

CALCULATION OF THE STANDARD DEVIATION OF THE STANDARDISED MEAN IN KIWIQC

If we had independent (non-correlated) random standardised variates, then the standard deviation of the standardised mean (\bar{z}_m) would be given by

$$\sigma_{\bar{z}} = \frac{1}{\sqrt{n}} \quad [1]$$

where n is the number of levels of QC. This assumes that the means and SDs of the individual QC levels are calculated from a large number of batches (>100).

Example. For three QC levels ($n = 3$) the “6 sigma” control limits are

$$\pm 3\sigma_{\bar{z}} = \frac{3}{\sqrt{3}} = \pm 1.732$$

However, this calculation is not appropriate for multiple QC levels, as they are usually correlated.

For non-independent (correlated) random standardised variates:

$$\sigma_{\bar{z}} \approx \frac{(1 + (n - 1)\bar{r})^{0.5}}{\sqrt{n}} \quad [2]$$

where $\bar{r} = (r_{12} + r_{13} + r_{23})/3$ and r is the correlation coefficient. This is the expression for the standard deviation of the standardised mean (z_m) used in KiwiQC

Example: for three QC levels (ie $n = 3$) where say $\bar{r} = 0.5$, the “6 sigma” control limits are

$$\pm 3\sigma_{\bar{z}} \approx \frac{3\sqrt{(1 + (3 - 1) \times 0.5)}}{\sqrt{3}} = \pm 2.449$$

Derivation of equation 2.

Variance of the mean ($\sigma_{\bar{x}}^2$) = expectation of the mean squared ($E(\bar{x})^2$).

$$\begin{aligned} E(\bar{x})^2 &= E(x_1 + x_2 + \dots + x_n)(x_1 + x_2 + \dots + x_n)/n^2 \\ &= E(x_1^2 + x_2^2 + \dots + x_n^2 + 2x_1x_2 + 2x_1x_3 + \dots + 2x_{n-1}x_n)/n^2 \end{aligned}$$

Now $E(x_i) = \sigma_i^2$ and $E(x_i x_j) = \sigma_{ij}$ where σ_i^2 is the variance of x_i and σ_{ij} is the covariance of x_i and x_j .

In KiwiQC, the QC values are standardised (mean = 0, sd = 1) so that all $\sigma_i^2 = 1$, and $\sigma_{ij} = r_{ij}$ where r is the correlation coefficient. If it is assumed that all the r_{ij} are equal, and using z to indicate standardised values, we have

$$E(\bar{z})^2 = \sigma_{\bar{z}}^2 = (n + 2r_{ij}n(n-1)/2)/n^2$$

Simplifying gives

$$\sigma_{\bar{z}}^2 = (1 + (n-1)r_{ij})/n$$

In fact the r_{ij} are not all equal in the QC situation, so as an approximation we replace r_{ij} in the formula by the \bar{r} , the mean of the $n(n-1)/2$ pair-wise correlations between the n levels of QC.

This gives

$$\sigma_{\bar{z}} \approx (1 + (n-1)\bar{r})^{0.5} / \sqrt{n}$$

Hence approximate $\pm 3sd$ control limits are given by

$$\pm 3(1 + (n-1)\bar{r})^{0.5} / \sqrt{n}$$

KiwiQC web page

<http://www.cdhb.govt.nz/chlabs/endo/KiwiQcWebPage.htm>

