

### Homework #2 Answer Key

1. Lumped heterozygosity =  $2\bar{p}\bar{q} - 2\text{var}(p)$  where  $\bar{p} = (.2 + .4)/2 = .3$ ,  $\bar{q} = 1 - .3 = .7$ , and  $\text{var}(p) = (.2^2 + .4^2)/2 - (.3)^2 = .01 \Rightarrow$  lumped heterozygosity =  $2(.3)(.7) - 2(.01) = .4$ . The expected heterozygosity if one random mating population =  $2\bar{p}\bar{q} = 2(.3)(.7) = .42$ .  $\Rightarrow$  observed heterozygosity < expected heterozygosity.

$\bar{H}_i(\text{observed}) = 2\bar{p}_i\bar{p}_j + 2\text{cov}(p_i, p_j) \Rightarrow \bar{H}_{12}(\text{observed}) = 2\bar{p}_1\bar{p}_2 + 2\text{cov}(p_1, p_2)$ .  
 $\bar{p}_1 = (.2 + .6)/2 = .4$ ,  $\bar{p}_2 = (.4 + .2)/2 = .3$ ,  
 $\text{cov}(p_1, p_2) = [(.2)(.4) + (.6)(.2)]/2 - (.4)(.3) = -.02 \Rightarrow$   
 $\bar{H}_{12}(\text{observed}) = 2(.4)(.3) + 2(-.02) = .2$ , which is less than  $\bar{H}_{12}(\text{expected}) = 2(.4)(.3) = .24$   
 $\bar{H}_{13}(\text{observed}) = 2\bar{p}_1\bar{p}_3 + 2\text{cov}(p_1, p_3)$ ,  $\bar{p}_3 = 1 - \bar{p}_1 - \bar{p}_2 = .3$ ,  
 $\text{cov}(p_1, p_3) = [(.2)(.4) + (.6)(.2)]/2 - (.4)(.3) = -.02 \Rightarrow \bar{H}_{13}(\text{observed}) = .2 <$   
 $\bar{H}_{13}(\text{expected}) = .24$ .  $\bar{H}_{23}(\text{observed}) = 2\bar{p}_2\bar{p}_3 + 2\text{cov}(p_2, p_3)$ ,  
 $\text{cov}(p_2, p_3) = [(.4)(.4) + (.2)(.2)]/2 - (.3)(.3) = .01 \Rightarrow$   
 $\bar{H}_{23}(\text{observed}) = 2(.3)(.3) + 2(.01) = .2 > \bar{H}_{23}(\text{expected}) = 2(.3)(.3) = .18$ .  
 Overall observed heterozygosity =  $.2 + .2 + .2 = .6 <$  overall expected heterozygosity =  $1 - (.4)^2 - (.3)^2 - (.3)^2 = .66$ .

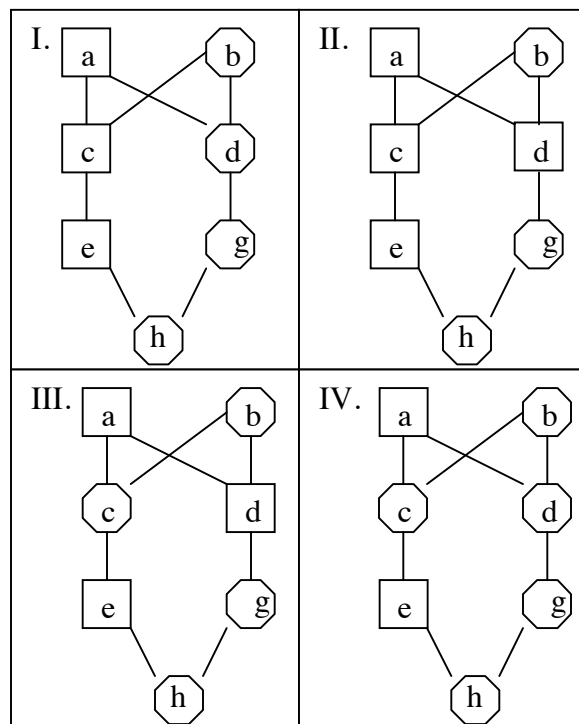
2. In general, the ratio of freq(recessive homozygote with inbreeding f) to freq(recessive homozygote with inbreeding f = 0) given a recessive allele of frequency q is

$$\frac{q^2 + fpq}{q^2} = 1 + f \frac{p}{q}$$

If  $f = .005$  and  $q = .005$ , then  $p = 1 - .005 = .995$  and the ratio is  $1 + (.005)(.995)/(.005) = 1.995$ .

Even a relatively small amount of inbreeding can significantly amplify the frequency of a rare recessive condition relative to a randomly mating population. In this case, an inbreeding coefficient of just 0.5% would almost double the frequency of the recessive disease.

3. The four possible first-cousin pedigrees are shown at right. Note that pedigrees I and II contain successive males in all lines leading to offspring h and so have inbreeding coefficients  $f = 0$ . In pedigree III, the only chain that contributes to inbreeding is “e-c-b-d-g” since the chain through “a” contains successive males. The inbreeding coefficient for this case is  $f = 1 \cdot (1/2) \cdot (1/2) \cdot 1 \cdot (1/2) = 1/8$ . In case IV, there are two feasible chains: “e-c-b-d-g” and “e-c-a-d-g”. (Note that neither contains successive males.) Inbreeding



via the first chain is  $1 \cdot (1/2) \cdot (1/2) \cdot (1/2) \cdot (1/2) = 1/16$  and  $1 \cdot (1/2) \cdot 1 \cdot (1/2) \cdot (1/2) = 1/8$  via the second chain. Summing these gives  $f = 3/16$ .

## 4. Case 1:

$$H_I = (.5 + .3)/2 = .4$$

$$\bar{H}_S: H_{S,1} = 2p_1q_1 = 2(.5)(.5) = .5; H_{S,2} = 2p_2q_2 = 2(.5)(.5) = .5 \Rightarrow \bar{H}_S = (.5 + .5)/2 = .5$$

$$H_T: q_1 = q_2 = 0.5 \Rightarrow \bar{q} = (.5 + .5)/2 = .5 \Rightarrow H_T = 2\bar{p}\bar{q} = 2(.5)(.5) = .5.$$

$$\text{So } F_{IS} = \frac{\bar{H}_S - H_I}{\bar{H}_S} = (.5 - .4)/.5 = .2; F_{ST} = \frac{H_T - \bar{H}_S}{H_T} = (.5 - .5)/.5 = 0;$$

$$F_{IT} = \frac{H_T - H_I}{H_T} = (.5 - .4)/.5 = .2.$$

## Case 2:

$$H_I = (.5 + .42)/2 = .46$$

$$\bar{H}_S: H_{S,1} = 2p_1q_1 = 2(.5)(.5) = .5; H_{S,2} = 2p_2q_2 = 2(.7)(.3) = .42 \Rightarrow \bar{H}_S = (.5 + .42)/2 = .46$$

$$H_T: q_1 = 0.5, q_2 = 0.3 \Rightarrow \bar{q} = (.5 + .3)/2 = .4 \Rightarrow H_T = 2\bar{p}\bar{q} = 2(.6)(.4) = .48$$

$$\text{So } F_{IS} = \frac{\bar{H}_S - H_I}{\bar{H}_S} = (.46 - .46)/.46 = 0; F_{ST} = \frac{H_T - \bar{H}_S}{H_T} = (.48 - .46)/.48 = .0417;$$

$$F_{IT} = \frac{H_T - H_I}{H_T} = (.48 - .46)/.48 = .0417$$

5. Subpopulation 1:  $p_1 = p_2 = \dots = p_5 = 1/5$ ,  $p_6 = p_7 = \dots = p_{10} = 0$ 

Subpopulation 2:  $p_1 = p_2 = \dots = p_5 = 0$ ,  $p_6 = p_7 = \dots = p_{10} = 1/5$

Averages:  $\bar{p}_1 = \bar{p}_2 = \dots = \bar{p}_{10} = 1/10$

$$H_{S,1} = H_{S,2} = 1 - \sum_{i=1}^{10} p_i^2 = 1 - 5(1/5)^2 = 4/5 \Rightarrow \bar{H}_S = 4/5.$$

$$H_T = 1 - \sum_{i=1}^{10} \bar{p}_i^2 = 1 - 10(1/10)^2 = 9/10 \Rightarrow G_{ST} = (.9 - .8)/.9 = .111$$

On the surface, this value of  $G_{ST}$  is not particularly large despite the two subpopulations not sharing even a single allele! The morale of the exercise is that any particular value of  $F_{ST}$  ( $G_{ST}$ ) has limitations as an indicator of population substructuring. One must also consider the number and distribution of alleles on which a given fixation index computation is based.

$$6. N_e(\text{diploid}) = \frac{4N_f N_m}{N_f + N_m} = \frac{4(5)(1)}{5 + 1} = 3\frac{1}{3}$$

$$N_e(\text{haplo-diploid}) = \frac{9N_f N_m}{2N_f + 4N_m} = \frac{9(1)(10)}{2(1) + 4(10)} = 2.14$$

7. Exact:  $N_e = \frac{1}{2} \left\{ 1 - \left[ (1 - 1/10)(1 - 1/100)(1 - 1/20)(1 - 1/200) \right]^{1/4} \right\}^{-1} = 11.9;$

Approximate:  $N_e \approx \frac{4}{1/5 + 1/50 + 1/10 + 1/100} = 12.1$ . The exact and approximate values of  $N_e$  are obviously very close to one another!

8.  $N_e(\text{mtDNA}) = \frac{N_{ef}}{2} = \frac{100}{2} = 50;$

$$N_e(\text{Y chrom}) = \frac{N_m}{2} = \frac{10}{2} = 5;$$

$$N_e(\text{autosomal}) = \frac{4N_{ef}N_{em}}{N_{ef} + N_{em}} = \frac{4(100)(10)}{100 + 10} = 36.4$$