Biostat 140.668 Problem set 5

## QTL mapping: other single-QTL models

1. Consider mapping a simple Mendelian disease gene using a backcross. Suppose you have essentially complete genotype data, so that the exact locations of crossovers can be determined. Let *L* denote the length (in cM) of the interval to which you will have mapped the gene with *n* backcross individuals (i.e., the distance between the two closest flanking recombination events).

- (a) How many backcross individuals will you need so that  $E(L) \le 1$  cM?
- (b) How many backcross individuals will you need so that  $Pr(L \le 1 \text{ cM}) \ge 80\%$ ?
- 2. Derive the extension of the Kruskal-Wallis statistic discussed in lecture, for nonparametric interval mapping.

Let  $R_i$  denote the ranks of the phenotypes, and let  $p_{ig}$  denote the probability that mouse i has QTL genotype g, given the available marker data. Define  $\bar{S}_g = \sum_i R_i p_{ig} / \sum_i p_{ig}$ 

- (a) Show that  $E(\bar{S}_g \mid \text{marker data, no linkage}) = (n+1)/2$ .
- (b) Show that

$$\operatorname{var}(\bar{S}_g \mid \operatorname{marker data}, \operatorname{no linkage}) = \left[\frac{n+1}{12}\right] \left[\frac{n\sum_i p_{ig}^2 - (\sum_i p_{ig})^2}{(\sum_i p_{ig})^2}\right]$$

(c) Put it all together.