An introduction to *outbreaker* 1.1-0

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Abstract

This vignette introduces the main functionalities of *outbreaker* [1], a package implementing a model for disease outbreak reconstruction using epidemiological data and pathogen genome sequences. The emphasis of this document is put on using *outbreaker* and on the visualization and interpretation of results.

More resources for disease outbreaks analysis in R are available on the *R-epi project*: http://sites.google.com/site/therepiproject. Questions about *outbreaker*, alonside any question relating to the analysis of epidemics, should be asked on the R-epi forum. To do so, send an email to: r-epi@googlegroups.com, or visit the website: http://groups.google.com/forum/#!forum/r-epi.

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1 Running outbreaker

1.1 A simple example

In this vignette, we shall use the toy dataset fakeOutbreak distributed with *outbreaker*. We first load the package, the dataset, and look at the object's content:

```
library(outbreaker)
data(fakeOutbreak)
class(fakeOutbreak)

## [1] "list"

names(fakeOutbreak)

## [1] "dat" "w" "collecDates" "res"

class(fakeOutbreak$dat)

## [1] "simOutbreak"

fakeOutbreak$w

## [1] 0.00 0.50 1.00 0.75

fakeOutbreak$collecDates

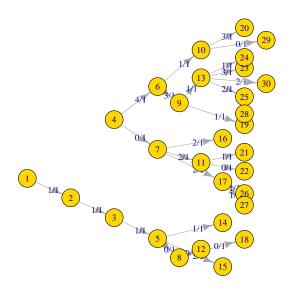
## [1] 3 5 6 6 7 9 8 9 9 9 11 10 10 10 10 11 11 12 11 13 12 13 11 12 11

## [26] 11 13 12 14 14
```

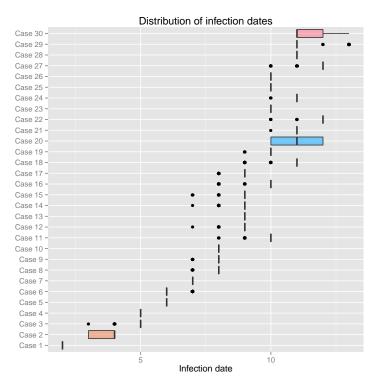
fakeOutbreak is a list containing dat, a simulated outbreak (obtained by simOutbreak), a generation time distribution (w), collection dates for the DNA sequences (collecDates), and results from *outbreaker* (res). We shall need only the first 3 items, and re-create the fourth.

```
dat <- fakeOutbreak$dat
w <- fakeOutbreak$w
collecDates <- fakeOutbreak$collecDates
plot(dat, main="Simulated outbreak")</pre>
```

Simulated outbreak



barplot(w, main="Generation time distribution", ylab="probability", xlab="days", names=0:3)



We run outbreaker on these data, using the parallel version, and specifying that we want to run 4 MCMC in parallel and 100,000 chains for each MCMC.

```
res <- outbreaker.parallel(n.runs=4, dna=dat$dna, dates=collecDates,w.dens=w, n.iter=1e5)
```

```
names(res$chains)
     [1] "step"
                   "post"
                                        "prior"
##
                              "like"
                                                   "mu1"
                                                              "mu2"
                                         "spa2"
     [7] "gamma"
                   "pi"
                              "spa1"
                                                   "Tinf_1"
                                                              "Tinf_2"
##
                                        "Tinf_6"
                   "Tinf_4"
                              "Tinf_5"
    [13] "Tinf_3"
                                                   "Tinf_7"
                                                              "Tinf_8"
##
   [19] "Tinf_9"
                   "Tinf_10" "Tinf_11"
                                        "Tinf_12"
                                                   "Tinf_13"
                                                              "Tinf_14"
##
   [25] "Tinf_15" "Tinf_16" "Tinf_17" "Tinf_18" "Tinf_19"
                                                              "Tinf_20"
   [31] "Tinf_21" "Tinf_22" "Tinf_23" "Tinf_24" "Tinf_25"
                                                              "Tinf_26"
##
                                                              "alpha_2"
##
   [37] "Tinf_27" "Tinf_28" "Tinf_29" "Tinf_30" "alpha_1"
##
   [43] "alpha_3" "alpha_4" "alpha_5" "alpha_6" "alpha_7" "alpha_8"
   [49] "alpha_9" "alpha_10" "alpha_11" "alpha_12" "alpha_13" "alpha_14"
##
   [55] "alpha_15" "alpha_16" "alpha_17" "alpha_18" "alpha_19" "alpha_20"
##
   [61] "alpha_21" "alpha_22" "alpha_23" "alpha_24" "alpha_25" "alpha_26"
##
   [67] "alpha_27" "alpha_28" "alpha_29" "alpha_30" "kappa_1" "kappa_2"
##
   [73] "kappa_3" "kappa_4" "kappa_5" "kappa_6" "kappa_7"
   [79] "kappa_9" "kappa_10" "kappa_11" "kappa_12" "kappa_13" "kappa_14"
   [85] "kappa_15" "kappa_16" "kappa_17" "kappa_18" "kappa_19" "kappa_20"
##
  [91] "kappa_21" "kappa_22" "kappa_23" "kappa_24" "kappa_25" "kappa_26"
   [97] "kappa_27" "kappa_28" "kappa_29" "kappa_30" "run"
```

The object res is a list with a number of named items, described in ?outbreaker. The most important one is res\$chains, containing the MCMC outputs:

```
class(res$chains)
## [1] "data.frame"
dim(res$chains)
## [1] 804 101
names(res$chains)
```

```
[1] "step"
                     "post"
                                "like"
                                            "prior"
                                                        "mu1"
                                                                   "mu2"
                     "pi"
                                "spa1"
                                            "spa2"
                                                        "Tinf_1"
                                                                   "Tinf_2"
##
     [7] "gamma"
        "Tinf_3"
                     "Tinf_4"
                                "Tinf_5"
                                            "Tinf_6"
##
    [13]
                                                        "Tinf_7"
                                                                   "Tinf_8"
##
    [19] "Tinf_9"
                     "Tinf_10"
                                "Tinf_11"
                                            "Tinf_12"
                                                       "Tinf_13"
                                                                   "Tinf_14"
##
    [25] "Tinf 15"
                     "Tinf 16"
                                "Tinf 17"
                                            "Tinf_18"
                                                        "Tinf 19"
                                                                   "Tinf 20"
##
    [31]
         "Tinf_21"
                     "Tinf_22"
                                "Tinf_23"
                                            "Tinf_24"
                                                        "Tinf_25"
                                                                   "Tinf_26"
##
    [37]
         "Tinf_27"
                     "Tinf_28"
                                "Tinf_29"
                                            "Tinf_30"
                                                        "alpha_1"
                                                                   "alpha_2"
                                                        "alpha_7"
    [43] "alpha_3"
                                "alpha_5"
                                            "alpha_6"
                                                                   "alpha_8"
##
                     "alpha_4"
##
    [49] "alpha_9"
                     "alpha_10"
                                "alpha_11"
                                            "alpha_12"
                                                        "alpha_13"
                                                                   "alpha_14"
    [55] "alpha_15" "alpha_16"
##
                                "alpha_17"
                                            "alpha_18"
                                                        "alpha_19"
                                                                   "alpha_20"
##
    [61] "alpha_21" "alpha_22" "alpha_23" "alpha_24" "alpha_25"
                                                                   "alpha_26"
##
    [67] "alpha_27" "alpha_28" "alpha_29" "alpha_30" "kappa_1"
                                                                   "kappa_2"
                                "kappa_5"
                                            "kappa_6"
                                                        "kappa_7"
##
    [73] "kappa_3"
                     "kappa_4"
                                                                   "kappa_8"
##
    [79]
         "kappa_9"
                     "kappa_10" "kappa_11" "kappa_12" "kappa_13"
                                                                   "kappa_14"
##
    [85] "kappa_15" "kappa_16" "kappa_17" "kappa_18" "kappa_19" "kappa_20"
    [91] "kappa_21" "kappa_22" "kappa_23" "kappa_24" "kappa_25" "kappa_26"
##
    [97] "kappa_27" "kappa_28" "kappa_29" "kappa_30" "run"
res$chains[1:10,1:10]
              post
##
      step
                       like prior
                                        mu1
                                                   mu2 gamma
                                                                  pi spa1 spa2
## 1
         1 -1106.3 -1108.4 2.093 5.000e-05 5.000e-05
                                                            1 0.9770
                                                                        0
                                                                              0
## 2
       500
            -468.0
                    -470.2 2.144 5.021e-05 5.021e-05
                                                            1 0.9826
                                                                        0
                                                                              0
            -446.9
                    -448.8 1.909 5.038e-05 5.038e-05
## 3
      1000
                                                            1 0.9572
                                                                              0
## 4
      1500
            -446.7
                    -448.9 2.294 5.060e-05 5.060e-05
                                                            1 0.9990
                                                                        0
                                                                              0
      2000
            -446.5 -448.7 2.227 5.087e-05 5.087e-05
                                                            1 0.9916
## 5
                                                                        0
                                                                              0
## 6
      2500
            -446.6 -448.7 2.147 5.080e-05 5.080e-05
                                                            1 0.9829
                                                                        0
                                                                              0
  7
      3000
            -446.9
                    -449.1 2.216 5.138e-05 5.138e-05
                                                              0.9905
                                                                        0
                                                                              0
      3500
            -448.5
                    -450.6 2.168 5.078e-05 5.078e-05
                                                            1 0.9852
                                                                        0
                                                                              0
## 8
## 9
      4000
            -446.8 -449.1 2.233 5.309e-05 5.309e-05
                                                            1 0.9923
                                                                        0
                                                                              0
            -452.0 -453.8 1.855 5.323e-05 5.323e-05
## 10 4500
                                                            1 0.9515
                                                                              0
```

The columns of this data.frame store the following outputs:

- step: the MCMC iteration of the sample
- post/like/prior: log values for posterior, likelihood, and prior densities
- mu1: in mutation model 1, mutation rate; otherwise, the rate of transitions, per site and generation
- mu2: in mutation model 1, mutation rate (mu1=mu2); otherwise, the rate of transversions, per site and generation
- gamma: the ratio between transversions and transitions (μ_2/μ_1)
- pi: the proportion of the transmission tree sampled
- Tinf_[number]: dates of infection
- alpha_[number]: the index of the ancestral cases (infectors)
- kappa_[number]: the number of generations between cases and their most recent sampled ancestor (here, fixed to 1)
- run: for parallel runs, the index of the run.

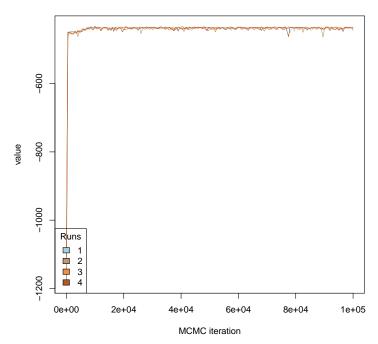
We shall see how this information can be used, visualized and interpreted over the following sections.

1.2 Assessing convergence and determining the burnin

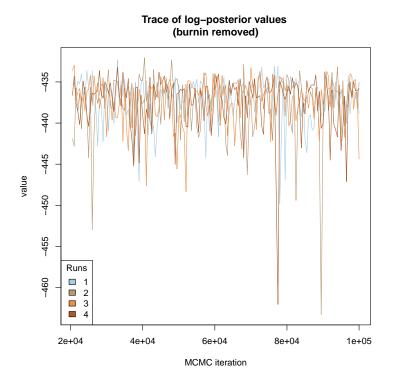
A MCMC is said to converge when it reaches a stationary state, i.e. its distributional properties are constant over time (mean and variance don't depend on which window of the MCMC you consider). Convergence of the chains is best assessed by comparing parallel runs. This can be done using plotChains, which we to visualize the trace of the log-posterior values of the model:

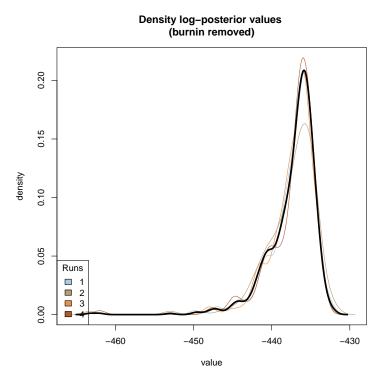
plotChains(res, main="Trace of log-posterior values")





We set the burnin to a conservative 20,000 steps, and plot the MCMC output as trace and as densities:





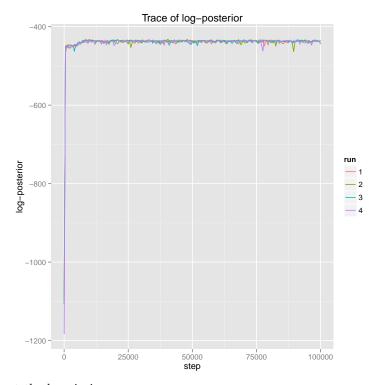
Here, all four runs have sampled from the same distribution, confirming convergence of the MCMCs. Note that this can also be tested using a simple ANOVA, which we use to check that log-posterior values do no differ from one run to another:

These graphs can be slightly improved by using ggplot2 [2]. We first load the package, and make sure that run is treated as a factor:

```
library(ggplot2)
library(reshape2)
library(mgcv)
x <- res$chains
x$run <- factor(x$run)</pre>
```

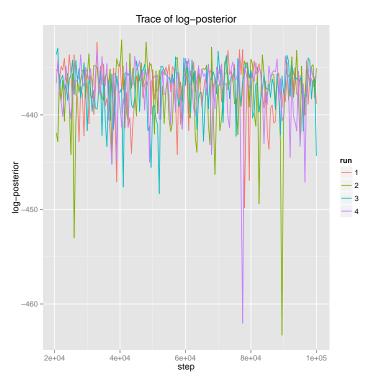
The basic plot of the log-posterior trace is obtained by:

```
p <- ggplot(x, aes(x=step)) +
    geom_line(aes(y=post, colour=run)) +
    labs(title="Trace of log-posterior", y="log-posterior")
p</pre>
```



The version without the burnin is:

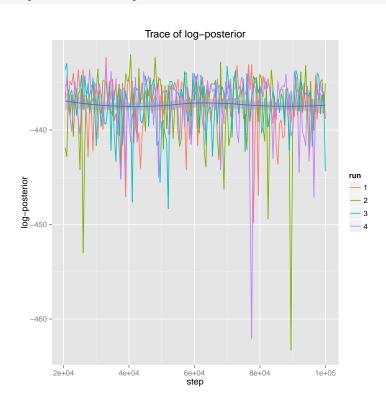
```
p <- ggplot(x[x$step>2e4,], aes(x=step)) +
    geom_line(aes(y=post, colour=run)) +
    labs(title="Trace of log-posterior", y="log-posterior")
p
```



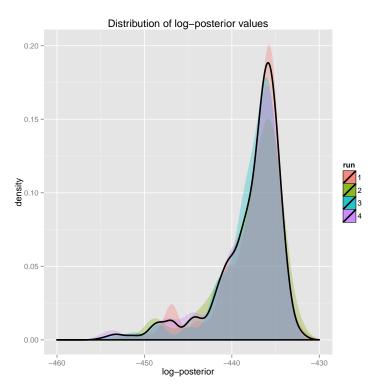
A model of the mean can be added easily:

```
p + geom_smooth(aes(y=post))
```

$geom_smooth$: method="auto" and size of largest group is <1000, so using loess. Use 'method = x' to change the smoothing method.

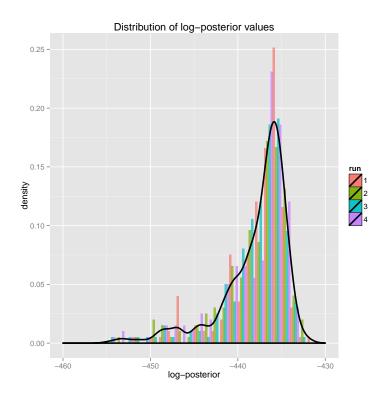


ggplot 2 is also pretty good for plotting distributions. Here is an example using 1-dimensional density estimation:



and another version using histograms:

```
p + geom_histogram(aes(x=post, fill=run, y=..density..), alpha=.7, colour=NA, position="dodge") +
    geom_density(aes(x=post), size=1, colour="black", shape=2, alpha=.8)
## stat_bin: binwidth defaulted to range/30. Use 'binwidth = x' to adjust this.
```



2 Interpreting the results

outbreaker can provide information on:

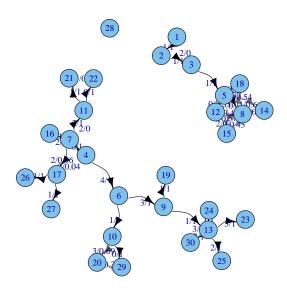
- the transmission tree ("who infected whom"); ancestry for case i is noted α_i
- the dates of infection; for case i, noted T_i^{inf}
- the mutation rate per generation of infection (μ)
- the proportion of the outbreak sampled (π)
- effective reproduction numbers over time or at an individual level [†]
- incidence curves †
- the mutation rate per unit of time †

Items indicated with a † are not explicitely modelled by *outbreaker*, but can be derived from the posterior samples of trees and parameters.

2.1 Visualizing reconstructed transmission trees

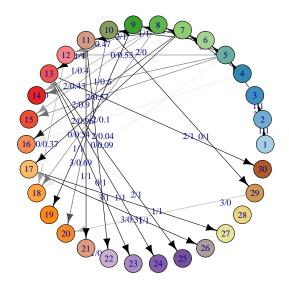
outbreaker being a Bayesian approach, it does not return a single tree, but a distribution of plausible transmission trees. A set of ancestries can be visualized by transGraph

```
library(igraph)
library(adegenet)
```



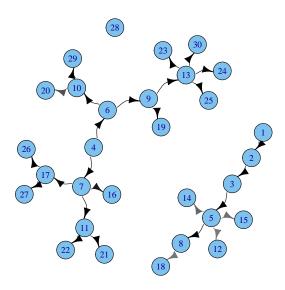
Annotations represent the number of mutations for the ancestries, and their support (frequency in the posterior samples). Note that this function returns an *igraph* object, which can be used for further plotting and customization:

```
## IGRAPH DN-- 30 40 --
## + attr: name (v/c), dates (v/n), label (v/n), color (e/c), support
## (e/n), curved (e/x), nb.mut (e/n), label (e/c)
plot(g, layout=layout.circle, edge.curved=FALSE, vertex.color=funky(30))
```



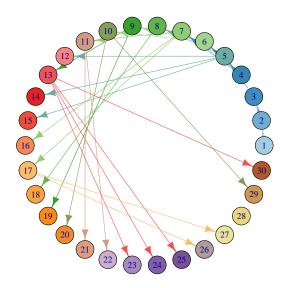
see <code>?plot.igraph</code> and <code>igraph.plotting</code> for more information on how to customize these graphics. We illustrate some possibilities below:

g <- transGraph(res, thres=0.5, annot="")



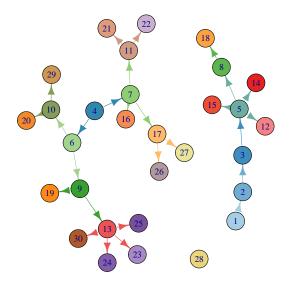
```
edge.colors <- funky(30)[as.numeric(get.edgelist(g)[,1])]
plot(g, layout=layout.circle, edge.curved=FALSE, vertex.color=funky(30),
        edge.color=edge.colors)
title("Ancestries with support >50% - circular graph")
```

Ancestries with support >50% - circular graph



```
plot(g, layout=layout.auto, edge.curved=FALSE, vertex.color=funky(30),
        edge.color=edge.colors)
title("Ancestries with support >50% - other layout")
```

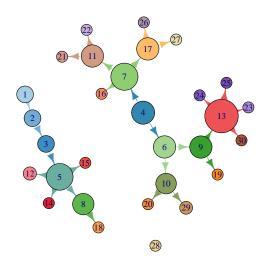
Ancestries with support >50% - other layout



Here we ensure that cases with higher reproduction numbers look bigger:

```
case.size <- 10+apply(get.R(res),2,mean)*5
plot(g, layout=layout.auto, edge.curved=FALSE, vertex.color=funky(30),
        edge.color=edge.colors, vertex.size=case.size)
title("Ancestries with support >50% \n(node size reflects R)")
```

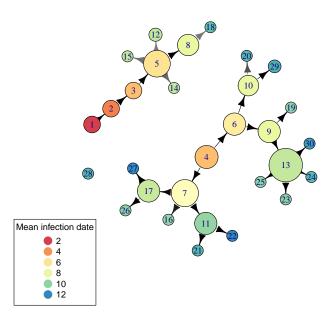
Ancestries with support >50% (node size reflects R)



Same idea, but this time colors represent dates of infection:

```
Tinf <- x[x$step>=2e4,grep("Tinf", names(x))]
case.color <- any2col(apply(Tinf,2,mean), col.pal=spectral)
plot(g, layout=layout.auto, edge.curved=FALSE, vertex.color=case.color$col,
    vertex.size=case.size)
title("Ancestries with support >50% \n(node size reflects R)")
legend("bottomleft", col=case.color$leg.col, leg=case.color$leg.txt,
    title="Mean infection date", pch=20, pt.cex=3, inset=-.1)
```

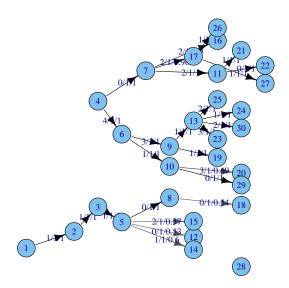
Ancestries with support >50% (node size reflects R)



The same can be done with a tree formed by consensus ancestries, obtained by get.tTree:

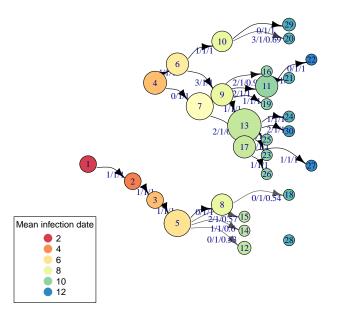
```
plot(get.tTree(res), main="Consensus ancestries - basic plot")
```

Consensus ancestries - basic plot



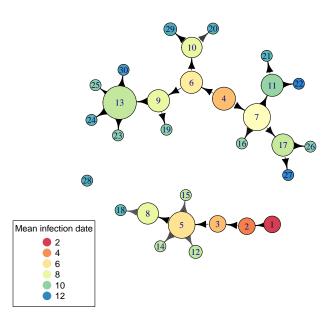
For a customized version where colors represent the mean infection dates:

Consensus ancestries (x-axis represents time)



and another, without the time axis:

Consensus ancestries



Note that all these plots can be visualized interactively using tkplot (just replace plot with tkplot in the above command lines). In all these graphs, we can see that cases 1, 4 and 28 have been classified as imported cases. For a more systematic assessment of imported cases, we can just look for cases for which the ancestor is unknown (NA) in the consensus tree:

```
temp <- get.tTree(res)
temp$ances

## [1] NA 1 2 NA 3 4 4 5 6 6 7 5 9 5 5 7 7 8 9 10 11 11 13 13 13
## [26] 17 17 NA 10 13

which(is.na(temp$ances))
## [1] 1 4 28</pre>
```

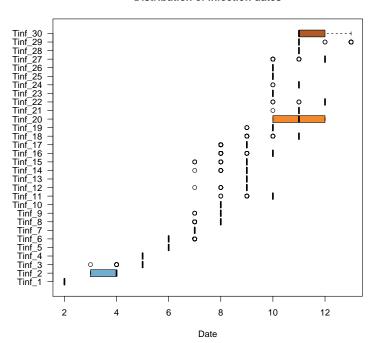
2.2 Plotting dates of infection

Dates of infection are stored in the table of MCMC outputs, with columns starting with Tinf:

```
Tinf <- x[x$step>=2e4,c(1,ncol(x),grep("Tinf", names(x)))]
Tinf[1:5,1:6]
       step run Tinf_1 Tinf_2 Tinf_3 Tinf_4
##
## 41 20000
                      2
                             4
                                    5
## 42 20500
                      2
                             4
                                    5
                                            5
              1
## 43 21000
              1
                      2
                             4
                                    5
                                            5
## 44 21500
              1
                      2
                             4
                                    5
                                            5
                      2
                             3
## 45 22000
                                    5
```

This information can be visualized easily using the basic boxplot:

Distribution of infection dates

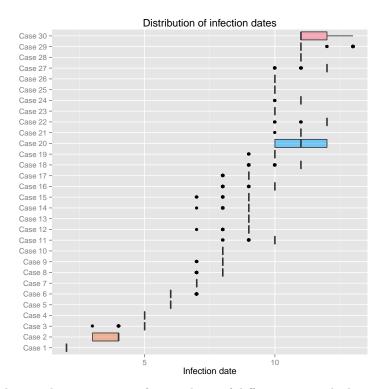


We can also use *ggplot2*, but this demands a slight reformating of the data:

```
Tinf <- melt(Tinf, id=1:2)</pre>
names(Tinf)[3:4] <- c("case", "date")</pre>
Tinf$case <- sub("Tinf_","Case ", Tinf$case)</pre>
Tinf$case <- factor(Tinf$case, levels=paste("Case",1:30))</pre>
head(Tinf)
##
      step run
                   case date
## 1 20000
              1 Case 1
                            2
## 2 20500
                            2
              1 Case 1
                            2
## 3 21000
              1 Case 1
## 4 21500
              1 Case 1
                            2
## 5 22000
                            2
              1 Case 1
                            2
## 6 22500
              1 Case 1
tail(Tinf)
##
                          case date
            step run
## 19315
           97500
                    4 Case 30
                                 11
## 19316
           98000
                    4 Case 30
                                 12
## 19317
           98500
                    4 Case 30
                                 12
## 19318
                    4 Case 30
                                 12
           99000
## 19319
           99500
                    4 Case 30
                                 12
## 19320 100000
                                 11
                   4 Case 30
```

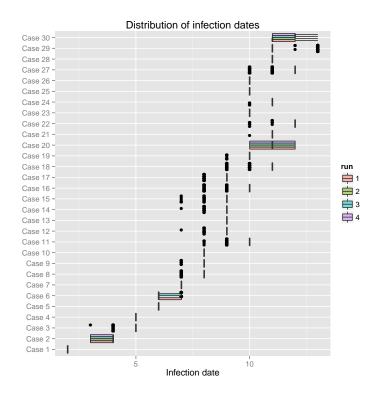
In this case, little is gained in the new plot:

```
p <- ggplot(data=Tinf) + geom_boxplot(aes(x=case,y=date,fill=case), alpha=.5) +
    coord_flip() + labs(y="Infection date", x="", title="Distribution of infection dates")
p + guides(fill=FALSE)</pre>
```



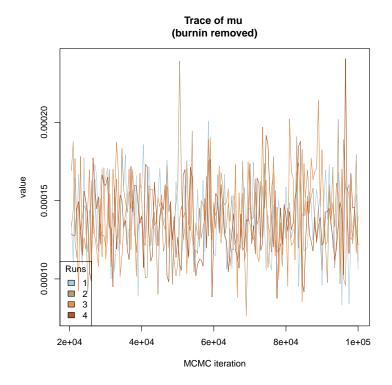
However, it would be simple to compare infection dates of different runs, which may be useful if different runs provide slightly different results:

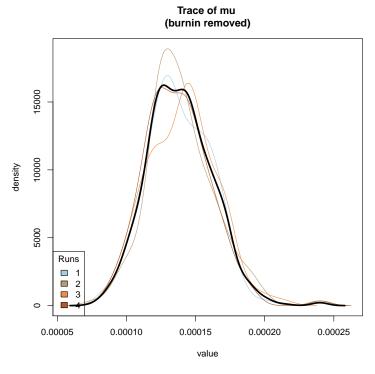
```
ggplot(data=Tinf) + geom_boxplot(aes(x=case,y=date,fill=run),alpha=.5) + coord_flip() +
    labs(y="Infection date", x="", title="Distribution of infection dates")
```



2.3 Accessing posterior distributions

Any element of the model in res\$chains can be plotted using plotChains: it just needs to be named in the argument what. For instance, the mutation rate:





(note that in this case, the plotted information is the mutation rate per generation of infection, and not per unit of time. See section on mutation rates below for an estimation of the rates per unit of time.

To derive statistics for a given distribution, one just needs to extract the relevant column, making sure to remove the burnin:

```
mu <- res$chains$mu1[res$chains$step>2e4]
head(mu)

## [1] 0.0001356 0.0001104 0.0001644 0.0001324 0.0001667 0.0001252

length(mu)

## [1] 640
head(mu)

## [1] 0.0001356 0.0001104 0.0001644 0.0001324 0.0001667 0.0001252

summary(mu)

## Min. 1st Qu. Median Mean 3rd Qu. Max.
## 7.66e-05 1.22e-04 1.38e-04 1.39e-04 1.54e-04 2.41e-04
```

Here we have 640 values to estimate parameters of the distribution of mu. However, the effective sample size might be smaller if successive values are correlated (autocorrelated chains). This can be tested easily:

```
cor(mu[-length(mu)], mu[2:length(mu)])
## [1] 0.02841
cor.test(mu[-length(mu)], mu[2:length(mu)])
##
##
   Pearson's product-moment correlation
##
## data: mu[-length(mu)] and mu[2:length(mu)]
## t = 0.7173, df = 637, p-value = 0.4735
## alternative hypothesis: true correlation is not equal to 0
## 95 percent confidence interval:
## -0.04926 0.10574
## sample estimates:
##
       cor
## 0.02841
```

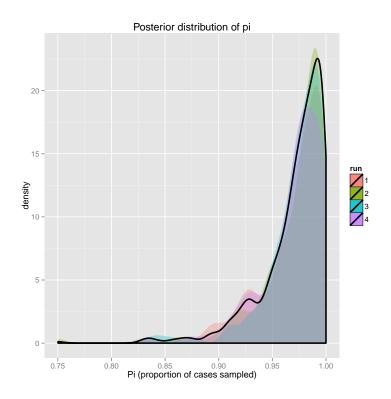
No, there is no correlation between successive values.

As before, ggplot2 versions of the plot can be obtained; here, for the parameter π (proportion of the outbreak sampled):

```
library(ggplot2)
library(reshape2)
x <- res$chains[x$step>2e4,]
x$run <- factor(x$run)</pre>
```

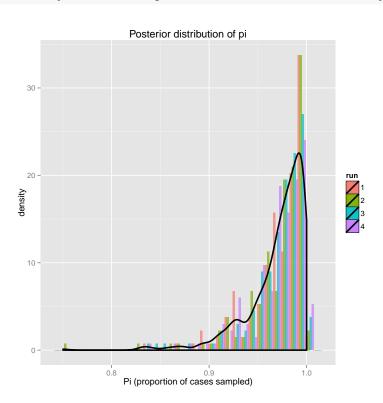
To plot densities:

```
p <- ggplot(data=x) + labs(title="Posterior distribution of pi", x="Pi (proportion of cases sampled)")
p + geom_density(aes(x=pi, fill=run), alpha=.3, colour=NA) +
    geom_density(aes(x=pi), size=1, colour="black", shape=2, alpha=.8)</pre>
```



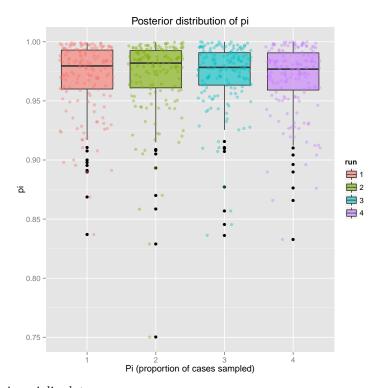
${\bf Histograms:}$

```
p + geom_histogram(aes(x=pi, fill=run, y=..density..), alpha=.7, colour=NA, position="dodge") +
    geom_density(aes(x=pi), size=1, colour="black", shape=2, alpha=.8)
## stat_bin: binwidth defaulted to range/30. Use 'binwidth = x' to adjust this.
```



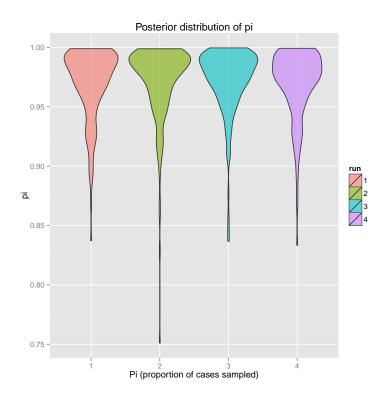
Using different boxplots for different runs:

```
p + geom_boxplot(aes(x=run, y=pi, fill=run),alpha=.6) +
    geom_jitter(aes(x=run, y=pi, col=run),alpha=.4)
```



Same idea, but using violinplots:

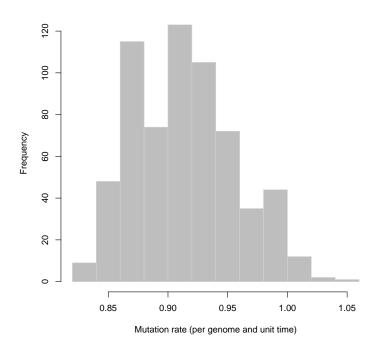
```
p + geom_violin(aes(x=run, y=pi, fill=run),alpha=.6)
```



2.4 Mutation rates

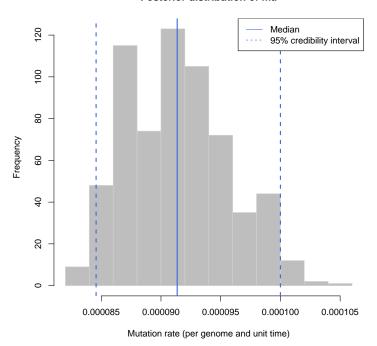
As mentioned before, mutation rates in *outbreaker*'s model are expressed per generation of infection. However, mutation rates per unit of time are biologically easier to interpret. These can be obtained using get.mu:

Posterior distribution of mu

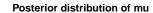


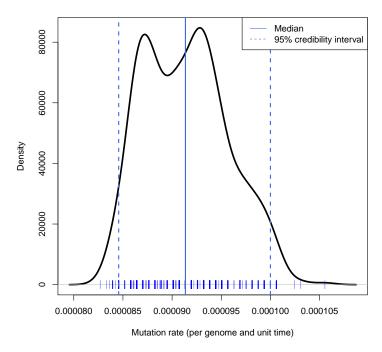
Re-expressing the rates per nucleotide, and adding the media and 95% credibility interval to the histogram:

Posterior distribution of mu



Same idea, using a one-dimensional density estimation:

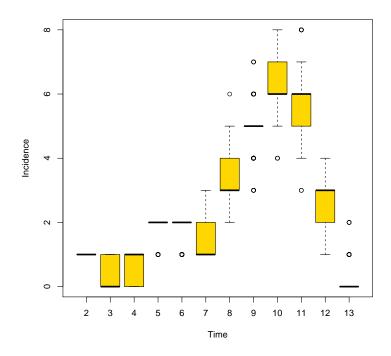




2.5 Incidence and reproduction numbers

Incidence curves and effective reproduction numbers can be derived from the results of *outbreaker*. Incidence curves can be obtained and visualized using get.incid:

incid <- get.incid(res, burnin=2e4)</pre>



```
class(incid)
## [1] "matrix"
dim(incid)
## [1] 12 640
incid[,1:10]
##
      init
## 2
         1 1 1 1 1 1 1 1 1 1
         0 0 0 1 0 0 1 0 0 0
## 3
## 4
         1 1 1 0 1 1 0 1 1 1
## 5
         2 2 2 2 2 2 2 2 2 2
         2 2 2 2 2 2 2 2 2 2
## 7
         1 1 1 1 1 1 1 1 1 1
## 8
         4 3 3 3 4 4 3 3 4 3
## 9
         4 5 6 6 4 4 5 7 4 6
## 10
         7 7 5 6 6 6 6 6 6 5
         6 5 6 5 7 7 6 4 7 7
## 11
## 12
         2 3 3 3 2 2 3 3 2 2
         0 0 0 0 0 0 0 0 0 0
## 13
```

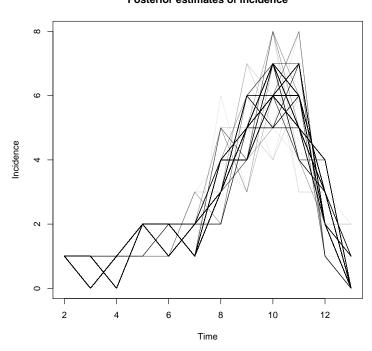
Each column in incid is a different realization of an incidence curve corresponding to one step of the MCMC. Other graphical options are available:

```
args(get.incid)
## function (x, burnin = 20000, plot = TRUE, type = c("boxplot",
## "lines"), lines = FALSE, fill.col = "gold", lines.col = transp("grey"),
```

```
## ...)
## NULL

incid <- get.incid(res, type="lines",lines.col=transp("black",.1))
title("Posterior estimates of incidence")</pre>
```

Posterior estimates of incidence

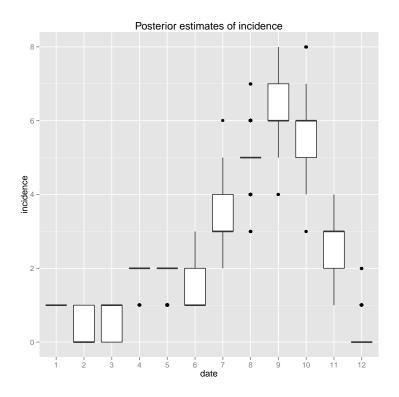


Because a lot of these trajectories overlap, visualizing them all is a bit tricky. ggplot2 will be helpful here. We first reformat the data:

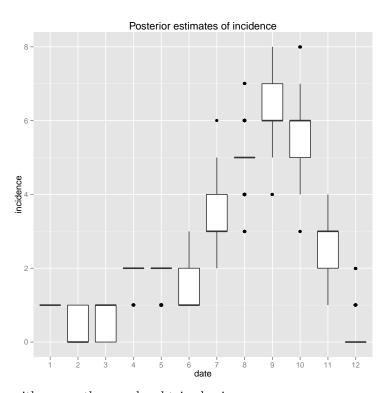
```
x <- data.frame(date=as.vector(row(incid)),</pre>
                 step=as.vector(col(incid)),
                 incidence=as.vector(incid))
head(x)
##
     date step incidence
## 1
        1
              1
## 2
              1
## 3
        3
              1
                         1
## 4
        4
              1
## 5
        5
              1
                         2
## 6
                         1
p <- ggplot(data=x, aes(x=date, y=incidence)) + labs(title="Posterior estimates of incidence")</pre>
```

This is for a basic boxplot:

```
p + geom_boxplot(aes(x=factor(date)))
```

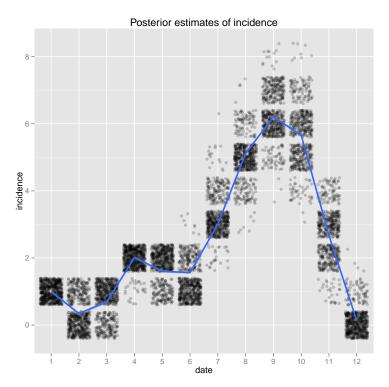


p + geom_boxplot(aes(x=factor(date)))



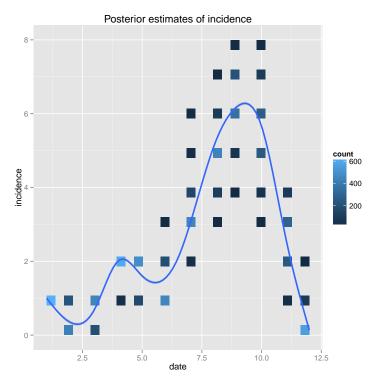
A noisified version with a smoother can be obtained using:

```
library(splines)
p + geom_jitter(aes(x=factor(date)), alpha=.2) + geom_smooth(method=lm, formula=y~ns(x,10), size=1)
```



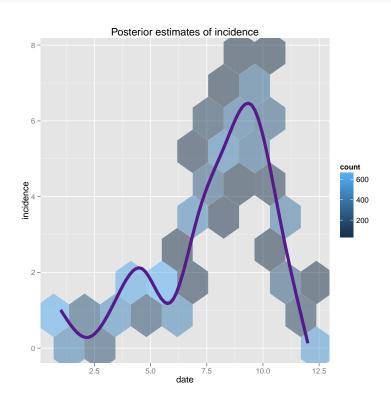
However, it adds artifactual variation and may be best avoided here. Another interesting option is offered by some density-based graphics:

```
p + geom_bin2d() + geom_smooth(method=lm, formula=y~ns(x,10), size=1)
```



or perhaps even better:

```
p + geom_hex(bins=8, alpha=.5) + geom_smooth(colour="purple4",size=2)  
## geom_smooth: method="auto" and size of largest group is >=1000, so using gam with formula: y \sim s(x, bs = "cs"). Use 'method = x' to change the smoothing method.
```



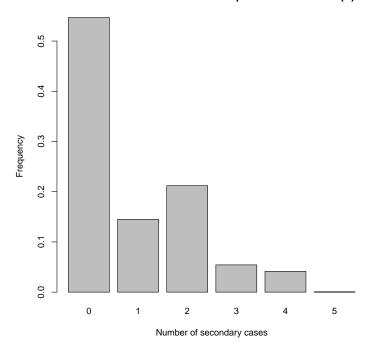
Effective reproduction numbers over time can be obtained using get.Rt, and given identical outputs to get.incid. Effective reproduction numbers per individuals can be obtained by get.R:

```
R <- get.R(res)</pre>
class(R)
## [1] "matrix"
dim(R)
## [1] 640 30
R[1:6,1:15]
     1 2 3 4 5 6 7 8 9 10 11 12 13 14 15
## 42 1 1 1 2 3 2 3 2 2 2 2
                              0
## 43 1 1 1 2 4 2 3 0 2 2
                           2
                              1
                                 4
                                     0
## 44 1 1 1 2 3 2 3 2 2 2 2
                              0
## 45 1 1 1 2 5 2 3 0 2 1
                            2
                              0
## 46 1 1 1 2 4 2 3 1 2
                         2
                            2
                               0
## 47 1 1 1 2 4 2 3 0 2 2
                           2
```

R is a matrix of effective reproduction numbers with one column per individual and one row for each retained step of the MCMC. This information can be pooled to get a rough idea of the overall distribution of R_i :

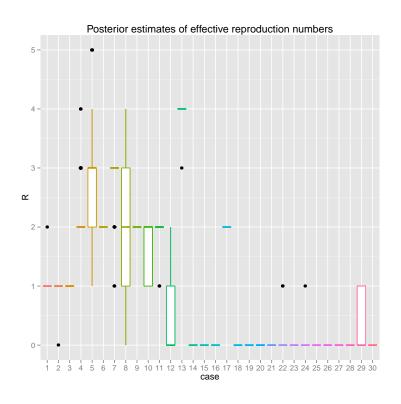
```
table(R)
## R.
       0
            1
                   2
                                     5
## 10498 2785 4070 1042
                             796
table(R)/length(R)
## R
##
           0
                     1
                               2
                                         3
## 0.5467708 0.1450521 0.2119792 0.0542708 0.0414583 0.0004687
barplot(table(R)/length(R), xlab="Number of secondary cases",
        main="Posterior estimates of effective reproduction numbers (R)", ylab="Frequency")
```

Posterior estimates of effective reproduction numbers (R)



And we can use ggplot2 to visualize the individual distributions:

```
x <- data.frame(case=factor(as.vector(col(R)), levels=as.character(1:30)),R=as.vector(R))</pre>
head(x)
##
     case R
## 1
        1 1
## 2
        1 1
## 3
        1 1
## 4
        1 1
## 5
        1 1
## 6
        1 1
tail(x)
##
         case R
## 19195
           30 0
## 19196
           30 0
           30 0
## 19197
## 19198
           30 0
           30 0
## 19199
## 19200
           30 0
p <- ggplot(data=x, aes(x=case, y=R)) + geom_boxplot(aes(colour=case))</pre>
p <- p + geom_boxplot(aes(colour=case)) + guides(colour=FALSE)</pre>
p + labs(title="Posterior estimates of effective reproduction numbers")
```



References

- [1] Jombart T, Cori A, Didelot X, Cauchemez S, Fraser C, and Ferguson N. Bayesian reconstruction of disease outbreaks by combining epidemiologic and genomic data. *PLoS Computational Biology*, accepted.
- [2] Hadley Wickham. ggplot2: elegant graphics for data analysis. Springer New York, 2009.