

$$S=10$$

$$R=100$$

$$I=300$$

Draft March 6, 2006

Statistical Analysis:

We tested the null hypothesis that readers' average accuracy with modality 1 is equivalent to readers' average accuracy with modality 2. The alternative hypothesis was that the average accuracies are not equivalent. We applied the multi-reader ROC analysis method of Obuchowski and Rockette [ref] which treats the study readers as randomly-selected readers from the population of readers, and the study patients as randomly-selected patients from the population of patients (i.e. random-reader effects model). A significance level of 0.05 was used. The details of the analysis follow.

Let R denote the total number of readers and let I denote the total number of patients in the study. Let R_s denote the number of readers in subgroup s , and I_s denote the total number of patients in subgroup s , such that $\sum_s R_s = R$ and $\sum_s I_s = I$. Let r_{sj} denote the j -th reader in subgroup s .

For each reader we constructed a ROC curve for each modality and estimated the area under the ROC curve and its variance using standard parametric methods [ref to Metz software]. We let $A_{s(j)}^m$ denote the estimated area under the ROC curve for reader j in subgroup s with modality m , where $m=1,2$. Let $V_{s(j)}^m$ denote the estimated variance of the area under the ROC curve for reader j in subgroup s . The covariance between the two estimated ROC areas of reader j in subgroup s is denoted by $V_{s(j)}^{1,2}$. Similarly, the covariance between the estimated ROC areas for modality m of readers j and k in subgroup s is denoted by $V_{s(i,k)}^{m,m}$. The covariance between the estimated ROC areas for reader j in modality 1 and reader k in modality 2, where both readers j and k are in subgroup s , is denoted by $V_{s(j,k)}^{1,2}$. Consider reader j in subgroup s and reader k in subgroup t , where $s \neq t$. Note that $V_{s(i),t(k)}^{m,m}=0$ and $V_{s(j),t(k)}^{1,2}=0$.

Define the symmetric $R \times R$ variance-covariance matrix B^m . The diagonal consists of the variances of the R readers in modality m , i.e. $V_{s(j)}^m$. The off-diagonal elements are the covariances between the R readers in modality m , i.e. $V_{s(i,k)}^{m,m}$ and $V_{s(i),t(k)}^{m,m}$.

Define the symmetric $R \times R$ covariance matrix C . The diagonal consists of the covariances of the R readers in modalities 1 and 2, i.e. $V_{s(j)}^{1,2}$. The off-diagonal elements are the covariances between the R readers in modality 1 and 2, i.e. $V_{s(i,k)}^{1,2}$ and $V_{s(i),t(k)}^{1,2}$.

The covariances in matrices B^1 , B^2 , and C were estimated using ROCKIT [ref to Metz software] by performing the appropriate pairwise comparisons of ROC areas. For example, to estimate $V_{s(j,k)}^{1,2}$ we ran ROCKIT to compare the ROC areas of reader j in modality 1 to reader k in modality 2. The covariance between these two ROC areas is estimated by ROCKIT and used here to construct matrix C .

OBUMRM2 [ref to website] is a FORTRAN program that performs the multi-reader ROC analysis method of Obuchowski and Rockette [ref]. Using the second input option in OBUMRM2, we input the $R \times 2$ estimated ROC areas, along with matrices B^1 , B^2 , and C . OBUMRM2 performed the analysis, and we present the results.