration? International Congress Series 1263 (2004) 281-283. - Vega T., Lozano J.E. et al. (2012) Influenza surveillance in Europe: establishing epidemic thresholds by the Moving Epidemic Method. Influenza and Other Respiratory Viruses, DOI:10.1111/j.1750-2659.2012.00422.x. - Vega T., Lozano J.E. et al. (2015) Influenza surveillance in Europe: comparing intensity levels calculated using the moving epidemic method. Influenza and Other Respiratory Viruses, DOI:10.1111/irv.12330.

======= MAP Curve

For a season with S time units (weeks), calculate the maximum fraction of accumulated news cases on windows of size r = 1, 2, ..., S.

$$C(r) = \max_{i=1,S-r+1} \sum_{t=i}^{t+r-1} I_t$$
$$p(r) = \frac{C(r)}{C(S)}$$

p(r): MAP curve

From this curve, different methods can be applied to define "non-influenza weeks" and then estimate the (pre-)epidemic threshold based on the corresponding case distribution. For instance, based on a percentile of the distribution, such as 90, for instance. The higher the percentile, the higher the threshold, lowering sensitivity but increasing specificity. And vice-versa by lowering the target percentile.







The optimal window is defined by minimizing the sum of normalized r and the normalized derivative of p as an approximation for inflection point.

$$\Delta p = p^{r+1} - p^r$$

$$\hat{r} = \frac{r + min}{r_{max} - r_{min}}$$

$$\hat{\Delta p} = \frac{\Delta p - \Delta p_{min}}{\Delta p_{max} - \Delta p_{min}}$$

Optimal window:

$$r\mid \sqrt{\hat{r}^2+\hat{\Delta p}^2-(\hat{r}+\hat{\Delta p})^2/2}$$
 is minimum



MAP Curve - Method 2

Define the optimal window as the lowest in which the derivative of p is less than a value defined based on sensitivity/specificity analisis. In general, a value between 2 and 4 (empirical).

$$\Delta p = p^{r+1} - p^r$$

Window:

 $\min r \mid \Delta p < \epsilon$

MAP Curve - Method 2



Inflection point

Time in which the second derivative of the incidence curve is null, thus changing the curve's concavity.

$$\begin{split} \Delta^2 I_t &= I_{t+1} - 2I_t + I_{t-1} = 0\\ \Delta^2 I_{t-1} &> 0 , \ \Delta^2 I_{t-1} < 0 \end{split}$$

Inflection point





All nice and sweet, but...

SARI cases notified in Rondônia state (Brazil), from 2020-2016.

