Questions on Delaneau et al (2013) Nature Methods 29:84-91.

Improved whole-chromosome phasing for disease and population genetic studies.

Make sure you read through section 9 of the supplementary methods.

- 1. In the background section, the authors summarize some previous strategies for generating haplotype estimation algorithms that do not scale quadratically with sample size. Summarize the algorithms implemented in MACH and IMPUTE2.
- 2. The SHAPEIT algorithm also avoids scaling quadratically with sample size. How?
- 3. The present paper tries to improve upon the original SHAPEIT algorithm by including some features of the IMPUTE2 approach. What prompted the authors to consider this?
- 4. The algorithm divides chromosomes into small segments, each with a number of heterozygous sites. What are the advantages of small segments? What are the advantages of long segments?
- 5. How do the segment boundaries change as the algorithm proceeds?
- 6. Do you think the proposed algorithm will be sensitive to missing data? If so, why?
- 7. Do you think the proposed algorithm could be adapted to low pass sequencing data? If so, why? If not, why not?
- 8. What is the danger of including closely related individuals in the sample? How can this be avoided?
- 9. What struck you most about the paper?