

## The new `summary.scanone`

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In R/qtl version 1.04, the function `summary.scanone` has been changed quite substantially. Also, the permutations with `scanone` have changed to allow the calculation of autosome- and X-chromosome-specific LOD thresholds, and to enable stratified permutation tests.

In this document, I describe the revisions and how to use the new functions. We'll first look at the `fake.f2` data as an example.

First we need to load the package and the data.

```
> library(qtl)
> data(fake.f2)
```

I'm going to use `scanone` with `method="hk"`. First I run `calc.genoprob`, and then `scanone` as before.

```
> fake.f2 <- calc.genoprob(fake.f2, step=2.5)
> out.f2 <- scanone(fake.f2, method="hk")
```

In `summary.scanone`, we can now get an indication of the number of degrees of freedom associated with the LOD scores. We use `df=TRUE`, as follows.

```
> summary(out.f2, threshold=3, df=TRUE)
```

Degrees of freedom: A:2 X:3

	chr	pos	lod
c1.loc27.5	1	27.5	5.12
c13.loc27.5	13	27.5	8.95
cX.loc10	X	10.0	7.20

There are a couple of improvements in the permutations performed by `scanone`. First, we can calculate autosome- and X-chromosome-specific LOD thresholds; this is important in this case, as the number of degrees of freedom is different for the X chromosome. Separate autosome and X chromosome permutations may be performed in `scanone` via `perm.Xsp=TRUE`. The X-chromosome-specific thresholds requires many more permutation replicates to get a threshold of equivalent precision. An increased number of permutations is chosen automatically.

Permutations can take a very long time, and so one might want to use a multi-processor computer or cluster and do multiple shorter runs in parallel. And so we have added a function `c.scanoneperm` for combining such runs together.

```
> operm1.f2 <- scanone(fake.f2, method="hk", n.perm=500, perm.Xsp=TRUE)
> operm2.f2 <- scanone(fake.f2, method="hk", n.perm=500, perm.Xsp=TRUE)
> operm.f2 <- c(operm1.f2, operm2.f2)
```

Getting the autosome- and X-chromosome-specific thresholds is a bit tricky, and so another improvement is the addition of the function `summary.scanoneperm` for calculating such thresholds. The argument `alpha` indicates the significance levels.

```
> summary(operm.f2, alpha=c(0.05, 0.20))
```

Autosome LOD thresholds (1000 permutations)

```
lod
5% 3.23
20% 2.58
```

X chromosome LOD thresholds (37374 permutations)

```
lod
5% 4.33
20% 3.42
```

Further, we may include the permutation results in the call to `summary.scanone` to automatically calculate thresholds and to have genome-scan-adjusted p-values displayed.

```
> summary(out.f2, perms=operm.f2, alpha=0.05, pvalues=TRUE)
```

```
chr pos lod pval
1 1 27.5 5.12 0.00103
2 13 27.5 8.95 0.00000
3 X 10.0 7.20 0.00000
```

Finally, one may prefer to do a stratified permutation test, permuting the phenotypes separately within each of the groups defined by sex and cross direction. This may be done in `scanone` with the argument `perm.strata`, which should be a numeric vector whose unique values define the separate strata.

We set of the strata as follows.

```
> sex <- fake.f2$pheno$sex
> pgm <- fake.f2$pheno$pgm
> strata <- sex + 2*pgm
> table(strata)
```

```
strata
  0  1  2  3
52 48 50 50
```

We then perform the permutation test in four pieces, and combine the results together, as follows.

```
> operm1.f2strat <- scanone(fake.f2, method="hk", n.perm=250,
+                           perm.Xsp=TRUE, perm.strata=strata)
> operm2.f2strat <- scanone(fake.f2, method="hk", n.perm=250,
+                           perm.Xsp=TRUE, perm.strata=strata)
> operm3.f2strat <- scanone(fake.f2, method="hk", n.perm=250,
+                           perm.Xsp=TRUE, perm.strata=strata)
> operm4.f2strat <- scanone(fake.f2, method="hk", n.perm=250,
+                           perm.Xsp=TRUE, perm.strata=strata)
> operm.f2strat <- c(operm1.f2strat, operm2.f2strat, operm3.f2strat,
+                   operm4.f2strat)
```

The new thresholds are as follows.

```
> summary(operm.f2strat, alpha=c(0.05, 0.20))
```

Autosome LOD thresholds (1000 permutations)

```
lod
5%  3.42
20% 2.69
```

X chromosome LOD thresholds (37376 permutations)

```
lod
5%  4.24
20% 3.40
```

The big changes to the `summary.scanone` function concern the case of results for multiple phenotypes. To illustrate this, we will look at the `fake.bc` data, which has two phenotypes. First we load the data.

```
> data(fake.bc)
```

Now let's run `calc.genoprob` and do a genome scan on the two phenotypes. Again, we use `method="hk"` for the sake of speed.

```
> fake.bc <- calc.genoprob(fake.bc, step=2.5)
> out.bc <- scanone(fake.bc, pheno.col=1:2, method="hk")
```

The results contain LOD scores for each of the phenotypes. By default, `summary.scanone` looks at the first of these, though it also shows the LOD score for the second phenotype at the locations of the LOD peaks for the first phenotype.

```
> summary(out.bc, threshold=3)
```

	chr	pos	pheno1	pheno2
c2.loc32.5	2	32.5	3.52	1.87
c5.loc17.5	5	17.5	7.90	2.71

If we use `lodcolumn=2`, we get the analogous results, looking at the second phenotype.

```
> summary(out.bc, threshold=3, lodcolumn=2)
```

	chr	pos	pheno1	pheno2
D5M394	5	9.8	7.42	3.9

If we use `format="allpheno"`, we get separate rows for the peaks of each of the phenotypes.

```
> summary(out.bc, threshold=3, format="allpheno")
```

	chr	pos	pheno1	pheno2
c2.loc32.5	2	32.5	3.52	1.87
D5M394	5	9.8	7.42	3.90
c5.loc17.5	5	17.5	7.90	2.71

Perhaps the most convenient output is obtained with `format="allpeaks"`, which gives a single row for each chromosome, with the maximum LOD score and its position for each of the phenotypes. A chromosome is displayed if the LOD score for at least one of the phenotypes exceeds its threshold. The `threshold` argument can be a single threshold, applied to all phenotypes, or we can give a vector with separate thresholds for each of the LOD score columns.

```
> summary(out.bc, threshold=c(3,2.5), format="allpeaks")
```

	chr	pos	pheno1	pos	pheno2
1	2	32.5	3.52	37.5	1.91
2	5	17.5	7.90	9.8	3.90

A permutation test may be performed as before. Since the `fake.bc` data has only autosomal data, use of `perm.Xsp=TRUE` would be ignored.

```
> operm.bc <- scanone(out.bc, pheno.col=1:2, method="hk", n.perm=1000)
```

We can again use `summary` to get LOD thresholds

```
> summary(operm.bc, alpha=0.05)
```

LOD thresholds (1000 permutations)

	pheno1	pheno2
5%	2.57	2.55

And again these can be used in `summary.scanone` to calculate thresholds and get genome-scan-adjusted p-values.

```
> summary(out.bc, perms=operm.bc, alpha=0.05, format="allpeaks",  
+         pvalues=TRUE)
```

	chr	pos	pheno1	pval	pos	pheno2	pval
1	2	32.5	3.52	0.005	37.5	1.91	0.195
2	5	17.5	7.90	0.000	9.8	3.90	0.002