

# Supplement to “Tail approximations for the Student $t$ –, $F$ –, and Welch statistics for non-normal and not necessarily i.i.d. random variables”

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This Appendix contains remarks on Theorem 1.1; an example of an application of Theorem 1.1 to real data; extended version of the literature review; comparison of the result of Theorem 3.1 with the exact distribution of the Welch statistic; proof of Theorem 5.1.

## 1.1 Remarks on Theorem 1.1

For the ease of presentation we repeat the statement of Theorem 1.1 below

**Theorem 1.1.** *For any fixed value of  $n$  and each of the three choices of  $T$ , there exists a functional  $K : \mathcal{G} \rightarrow \mathbb{R}^+$ , such that for all  $g_0, g_1 \in \mathcal{G}$  the limit expression*

$$\frac{P(T > u|H_1)}{P(T > u|H_0)} = \frac{K_{g_1}}{K_{g_0}} + o(1) \quad \text{as } u \rightarrow \infty \quad (1)$$

*holds with constants  $0 < K_{g_0} = K(g_0) < \infty$  and  $0 < K_{g_1} = K(g_1) < \infty$ . The exact expressions for  $K(g)$  are given in Sections 2, 3, and 5 for the three choices of the test statistic  $T$ .*

**Remark 1.** Standard assumption in the use of any of the test statistics described above is that  $\mathbf{X} \sim MVN(\mathbf{0}, \sigma^2 \mathbf{1}_n)$ , where  $MVN(\boldsymbol{\mu}, \boldsymbol{\Sigma})$  denote the multivariate normal distribution with mean vector  $\boldsymbol{\mu}$  and covariance matrix  $\boldsymbol{\Sigma}$ . It is easy to check that  $MVN(\mathbf{0}, \sigma^2 \mathbf{1}_n) \in \mathcal{G}$  and that  $K(MVN(\mathbf{0}, \sigma^2 \mathbf{1}_n)) = 1$ .

**Remark 2:** Theorem 1.1 holds also for the Welch test. In this case  $K(MVN(\mathbf{0}, \sigma^2 \mathbf{1}_n))$  is not necessarily equal to 1, see Table 2 in Section 4.

**Remark 3:** It might be tempting to think that if we start with  $g_1(\mathbf{x}) = g_0(\mathbf{x}) \in \mathcal{G}$ , then small perturbation of  $g_1(\mathbf{x})$  will induce small change in the right-hand side of (1), i.e. that it will deviate “just slightly” from 1. This is, generally, not the case. For example, it can be shown that if  $T$  is a  $Z$ –statistic,  $g_0 \sim MVN(\mathbf{0}, \mathbf{1}_n)$ , and  $g_1 \sim MVN(\mu \times \mathbf{1}, \mathbf{1}_n)$ , then the right-hand side of (1) becomes either 0, 1, or  $\infty$  for  $\mu < 0$ ,  $\mu = 0$ , and  $\mu > 0$  respectively. Theorem 1.1 therefore does not hold for  $Z$ –statistic.

**Remark 4:** It is important to emphasize that large sample approximations, that is asymptotic behavior for the case when  $u$  is fixed and  $n \rightarrow \infty$ , or when both  $n \rightarrow \infty$  and  $u_n \rightarrow \infty$ , are irrelevant to the topic of the present article. We always assume that  $u \rightarrow \infty$  and  $n$  is fixed, and preferably small. In particular, the convergence speed in (1) is faster for smaller sample sizes.

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## 1.2 An example of an application of Theorem 1.1

Let  $g_1(\mathbf{x})$  be the joint density of the generic pair  $\mathbf{X}$  described in the HTS study in Section 1.2, and let  $F_1(x)$  be the CDF of the  $p$ -value  $p = t(T(\mathbf{X}))$  that corresponds to the right-sided  $t$ -test. Assume also that  $g_1(\mathbf{x})$  is continuous and satisfies regularity constraints of Theorem 2.1 (which is most likely the case except for really “pathological” instances). Then, according to Theorem 1.1 with  $g_0 \sim MVN(\mathbf{0}, \sigma^2 \mathbf{1}_2)$  and Remark 1 it follows that  $F_1(0) = 0$  and the right-sided derivative of  $F_1(x)$  at zero is equal to

$$\begin{aligned} \lim_{x \rightarrow 0+} \frac{F_1(x) - F_1(0)}{x} &= \lim_{u \rightarrow \infty} \frac{F_1(t(u))}{t(u)} = \lim_{u \rightarrow \infty} \frac{\mathbf{P}(t(T(\mathbf{X})) \leq t(u))}{t(u)} \\ &= \lim_{u \rightarrow \infty} \frac{\mathbf{P}(T(\mathbf{X}) \geq u)}{t(u)} = \lim_{u \rightarrow \infty} \frac{\mathbf{P}(T(\mathbf{X}) > u | \mathbf{X} \sim g_1)}{\mathbf{P}(T(\mathbf{X}) > u | \mathbf{X} \sim g_0)} = K_{g_1}. \end{aligned}$$

Here we use continuity of  $g_1(\mathbf{x})$  to replace “greater or equal” by “strictly greater” in the probability in the numerator of the last fraction in the above equation. Finally, we conclude that  $K_{g_1}$  is the slope of the line tangent to the graph of  $F_1(x)$  at zero, that is,

$$K_{g_1} = F_1'(0+).$$

Using this simple linear approximation of the left tail of the distribution of the  $p$ -values, as shown in Figure 1, one can, for example, correct theoretical  $p$ -values for non-normal distribution of the LSC. Further applications include quantifying the impact of non-normality, dependence, or non-homogeneity of data from HTS experiments on the number of false positives; a detailed description of the methods lies beyond the scope of the present article and a curious reader is referred to the Introduction Part of Zholud (2011) and Rootzén and Zholud (2014).

## 1.3 Extended version of the literature review

There is an extensive literature on the behavior of the Student  $t$ - and  $F$ - statistic under deviation from normality. Below we focus mostly on the Student one-sample  $t$ -statistic, with references to the Student two-sample  $t$ -,  $F$ -, and Welch statistics given as needed.

A brief and quite recent introduction to the Student one-sample  $t$ -test can be found in ZABELL (2008), and Cressie (1980) is a review with emphasis on understanding of the behavior of the test statistic under non-normality.

A main theme in the literature is the *Normal Approximation*, which is commonly stated as follows: “if the sample size is large enough and the population distribution is in the domain of attraction of the normal law, then the Student one-sample  $t$ -statistic is approximately  $N(0, 1)$  distributed”, see e.g. Giné et al. (1997). The non-central  $t$ -statistic is discussed in Bentkus et al. (2007). However, the Normal Approximation is inaccurate for small sample sizes, which are the center of interest in this paper.

Additional accuracy in the Normal approximation can be obtained by using the first few terms of a Gram-Charlier series, Geary (1936), Bartlett (1935), or *Edgeworth expansion*, see e.g. Field and Ronchetti (1990), Hall (1987) and Gaen (1949, 1950). The Edgeworth expansion improves the Normal approximation and performs better for smaller sample sizes, but still is inaccurate in the extreme tail area.

A different approach is to use *Saddlepoint approximations* to the distribution of the test statistics, see e.g. Zhou and Jing (2006), Jing et al. (2004) and Daniels and Young (1991). The simulation study of Jing et al. (2004) showed that the Saddlepoint approximation for Student’s  $t$ -statistic is very accurate in the tail area, however it is not known whether the relative error of such approximation tends to zero as one goes far out in the tail. The formulas for the Saddlepoint approximation do not

give explicit formulas for the tail behavior of the test statistic and are computationally complex. In particular, Saddlepoint approximations require knowledge of the population density, which limits their use in statistical inference. Details on Saddlepoint approximations can be found in Kolassa (2006), Jensen (1995), Reid (1988) and Lugannani and Rice (1980).

The Student one-sample  $t$ -test is closely related to the so-called self-normalized sum, see e.g. Shao (2004), Shao (1997) and Logan et al. (1973). The exact distribution of  $t$ -statistic for some special cases is discussed in Eden and Yates (1933), Rider (1929), Perlo (1933) and Laderman (1939).

Fundamental results on the Student two-sample  $t$ -test and  $F$ -test were formulated by Fisher (1924, 1925, 1935a,b) and the behavior of these tests under deviation from standard assumptions was thoroughly studied. One particular case is known as the two-means Behrens-Fisher problem, see e.g. Sawilowsky (2002) and Kim and Cohen (1998) and the list of references in these papers, and addresses the question of using the Student two-sample  $t$ -test when the variances of the two populations are unequal. A common approach to the Behrens-Fisher problem is to use the Welch-Satterthwaite approximation, see Aspin and Welch (1949), Welch (1937, 1947), and Satterthwaite (1941, 1946). Exact distribution for the Welch  $t$ -test for odd sample sizes is given in Ray and Pitman (1961). The formulas in the latter paper can be easily modified to hold for the Student two-sample  $t$ -test as well. The effect of non-normality on the  $F$ -test was considered by e.g. Box (1953, 1954), Gaen (1950) and David and Johnson (1951).

## 1.4 Welch statistic

In this section we consider the tails of the Welch statistic  $T$  and discuss the accuracy of our asymptotic approximations for  $\mathbf{P}(T > u)$  as  $u \rightarrow \infty$ . For the rest of the section we assume that the data is a Gaussian zero-mean vector, and the components are independent, and identically distributed within the two samples  $(X_1, X_2, \dots, X_{n_1})$  and  $(X_{n_1+1}, X_{n_2+2}, \dots, X_n)$  having population variances  $\sigma_1^2$  and  $\sigma_2^2$  respectively.

Under the above assumptions Ray and Pitman (1961) showed that for odd sample sizes  $n_1$  and  $n_2$  the exact distribution of the Welch statistic  $T$  (also known as Fisher-Behrens-Welch or Welch-Aspin statistic) is given by the weighted sum of  $t$ -distributions. More precisely, formula (4.1) on page 380 of the cited reference (we use the notation  $k$  instead of  $n$ ) says that

$$\rho_T(v) = (2\pi)^{-\frac{1}{2}} \left\{ \sum_{r=0}^{m-1} \alpha_r \Gamma\left(r + \frac{3}{2}\right) \left(\frac{1}{2a} + \frac{v^2}{2}\right)^{-(r+3/2)} + \sum_{r=0}^{k-1} \beta_r \Gamma\left(r + \frac{3}{2}\right) \left(\frac{1}{2b} + \frac{v^2}{2}\right)^{-(r+3/2)} \right\}, \quad (2)$$

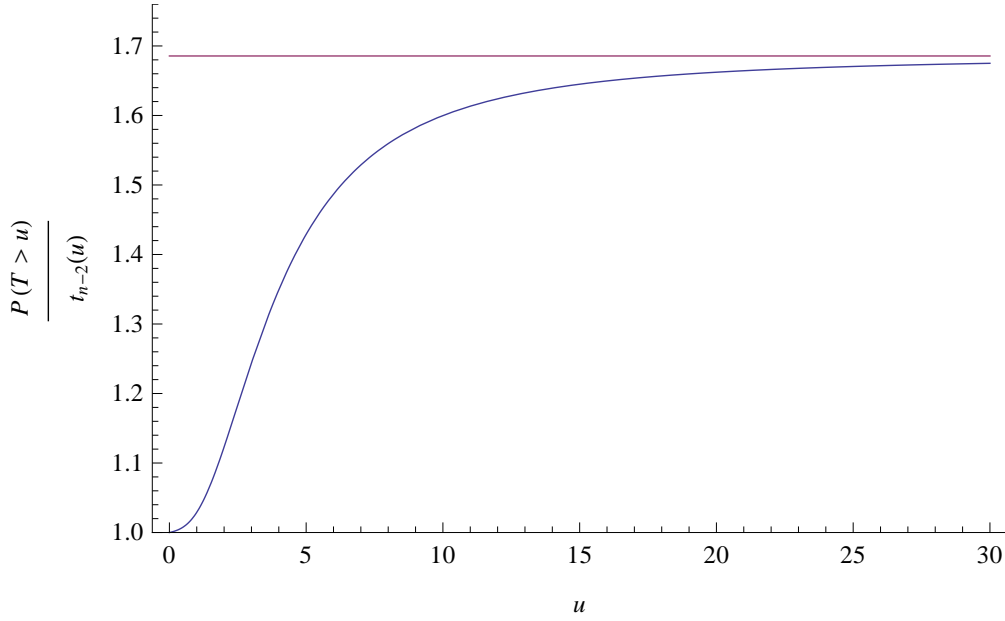
where  $\rho_T(v)$  is the p.d.f. of the Welch statistic  $T$  with odd sample sizes  $n_1 = 2m + 1$  and  $n_2 = 2k + 1$ . The constants  $a, b, \alpha_r$  and  $\beta_r$  depend on  $m, k, \sigma_1$  and  $\sigma_2$ , and the constants  $\alpha_r$  and  $\beta_r$  depend also on  $r$ .

In the light of the asymptotic representation of Theorem 3.1, the expression (2) for the exact density of  $T$  for odd sample sizes may look suspicious, since  $\rho_T(v)$  is a mixture of the Student  $t$ -densities that have tails heavier than  $t_{n-2}(u)$ , the tail of the Student  $t$ -density in Theorem 3.1. A more detailed investigation, however, revealed that  $\alpha_0 + \beta_0 = 0$ , and the main terms in the asymptotic expansion for the heaviest densities in (2) cancel out, bringing in summands of higher order, which, in turn, may cancel out the main terms of the asymptotic expansion of the next most heavy summands (i.e. the ones that correspond to  $r = 1$ ), and so on.

We tested our theory for the case  $n_1 = 3$ ,  $n_2 = 5$  and  $\sigma_1^2 = \sigma_2^2$  by computing the left-hand side of (11) using (2), see Figure 1, and also for other choices of  $n_1$  and  $n_2$  and for  $\sigma_1 \neq \sigma_2$ , see the attached Mathematica scripts in Supplementary Materials.

The figure below shows that the left-hand side of (11) converges to the constant  $K_g$ . Recall that the latter is obtained by substituting  $\alpha = 1/n_1$  and  $\beta = 1/n_2$  in (14). We also computed the first 10 terms in the Taylor series expansion for  $\rho_T(u)$  and  $K_g t_{n-2}(u)$  as  $u \rightarrow \infty$ , and found that the two expansions are in agreement with each other, see the corresponding Mathematica script in Supplementary Materials.

The asymptotic expressions for the Welch test under non-normality and dependence are obtained using the procedure similar to the one employed in the derivation of the asymptotic expressions for the Student two-sample  $t$ -test, see Section 3. We also study the accuracy of the asymptotic approximations of Corollary 3.1.1 for the Student two-sample  $t$ -test and Welch test using simulations, see Section 7.



**Figure 1.** Convergence in (11) for the Welch  $t$ -statistic. Sample sizes:  $n_1 = 3$  and  $n_2 = 5$ , variances equal. Blue curve is the left-hand side of (11) computed using the exact density of Welch's statistic, see (2). Horizontal line is the constant  $K_g \approx 1.68$  computed from (14).

### 1.5 Proof of the main theorem for the F-statistic for comparison of variances

This section contains the proofs of Theorem 5.1 and Corollary 5.1.1. For convenience we state them here as well.

**Theorem 1.2.** *If  $g$  is continuous and satisfies regularity constraints of Theorem 5.1, then*

$$\frac{\mathbf{P}(T > u)}{F_{n_1-1, n_2-1}(u)} = K_g + o(1) \quad \text{as } u \rightarrow \infty, \quad (3)$$

where  $F_{n_1-1, n_2-1}(u)$  is the tail of the  $F$ -distribution with parameters  $n_1 - 1$  and  $n_2 - 1$  and  $0 < K_g = K(g) < \infty$  is defined in (18) in Section 5.

*Proof.* Let  $A$  be an orthogonal linear operator defined on  $\mathbb{R}^{n_2}$  such that  $A\mathbf{e}_{n_2} = \mathbf{I}$ . We have

$$\mathbf{P}(T > u) = \int_D g(\mathbf{x}, A\mathbf{y}) d\mathbf{x} d\mathbf{y},$$

$$\text{where } D = \left\{ (\mathbf{x}, \mathbf{y}) : \frac{s_1^2(\mathbf{x})}{\sum_{i=1}^{n_2-1} y_i^2} > \frac{1}{n_2-1} u \right\}.$$

Changing variables  $y_i = (n_2 - 1)^{1/2} r s_1(\mathbf{x}) t_i$  for  $1 \leq i < n_2$  and  $y_{n_2} = r$  (though, formally, we should have considered the case  $r > 0$  and  $r < 0$  separately), we write

$$\mathbf{P}(T > u) = \int \cdots \int_{\sum_{i=1}^{n_2-1} t_i^2 < u^{-1}} G(\mathbf{t}) d\mathbf{t}, \quad (4)$$

where

$$G(\mathbf{t}) = (n_2 - 1)^{\frac{n_2-1}{2}} \int \cdots \int_{\mathbb{R}^{n_1}} s_1(\mathbf{x})^{n_2-1} \int_{-\infty}^{\infty} g(\mathbf{x}, r\mathbf{I} + s_1(\mathbf{x})A\mathbf{v}(\mathbf{t})) dr d\mathbf{x}$$

and

$$\mathbf{v}(\mathbf{t}) = (n_2 - 1)^{1/2} (t_1, t_2, \dots, t_{n_2-1}, 0).$$

Continuity of  $G$  at zero follows from the finiteness of integral (17) and continuity of  $g$  by the dominated convergence theorem, and Lemma 8.1 (A) in Section 1 of Appendix A implies that  $\mathbf{P}(T > u)$  is asymptotically proportional to  $t_{n_2-1}(\sqrt{u})$ . It can be shown that

$$\frac{t_{n_2-1}(\sqrt{u})}{F_{n_1-1, n_2-1}(u)} = \frac{\Gamma\left(\frac{n_2}{2}\right) \Gamma\left(\frac{n_1-1}{2}\right) (n_1 - 1)^{\frac{n_2-1}{2}}}{2\sqrt{\pi} \Gamma\left(\frac{n-2}{2}\right)} + o(1) \text{ as } u \rightarrow \infty,$$

and (3) and the expression for  $K_g$  in (18) follow.  $\square$

**Corollary 1.2.1** (Gaussian zero-mean case, independent samples). *If  $X$  and  $Y$  are independent zero-mean Gaussian random vectors with strictly non-degenerate covariance matrices  $\Sigma_1$  and  $\Sigma_2$ , then (3) holds with*

$$K_g = C \int \cdots \int_{\mathbb{R}^{n_1}} \frac{s_1(\mathbf{x})^{n_2-1}}{(1 + \mathbf{x}\Sigma_1^{-1}\mathbf{x}^T)^{n/2}} d\mathbf{x}, \quad (5)$$

where the constant  $C$  is given by

$$C = \frac{(n-2)(n_1-1)^{\frac{n_2-1}{2}} \Gamma\left(\frac{n_1-1}{2}\right) |\mathbf{I}\Sigma_2\mathbf{I}^T|^{1/2}}{2\pi^{\frac{n_1+1}{2}} |\Sigma_1|^{1/2} |\Sigma_2|^{1/2}}.$$

*Proof.* The regularity assumption (17) follows from Lemma 9.2 and the derivation of the constant  $K_g$  is a calculus exercise.  $\square$

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