Acute effects of ambient exposures

Time series and case-crossover studies

Brooke Anderson

February 3, 2016

Example data: Chicago NMMAPS

chicagoNMMAPS data

For the examples in this lecture, I'll use some data from Chicago on mortality, temperature, and air pollution. These data are available as part of the dlnm package. You can load them in R using the following code:

```
library(dlnm)
```

```
## This is dlnm 2.1.3. For details: help(dlnm) and vignette
## Important changes since version 2.0.0
```

See: 'file.show(system.file('Changesince200',package='d]

```
data("chicagoNMMAPS")
```

chicagoNMMAPS data

chic <- chicagoNMMAPS

To make the data a little easier to use, I'll rename the data frame as chic:

chicagoNMMAPS data

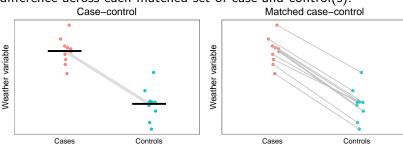
To find out more about this data, you can look at its help file:

?chicagoNMMAPS

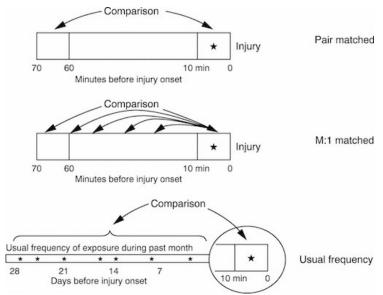
Concept: Case-crossover studies

Case-crossover models

Case-crossover model designs are based on the idea of matched case-control studies. For these, instead of comparing averages of exposure for cases versus controls, you compare the average difference across each matched set of case and control(s).



Types of case-crossover designs



Source: Sorock et al. 2001, Injury Prevention



Strata for case-crossover

Strata for a case-crossover: Year, month, day of week

3	1	2	3	4	5	6		5	6	7	8	9	10	11		2	3	4	5	6	7	8		23	24	1	2	3	4	5	3
3	4	5	6	7	1	2		4	8	9	1	2	3	4		19	20	21	15	16	17	1		23	24	25	26	27	28	22	5
3	4	5	6	7	1	2		4	8	9	10	11	12	13		19	20	21	15	16	17	18		23	24	25	26	27	28	22	10
3	4	5	6	7	1	2		4	8	9	10	11	12	13		19	20	21	15	16	17	18		23	24	25	26	27	28	22	15
3	4	5	6	7	1	2		4	8	9	10	11	12	13		19	20	21	15	16	17	18		23	24	25	26	27	28	22	20
26	27	28	29	30	1	2		31	8	9	10	11	12	13		28	29	30	15	16	17	18		26	27	28	29	30	31	22	25
May								June								July								August							

Implementation: Case-crossover studies

GLM method

One way to fit this type of model is using a GLM, but with control by strata of year-month-day of week.

Case-crossover fit using a GLM:

$$E(log(Y_t)) \sim \beta_0 + \beta_1 PM_t + \beta_2 Stratum_t$$

GLM method

To code this, first you need to create a column with the stratum. In R, you can use format with the date to do this easily, and then convert the formatted date for a factor class:

```
chic$casecross_stratum <- format(chic$date, "%Y-%m-%a")
chic$casecross_stratum <- factor(chic$casecross_stratum)
head(chic$casecross_stratum, 3)</pre>
```

```
## [1] 1987-01-Thu 1987-01-Fri 1987-01-Sat
## 1176 Levels: 1987-01-Fri 1987-01-Mon 1987-01-Sat 1987-01
```

Now you can include this factor in your model (note: this takes the place of model control for time trends and day of week in a typical time series model):

```
##
                                    Estimate Std. Error
## (Intercept)
                                4.0482946294 0.0855215089
                                0.0001909843 0.0001680322
## pm10
## casecross_stratum1987-01-Mon 0.1907876393 0.1137590495
## casecross_stratum1987-01-Sat 0.0855529446 0.1168756412
## casecross stratum1987-01-Sun 0.3300835895 0.1099033832
## casecross stratum1987-01-Thu 0.0462517003 0.1043859066
##
                                   Pr(>|t|)
## (Intercept)
                                0.00000000
                                0.255782198 ( ) ( ) ( ) ( )
## nm10
```

You can interpret the coefficients now in the same way as with the time series model:

```
pm_coef <- summary(mod_f)$coefficients["pm10", ]
100 * (exp(10 * pm_coef[1]) - 1)</pre>
```

```
## Estimate
## 0.1911668
```

Therefore, for this model, there is a 0.191% increase in mortality for an increase of 10 $\mu g/m^3$ PM10.

There are also other methods for fitting case-crossover models:

- Armstrong et al. (Conditional Poisson models: a flexible alternative to conditional logistic case cross-over analysis) suggest using a conditional Poisson regression model (gnm()) to speed up computational time.
- ► The casecross function in the season package by Adrian Barnett uses 28-day strata (rather than by month) and a Cox proportional hazards regression model to fit the model.

If you are using this method for a paper, it is worthwhile testing the different methods to see if you get similar results.

Using a conditional Poisson model:

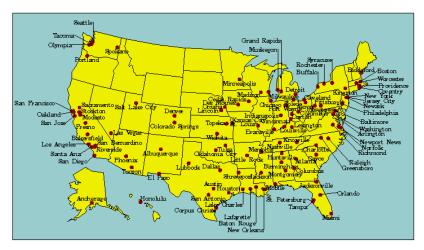
```
library(gnm)
mod_g \leftarrow gnm(cvd \sim pm10 + ns(temp, 4),
              eliminate = casecross_stratum,
              data = chic,
              family = quasipoisson())
pm_coef <- summary(mod_g)$coefficients["pm10", ]</pre>
100 * (exp(10 * pm coef[1]) - 1)
##
    Estimate
## 0.1911668
```

Using a Cox proportional hazards regression model:

```
library(season)
mod h <- casecross(cvd ~ pm10 + temp,
                    matchdow = TRUE.
                    data = chic)
## Note, irregularly spaced data...
## ...check your data for missing days
pm coef <- mod h$c.model$coefficients[1]</pre>
100 * (exp(10 * pm_coef[1]) - 1)
##
        pm10
## 0 4320228
```

Multi-city studies

NMMAPS



Source: www.ihapss.jhsph.edu

NMMAPS package

NMMAPSdata package

<u>Data</u>

- akr
- albu
- Anch

and 105 other US cities

 Meta-data on cities (population, location, counties, Census variables)

Functions

- readCity
- getMetaData and various other functions for different versions of the package

<u>Documentation</u>

- PDF users' manual
- Instructions for each function within R
- Examples for each function within R
- Website

Impact of NMMAPS

Research impacts of NMMAPS package

- ▶ As of November 2011, 67 publications had been published using this data, with 1,781 citations to these papers
- Research using NMMAPS has been used by the US EPA in creating regulatory impact statements for air pollution (particulates and ozone)
- "Thanks to NMMAPS, there is probably no other country in the world with a greater understanding of the health effects of air pollution and heat waves in its population."

Source: Barnett, Huang, and Turner, "Benefits of Publicly Available Data", Epidemiology 2012