

# Resampling Techniques and their Application

## -Class 9-

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  - Every subject (patient) is his/her own control
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  - Less costs (potentially)
- Measurements from the same subject **are not necessarily independent**

## Example

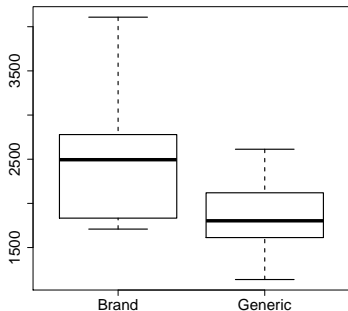
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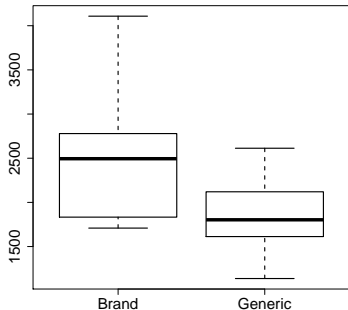
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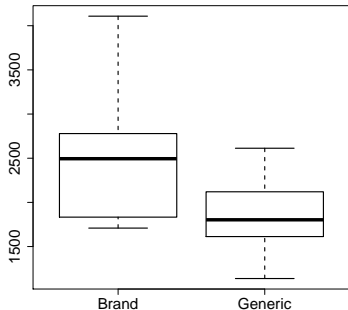
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- Aim:  $H_0 : \mu_1 = \mu_2$  and confidence interval

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- Measures the degree of the (linear) relationship between  $X_k$  and  $Y_k$
- On average,  $\underbrace{(X_k - \mu_1)}_{\leq 0} \underbrace{(Y_k - \mu_2)}_{\leq 0} \leq 0$

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- Reject  $H_0$ , if  $|T| \geq t_{1-\alpha/2}(n-1)$

## Example Evaluation

```
brand=c(4108,2526,2779,3852,1833, 2463,2059,1709,1829,2594)
generic=c(1755,1138,1613,2254,1310,2120,1851,1878,1682,2613)
plot(brand,generic,pch=19,cex=1.3)
n=length(brand)
x=cbind(brand,generic)
var(x)
```

```
diff=brand-generic
mD=mean(diff)
vd=var(diff)
```

```
T=sqrt(n)*mD/sqrt(vd)
pvalue=2*min(pt(T,n-1), 1-pt(T,n-1))
```

```
t.test(brand,generic,paired=TRUE)
```

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- Idea: Resample the distribution of  $T$
- But how? Differences?

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    - Resampling the differences
    - Resampling from all data and thus ignoring dependencies



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- In R: `sample(x, replace=TRUE)`
- Also known as **Nonparametric Bootstrap**

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  - Asymmetric signs  $P(W_1 = \frac{1+\sqrt{5}}{2}) = \frac{\sqrt{5}-1}{2\sqrt{5}}$  and  $P(W_1 = \frac{1-\sqrt{5}}{2}) = \frac{\sqrt{5}+1}{2\sqrt{5}}$

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- Note that centering is not necessary (why?)

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- This method is equivalent to....

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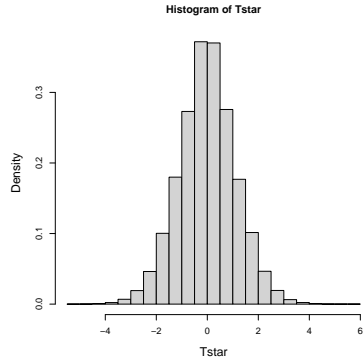
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- Reference: Konietschke and Pauly (2015)

# Illustration

```
x=brand
y=generic
plot(x,y,pch=19,cex=1.3)
n=10
d=x-y
T=sqrt(n)*mean(d)/sd(d)
pvalue=2*min(pt(T,n-1),1-pt(T,n-1))
pvalue
Tstar=c()
xy=c(x,y)
for(i in 1:100000){
  xstar=sample(xy) #permutation overall
  dstar=xstar[1:n]-xstar[(n+1):(2*n)]
  Tstar[i]= sqrt(n)*mean(dstar)/sd(dstar)
}
pstar= 2*min(mean(Tstar<=T),mean(Tstar>=T))
pstar
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- $\Phi(x)$ : CDF of  $N(0, 1)$

# Project

- In a paired data setting, permuting data overall and thus ignoring the dependency is somewhat counter intuitive. Verify the validity of the method for the paired  $t$ -test in a simulation study at 5% level of significance. Use  $n_{sim} = 10,000$  and  $n_{perm} = 10,000$  permutation runs. Generate bivariate normal data with variance  $\sigma_i^2 = 1$  and different covariances  $\sigma \in \{-0.95, -0.5, 0, 0.5, 0.95\}$  and sample sizes  $n \in 10, 20$ .