Commentary on: Budowle B, Shea B, Niezgoda S, Chakraborty R. CODIS STR loci data from 41 sample populations. J Forensic Sci 2001;46:453–489.

Sir,

Analysis of CODIS data from a large study of human populations by Budowle et al. (1) raised concerns articulated in a Commentary by Krane et al. (2) and rebutted in the Authors' Response by Chakraborty et al. (3). One aspect of this exchange merits additional discussion.

Krane et al. (2) objected to the use of a Bonferroni correction when a test of Hardy-Weinberg equilibrium (HWE) was performed on this dataset. While the original study convincingly demonstrated differentiation among population groups and the absence of structure within individual populations, the use of a Bonferroni correction, as defended in the Response, may be inappropriate. This adjustment has been repeatedly critiqued in recent ecological and evolutionary literature (4-7). Kinnison et al. (8) suggested that if one is interested in testing the combined multilocus evidence of HWE over the entire dataset, as was done in Budowle et al. (1), the binomial likelihood function is more appropriate. As the number of loci and/or populations increases, critical values can become exceptionally small making detection of truly significant results improbable, i.e., inflating type I errors and generating false negatives. This may not be acceptable if the researchers wish to identify multilocus evidence of populations that exhibit structure or multipopulation evidence of problematic loci. For example, consider a sample that deviated significantly (p = 0.010) from HWE at each of 12 loci examined. In a general test of HWE using a Bonferroni corrected critical value, such as performed by Budowle et al. (1), these deviations would be deemed inconsequential if this sample were examined simultaneously with 40 additional populations. This same outcome results for testing HWE after correction in the hypothetical population. However, the systematic deviation would be obvious by visual inspection of the uncorrected probabilities, a "vote-counting" approach. As noted by Chakraborty et al. (3), Bonferroni-type adjustments are warranted if researchers wish to minimize detection of false positives in genomic scans. More generally, controlling false discovery rate (9) has been advocated as a powerful approach for a variety of studies in this arena (10,11).

References

- 1. Budowle B, Shea B, Niezgoda S, Chakraborty R. CODIS STR loci data from 41 sample populations, J Forensic Sci 2001;46:453–89. [PubMed]
- Krane DE, Doom TE, Mueller L, Raymer ML, Shields WM, Thompson WC. Commentary on Budowle B, Shea B, Niezgoda S, Chakraborty R. CODIS STR data from 41 sample populations. J Forensic Sci 2004;49: 1388–9.
- Chakraborty R, Lee HS, Budowle B. Authors' Response. J Forensic Sci 2004;49:1389–93.
- Moran MD. Arguments for rejecting the sequential Bonferroni in ecological studies. Oikos 2003;100:403–5.
- Garcia LV. Controlling the false discovery rate in ecological research. Trends Ecol Evol 2003;18:553–54.
- 6. Garcia LV. Escaping the Bonferroni iron claw in ecological studies. Oikos 2004;105:657–63.
- Nakagawa S. A farewell to Bonferroni: the problems of low statistical power and population bias. Behav Ecol 2004;15:1044–5.
- Kinnison MT, Bentzen P, Unwin MJ, Quinn TP. Reconstructing recent divergence: evaluating non equilibrium population structure in New Zealand Chinook salmon. Mol Ecol 2002;11:739–54. [PubMed]
- Benjamini Y, Hochberg Y. Controlling the false discovery rate: a practical and powerful approach to multiple testing. J Roy Stat Soc 1995; B57:289–300.
- Storey JD, Tibshirani IR. Statistical significance for genome-wide studies. Proc Natl Acad Sci USA 2003;100:9440–5. [PubMed]
- Manley KF, Nettleton D, Hwang JTG. Genomics, prior probability, and statistical tests of multiple hypotheses. Genome Res 2004;14:997–1001. [PubMed]

Irving L. Kornfield, Ph.D. Molecular Forensics Laboratory University of Maine Orono, ME 04469