

## Standardization of Assays from Multiple Labs for a Given Analyte

Assume one lab/method ('lab 0') has a special status, by virtue of presumed assay quality or frequency of use in WHI samples or some other criterion, and consider the standardization of measures from other labs/methods so that they 'align' with lab 0 measures:

For a specific analyte (e.g. CRP; vitamin D) let  $Y(i,j)$  denote the log-transformed assessment for the  $j$ th individual from lab/method  $i$ ;  $i=0,1,2,\dots$

Apply a simple statistical model

$$Y(i,j) = a_0 + a_i X(i) + b \cdot Z(i,j) + s_i e(i,j),$$

Where  $X(i)=1$  for lab  $i>0$ , and  $=0$  otherwise,

$Z(i,j)$  is a vector of study subject characteristics that may associate with the  $Y$ , for the  $j$ th participant and the  $i$ th lab/method (e.g. case versus control status in contributing project, risk factors for elevated values of analyte in question),

$s_i$  is a scale parameter for the  $i$ th lab, and

$e(i,j)$  is a noise term.

After **model fitting** the standardized values would be:

$Y(i,j)$  for lab 0,

$Y(i,j) + a_i \hat{a}$ , for lab  $i$  measurements, where 'hat' denotes estimate, in the absence of evidence against  $s(i)=s(0)$ , and either

$(Y(i,j)+a_i \hat{a}) / (s_i \hat{a} / s_0 \hat{a})$  for lab/method  $i$  measurements if there is evidence against  $s(i)=s(0)$ , or decide to keep lab/method  $i$  data separate (no standardized value developed).

Label standardized data a 'standardized to lab 0 assessments'.

Some additional useful analyses can be entertained if multiple labs/methods were used on the same specimens. For example, differences between  $Y(i,j)$  values from the same specimen (ratios on the original scale) are free of dependence on  $Z(i,j)$  values under the above model, and if sufficiently numerous, can support simple estimates of  $a_i$  and  $s_i$  values.

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