6. Writing Big Loops

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Writing loops in \texttt{R}

We saw that \texttt{replicate()}, and \texttt{apply()}, \texttt{sapply()} are \texttt{R}'s preferred way of looping (i.e. doing the same thing many times).

Even for expert users, their use requires some careful thought; debugging code may be complex. Also there are some jobs, such as iteration, that these loops cannot do.

In this session we’ll talk about some alternatives, and their application to high-throughput genome-wide studies.
Your first computer program?

for(i in 1:100){
    print("Hello world!")
    print(i*i)
}

- Everything inside the curly brackets {...} is done 100 times

- Looped commands can depend on i (or whatever you called the counter)

- R creates a vector i with 1:100 in it. You could use any vector that's convenient
for() loops

for() loops are very intuitive, but have some drawbacks;

- They can be slow;
  - ‘growing’ a large dataset is a bad idea;
    mydata <- cbind(mydata, rnorm(1000, mean=i))
  - so set up blank output first, then ‘fill it in’

- apply() is interpreted slightly faster than for() – but typically this will not matter, contrary to some urban myths

- for() loops require more typing than apply()! For tasks which will be repeated, writing a function really is the Right Thing to do, in the long run.

Using for(i in 1:N) sets up a vector (i) of length N. Do you really need this?
for() loops

Two alternatives; (see ?Control for details)

```r
i <- 1; my.mat <- matrix(NA, N, 3)
while(i <= N){
  z <- work.on.gene(i)
  my.mat[i,] <- summary(z)
  i <- i+1
}
```

- note that we avoided ‘growing’ the output

```r
i <- 1; my.mat <- matrix(NA, N, 3)
repeat{
  z <- work.on.gene(i)
  my.mat[i,] <- summary(z)
  i <- i+1
  if(i>=N) break
}
```

Use `apply()`, `sapply()` to avoid the ‘setup’ stage
Whole genome studies can look very intimidating...

Genome-wide association study identifies new susceptibility loci for Crohn disease and implicates autophagy in disease pathogenesis

Application to whole-genome study

... however, each \( p \)-value on that picture comes from a single logistic regression.

There may be 2,500,000 tests in total; if each one takes 1/10 sec, the analysis is done in under an hour;

<table>
<thead>
<tr>
<th>Time per test</th>
<th>Total time</th>
</tr>
</thead>
<tbody>
<tr>
<td>0.01 sec</td>
<td>7 hrs</td>
</tr>
<tr>
<td>0.1 sec</td>
<td>2 days 22 hrs</td>
</tr>
<tr>
<td>1 sec</td>
<td>28 days</td>
</tr>
<tr>
<td>5 sec</td>
<td>5 months</td>
</tr>
<tr>
<td>5 mins</td>
<td>24 yrs</td>
</tr>
</tbody>
</table>

Cutting time per test from 1 sec \( \rightarrow \) 0.1 sec is clearly worthwhile – but *proposing* analyses where each test takes \( > 5 \) secs may be silly.
Making code run faster, part 1

Some easy ‘streamlining’ ideas;

- Write a function to do just the analysis you want
  
  ```r
  > my.output <- apply(my.data, 1, my.function)
  ```

- Make a compiled version of it – use `cmpfun()` in the (standard) `compiler` package

- Pre-process/‘clean’ your data before analysis; e.g. `sum(x)/length(x)` doesn’t error-check like `mean(x)`

- Similarly, you can streamline `glm()` to just `glm.fit()` [as we’ll see, in some examples]

- Use vectorized operations, where possible

- Store data as matrices, not data.frames
Streamlining, for experts;

- Write small but *important* pieces of code in C, and call these from R

- *Batch mode* processing lets you break down e.g. the whole genome into 23 chromosomes – great if you have 23 processors to use.
  - Save your analysis in 23 output files
  - read in the answers
  - finally produce e.g. multi-color pictures

Various packages help implement this, but use is platform-specific
Timing

“Premature optimization is the root of all evil”

Donald Knuth

Do you *need* to optimize your code? Running 2 or 3 times faster may not be worth the time spent coding/debugging!

But going at least an order of magnitude faster is almost always A Good Thing.

**After** you have code that works, you may need to speed it up. Experienced users may be able to ‘eyeball’ the problem... but actual measurement is an easier *and* more reliable approach!
Timing

- `proc.time()` returns the current time. Save it before a task and subtract from the value after a task.

- `system.time()` times the evaluation of expression

- R has a *profiler*; this records which functions are being run, many times per second. `Rprof(filename)` turns on the profiler, `Rprof(NULL)` turns it off. `summaryRprof(filename)` reports how much time was spent in each function.

It’s better to improve the time-consuming functions, if possible. Remember that a 1000-fold speedup in a function used 10% of the time is less helpful than a 30% speedup in a function used 50% of the time.
High-throughput code – caveats

We saw earlier that ‘weird’ datasets can crash your code. Inevitably, these will appear in high-throughput work, and a crash at e.g. SNP #2,999,999,999 will be very frustrating.

- Some ‘weirdness’ is easy to spot;
  - Everyone is homozygous
  - All cases missing
  - No variation in outcome ...

- In more complex models, it’s easier to ‘try it and see’. Use `tryCatch()`

- When ‘weirdness’ is found, high-throughput code should;
  - Produce sensible output (\texttt{NA}, \texttt{-999} etc)
  - Handle these appropriately in summary output