

STAT 850 Notes

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1 Randomized Block Design

Problem Setting:

- One block factor with b levels. and one treatment factor with t levels.
- Treatments randomized with blocks.
- No replicate observations. Our observed data y_{ij} represents the data in i th block with j th factor.

	Treatment				
	A	B	C	D	Mean
Block 1	89	88	97	94	92
Block 2	84	77	92	79	83
Block 3	81	87	87	85	85
Block 4	87	92	89	84	88
Block 5	79	81	80	88	82
Mean	84	85	89	86	86

Table 1.

Model without block-treatment interactions:

$$y_{ij} = \mu + b_i + t_j + \epsilon_{ij} \quad i \in 1:B, j = 1:T$$

with $\epsilon_{ij} \sim \text{iid } N(0, \sigma^2)$ and $\sum_j t_j = 0$. b_i s and t_j s indicate block and factor effects.

There are two settings for the block effects in the model:

- **Fixed block effects:** $\sum_{i=1}^N b_i = 0$ (sum to zero constraint for block effects)
- **Random block effects:** $b_1, \dots, b_B \sim \text{iid } N(0, \sigma_B^2)$

Data decomposition: the observed data can be decomposed to a summation of the (estimated) effects:

$$\begin{aligned} y_{ij} &= y_{..} + (y_{i.} - y_{..}) + (y_{.j} - y_{..}) + (y_{ij} - y_{i.} - y_{.j} + y_{..}) \\ &= \hat{\mu} + \hat{b}_i + \hat{t}_j + \hat{\epsilon}_{ij} \end{aligned}$$

ANOVA decomposition: the total sum of squares can be decomposed to:

$$\underbrace{\sum_i \sum_j (y_{ij} - y_{..})^2}_{\text{Total sum of squares}} = t \underbrace{\sum_i (y_{i.} - y_{..})^2}_{\text{Blocks}} + b \underbrace{\sum_j (y_{.j} - y_{..})^2}_{\text{Treatments}} + \underbrace{\sum_i \sum_j (y_{ij} - y_{i.} - y_{.j} + y_{..})^2}_{\text{Errors}}$$

Source	SS	df	$\mathbb{E}(\text{MS} = \text{SS}/\text{df})$
Fixed block effects			
Blocks	$t \sum_i (y_{i.} - y_{..})^2$	$b - 1$	$\sigma^2 + t(b-1)^{-1} \sum_i b_i^2$
Treatments	$b \sum_j (y_{.j} - y_{..})^2$	$t - 1$	$\sigma^2 + b(t-1)^{-1} \sum_j t_j^2$
Error	$\sum_{ij} (y_{ij} - y_{i.} - y_{.j} + y_{..})^2$	$(b-1)(t-1)$	σ^2
Total	$\sum_{ij} (y_{ij} - y_{..})^2$	$bt - 1$	
Random block effects			
Blocks	$t \sum_i (y_{i.} - y_{..})^2$	$b - 1$	$\sigma^2 + t \sigma_b^2$
Treatments	$b \sum_j (y_{.j} - y_{..})^2$	$t - 1$	$\sigma^2 + b(t-1)^{-1} \sum_j t_j^2$
Error	$\sum_{ij} (y_{ij} - y_{i.} - y_{.j} + y_{..})^2$	$(b-1)(t-1)$	σ^2
Total	$\sum_{ij} (y_{ij} - y_{..})^2$	$bt - 1$	

Table 2. ANOVA table for Randomized Block Design

2 Factorial treatment structure

2.1 Model Setting

Consider an experiment with two factors P and Q (**P and Q may have interactions**) with levels $j = 1, 2, \dots, p$, and $k = 1, 2, \dots, q$, replicated r times ($l = 1, 2, \dots, r$), with model

$$y_{jkl} = \mu_{jk} + \epsilon_{jkl}$$

Group effect parameters:

- Grand mean: $\mu_{..} = (pq)^{-1} \sum_j \sum_k \mu_{jk}$
- Group means for factor P : $\mu_{j.} = q^{-1} \sum_k \mu_{jk}$
- Group means for factor Q : $\mu_{.k} = p^{-1} \sum_j \mu_{jk}$
- Effect of factor P : $p_j = \mu_{j.} - \mu_{..}$
- Effect of factor Q : $q_k = \mu_{.k} - \mu_{..}$

We have sum to zero constraints under this setting:

$$\sum_j p_j = 0, \quad \sum_k q_k = 0$$

Interaction effect parameters:

$$(pq)_{jk} = \mu_{jk} - (\mu_{..} + p_j + q_k) = (\mu_{jk} - \mu_{.k}) - (\mu_{j.} - \mu_{..})$$

Also, we have

$$\sum_j (pq)_{jk} = 0 \quad \text{for all } k, \quad \sum_k (pq)_{jk} = 0 \quad \text{for all } j$$

Then the model can be expand as:

$$\mu_{jk} = \mu_{..} + p_j + q_k + (pq)_{jk}. \tag{1}$$

Remark 1. p and q without subscripts denote the number of levels for factor P and Q , while p_i and q_j denote the effect parameter for each level.

Example 1. (4×4 Design)

μ_{jk}				$\mu_{j.}$	p_j	$(pq)_{jk}$			
4	10	20	30	16	-6	-3	-4	1	6
14	20	40	50	31	9	-8	-9	6	11
14	20	10	20	16	-6	7	6	-9	-4
20	30	30	20	25	3	4	7	2	-13
$\mu_{.k}$	13	20	25	30	22				
q_k	-9	-2	3	8					

Table 3. 4×4 design data table

2.2 Compare differences between treatments

To compare differences between treatments, we define **contrast** and **interaction contrast**:

Definition 1. A contrast for the main effects of factor P is defined as

$$C_P = \sum_{j=1}^p l_j \mu_{j.},$$

where l_1, \dots, l_p are coefficients with $\sum_{j=1}^p l_j = 0$.

Example 2. (Simple Pairwise Comparison)

$$C_P = \mu_{1.} - \mu_{2.}$$

Definition 2. An interaction contrast is defined as:

$$C_{PQ} = \sum_{j=1}^p \sum_{k=1}^q l_j m_k \mu_{jk},$$

where m_1, \dots, m_q are also coefficients with $\sum_{k=1}^q m_k = 0$.

Example 3. Test whether the difference between levels of P depends on the level of Q .

$$C_{PQ} = (\mu_{11} - \mu_{12}) - (\mu_{21} - \mu_{22})$$

Interpretation of main and interaction effects:

1. **Always start by checking main effects.** Interactions modify these effects and only make sense in that context.
2. **If interactions are negligible,** simplify the interpretation and focus on main effects.
3. **If 3 or higher order interactions are negligible,** but second-order interactions are significant, then we should focus on both main effects and second-order interactions.
4. If a two-factor interaction is **very important**, and its **mean square (MS) value is similar to the MS values for main effects**, then the best way to interpret results is by **looking at the mean values for two-factor combinations** rather than just reporting main effects.

5. If a **two-factor interaction is significant**, but **one or both main effects are much larger than the interaction**, then the interpretation should consider main effects first, with adjustments for interaction effects.

2.3 Least-squares estimation for an unreplicated 2×3 design

Consider a two-way factorial design with $p=2$ and $q=3$

$$\mu_{jk} = \mu_{..} + p_j + q_k + (pq)_{jk}$$

	1	2	3
1	y_{11}	y_{12}	y_{13}
2	y_{21}	y_{22}	y_{23}

Table 4.

$$\mathbf{y} = \begin{pmatrix} y_{11} \\ y_{12} \\ y_{13} \\ y_{21} \\ y_{22} \\ y_{23} \end{pmatrix} = \mathbf{X}\beta = \begin{pmatrix} 1 & 1 & 1 & 0 & 1 & 0 \\ 1 & 1 & 0 & 1 & 0 & 1 \\ 1 & 1 & -1 & -1 & -1 & -1 \\ 1 & -1 & 1 & 0 & -1 & 0 \\ 1 & -1 & 0 & 1 & 0 & -1 \\ 1 & -1 & -1 & -1 & 1 & 1 \end{pmatrix} \begin{pmatrix} \mu_{..} \\ p_1 \\ q_1 \\ q_2 \\ (pq)_{11} \\ (pq)_{12} \end{pmatrix} + \epsilon$$

The columns of \mathbf{X} w.r.t. different parameter groups are orthogonal. In this case, the columns with respect to p and columns with respect to q are orthogonal. Also, they are orthogonal to the columns w.r.t. pq . Therefore, $\mathbf{X}'\mathbf{X}$ appears to be block diagonal:

$$\mathbf{X}'\mathbf{X} = \begin{pmatrix} 6 & 0 & 0 & 0 & 0 & 0 \\ 0 & 6 & 0 & 0 & 0 & 0 \\ 0 & 0 & 4 & 2 & 0 & 0 \\ 0 & 0 & 2 & 4 & 0 & 0 \\ 0 & 0 & 0 & 0 & 4 & 2 \\ 0 & 0 & 0 & 0 & 2 & 4 \end{pmatrix}$$

$$(\mathbf{X}'\mathbf{X})^{-1} = \begin{pmatrix} 1/6 & 0 & 0 & 0 & 0 & 0 \\ 0 & 1/6 & 0 & 0 & 0 & 0 \\ 0 & 0 & 1/3 & -1/6 & 0 & 0 \\ 0 & 0 & -1/6 & 1/3 & 0 & 0 \\ 0 & 0 & 0 & 0 & 1/3 & -1/6 \\ 0 & 0 & 0 & 0 & -1/6 & 1/3 \end{pmatrix}$$

The least square solution gives

$$\hat{\beta} = (\mathbf{X}'\mathbf{X})^{-1}\mathbf{X}'\mathbf{y} = \begin{pmatrix} y_{..} \\ y_{1.} - y_{..} \\ y_{.1} - y_{..} \\ y_{.2} - y_{..} \\ y_{11} - y_{1.} - y_{.1} + y_{..} \\ y_{12} - y_{1.} - y_{.2} + y_{..} \end{pmatrix}.$$

Note that the number of parameter equals to the number of observations, \mathbf{X} is invertable, $\hat{\beta}$ is the solution to $\mathbf{X}\beta = \mathbf{y}$.

Now we consider an additive model without interactions:

$$\mu_{jk} = \mu_{..} + p_j + q_k,$$

the least square solution gives:

$$\hat{\beta} = (\mathbf{X}^T \mathbf{X})^{-1} \mathbf{X} \mathbf{y} = (y_{..} \quad y_{1.} - y_{..} \quad y_{.1} - y_{..} \quad y_{.2} - y_{..})^T,$$

which align with the solution of the model with interactions. This is due to the orthogonality.

2.4 Experiment with Replication

The model for an experiment with replication can be written as:

$$y_{jkl} = \mu + p_j + q_k + (pq)_{jk} + \epsilon_{jkl}, \quad j = 1, \dots, p; k = 1, \dots, q, l = 1, \dots, r,$$

where ϵ_{jkl} are i.i.d. $\mathcal{N}(0, \sigma^2)$. We can decompose data as:

$$\begin{aligned} y_{jkl} &= \hat{\mu} + \hat{p}_j + \hat{q}_k + (\widehat{pq})_{jk} + \hat{\epsilon}_{jkl} \\ &= y_{...} + (y_{j..} - y_{...}) + (y_{.k.} - y_{...}) + (y_{jk.} - y_{j..} - y_{.k.} + y_{...}) + (y_{jkl} - y_{jk.}). \end{aligned}$$

The sum of squares are defined as:

$$\begin{aligned} S_P &= q r \sum_j (y_{j..} - y_{...})^2 \\ S_Q &= p r \sum_k (y_{.k.} - y_{...})^2 \\ S_{PQ} &= r \sum_j \sum_k (y_{jk.} - y_{j..} - y_{.k.} + y_{...})^2 \\ S_R &= \sum_j \sum_k \sum_l (y_{jkl} - y_{jk.})^2 \\ S_D &= \sum_j \sum_k \sum_l (y_{jkl} - y_{...})^2 \end{aligned}$$

Source	SS	df	MS	Ratio
Factor P	$S_P = 1.03301$	$p - 1 = 2$	$s_P^2 = 0.51651$	$s_P^2 / s_R^2 = 23.22$
Factor Q	$S_B = 0.92121$	$q - 1 = 3$	$s_Q^2 = 0.30707$	$s_Q^2 / s_R^2 = 13.81$
Interaction	$S_{PQ} = 0.25014$	$(p - 1)(q - 1) = 6$	$s_{PQ}^2 = 0.04169$	$s_{PQ}^2 / s_R^2 = 1.87$
Residual	$S_R = 0.80073$	$p q (r - 1) = 36$	$s_R^2 = 0.02224$	
Total	$S_D = 3.00508$	$p q r - 1 = 47$		

Table 5. ANOVA table for two factors experiments with replications

2.5 Model Checking

1. Define the estimated value of μ_{jk} under the full model as $\hat{\mu}_{jk} = y_{jk.}$. Let $\tilde{\mu}_{jk} = y_{j..} + y_{.k.} - y_{...}$ denote the estimated value of μ_{jk} assuming no interactions.
2. To assess the homogeneity of variance in interactions, create a plot of the residuals $y_{jkl} - \hat{\mu}_{jk}$ against the fitted values $\hat{\mu}_{jk}$. A consistent spread of residuals across different values of $\hat{\mu}_{jk}$ suggests homogeneity, while a pattern or funnel shape may indicate variance issues.
3. To detect possible nonadditivity, plot $y_{jk.} - \tilde{\mu}_{jk}$ against $\tilde{\mu}_{jk}$. If the plot exhibits a curvilinear pattern, this suggests the presence of transformable nonadditivity, meaning that a transformation of the response variable may be necessary for a better model fit.

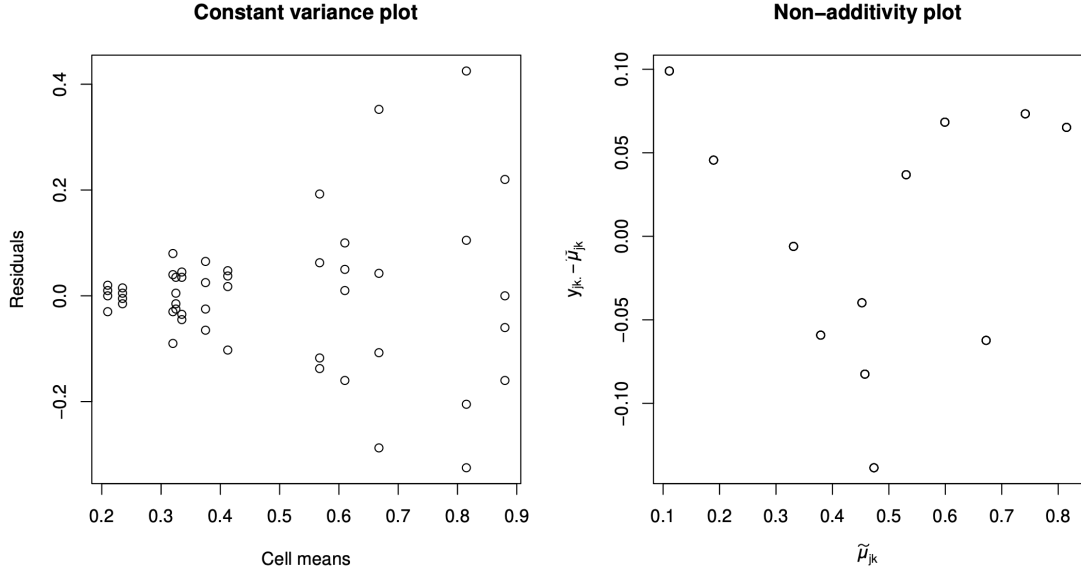


Figure 1. Model checking plots suggesting heteroscedasticity and non-additivity

2.6 Transformations

2.6.1 Taylor Power Transformation

Consider heterogeneous data $\text{Var}(y_{jkl}) = \sigma_{jk}^2$ and assume that $\sigma_{jk}^2 \propto \mu_{jk}^\beta$ for some β . We can use **Taylor power transformation** to deal with the heteroscedasticity (Assume $y_{jk} > 0$).

1. Calculate the sample mean y_{jk} and sample standard deviation s_{jk} for each (j, k) cell.
2. Fit a linear regression on $\log s_{jk} \sim \log y_{jk}$.
3. The fitted slope $\hat{\beta}$ is an estimate of β .
4. Use the transformation $(y^\lambda - 1)/\lambda$ with $\lambda := 1 - \hat{\beta}$.
5. If $\lambda = 0$, apply the log transformation $y_{jkl}^* = \log(y_{jkl})$.

Remark 2. Simple power transformation uses y^λ , which doesn't smoothly transit to $\log y$ as $\lambda \rightarrow 0$.

Justification for the method:

Define

$$z = f_\lambda(y) = \begin{cases} (y^\lambda - 1)/\lambda, & \lambda \neq 0 \\ \log y, & \lambda = 0 \end{cases} \quad (2)$$

By Taylor expansion:

$$z_{jkl} = f_\lambda(y_{jkl}) \approx f_\lambda(\mu_{jk}) + f'_\lambda(\mu_{jk})(y_{jkl} - \mu_{jk}).$$

Since $f'(y) = y^{\lambda-1}$,

$$\begin{aligned} \text{Var}(z_{jkl}) &\approx (f'_\lambda(\mu_{jk}))^2 \text{Var}(y_{jkl}) \\ &= \mu_{jk}^{2(\lambda-1)} \sigma_{jk}^2 \\ &\propto \mu_{jk}^{2(\lambda-1)} \mu_{jk}^{2\beta} \\ &= \mu_{jk}^{2(\lambda-1+\beta)}, \end{aligned}$$

and $\text{Var}(z_{jkl})$ becomes a constant if $\lambda = 1 - \beta$.

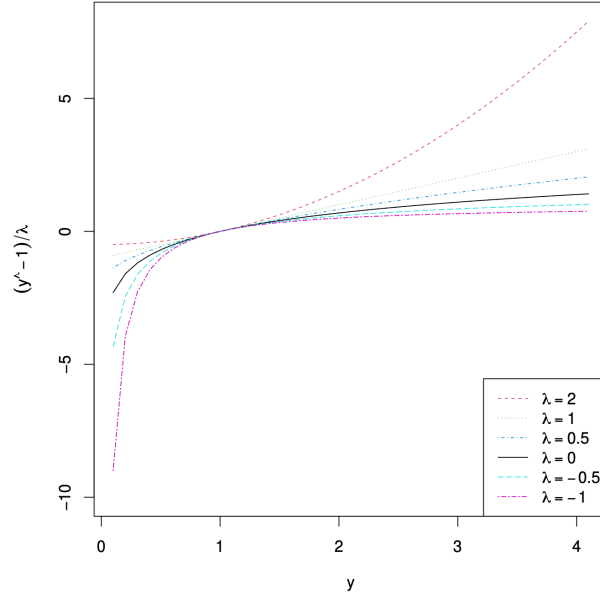


Figure 2. Power transformations

2.6.2 Box-Cox Transformation

Assumptions: There exists a λ such that $\{f_\lambda(y_i)\}_{i=1}^n$:

- are mutually independent
- are normally distributed
- have constant variance
- satisfy a linear model $f_\lambda(\mathbf{y}) = \mathbf{X}\beta + \epsilon$

2.6.3 Maximum Likelihood Estimation of Box-Cox λ

Assume $\mathbf{y}^{(\lambda)} = f_\lambda(\mathbf{y}) = \mathbf{X}\beta + \epsilon$ where f_λ is defined in (2) with $\epsilon \sim \mathcal{N}(0, \sigma^2 \mathbf{I})$ for some λ . Then the likelihood function for the untransformed data follows:

$$L(\lambda, \beta, \sigma) = \frac{1}{(2\pi\sigma^2)^{n/2}} \exp \left\{ -\frac{(\mathbf{y}^{(\lambda)} - \mathbf{X}\beta)^T (\mathbf{y}^{(\lambda)} - \mathbf{X}\beta)}{2\sigma^2} \right\} J(\lambda, \mathbf{y})$$

with Jacobian

$$J(\lambda, y) = \prod_{i=1}^n y_i^{\lambda-1}$$

We first find the LSEs of β and σ_λ for fixed λ :

$$\hat{\beta}_\lambda = (X^T X)^{-1} X^T \mathbf{y}^{(\lambda)}, \quad \hat{\sigma}_\lambda = \frac{(\mathbf{y}^{(\lambda)} - \mathbf{X}\hat{\beta}_\lambda)^T (\mathbf{y}^{(\lambda)} - \mathbf{X}\hat{\beta}_\lambda)}{n - p - 1},$$

then we have

$$\begin{aligned} \ell(\lambda, \hat{\beta}_\lambda, \hat{\sigma}_\lambda) &= \frac{\exp(-(n-p-1)/2)}{(2\pi)^{n/2} \hat{\sigma}_\lambda^n} J(\lambda, \mathbf{y}) \\ &= \hat{\sigma}_\lambda^{-n} \prod_{i=1}^n y_i^{\lambda-1} \frac{\exp(-(n-p-1)/2)}{(2\pi)^{n/2}} \end{aligned}$$

Let $\ell(\lambda) = \log L(\lambda, \hat{\beta}_\lambda, \hat{\sigma}_\lambda)$, and find the MLE $\hat{\lambda}$ that maximizes $\ell(\lambda)$.

The **confidence interval** follows from the standard result that the log-likelihood ratio statistic follows a chi-square distribution with 1 degree of freedom

$$2(\ell(\hat{\lambda}) - \ell(\lambda)) \sim \chi_1^2.$$

Then the confidence interval is the root for $\ell(\lambda) = \ell(\hat{\lambda}) - 0.5\chi_{1,\alpha}^2$.

Remark 3. The `boxcox` function in MASS library gives the MLE of λ .

```
library(MASS)
bc <- boxcox(y ~ p+q)
title(paste("Without interaction, lambda =", round(bc$x[which.max(bc$y)], 2)))
bc <- boxcox(y ~ p*q)
title(paste("With interaction, lambda =", round(bc$x[which.max(bc$y)], 2)))
```

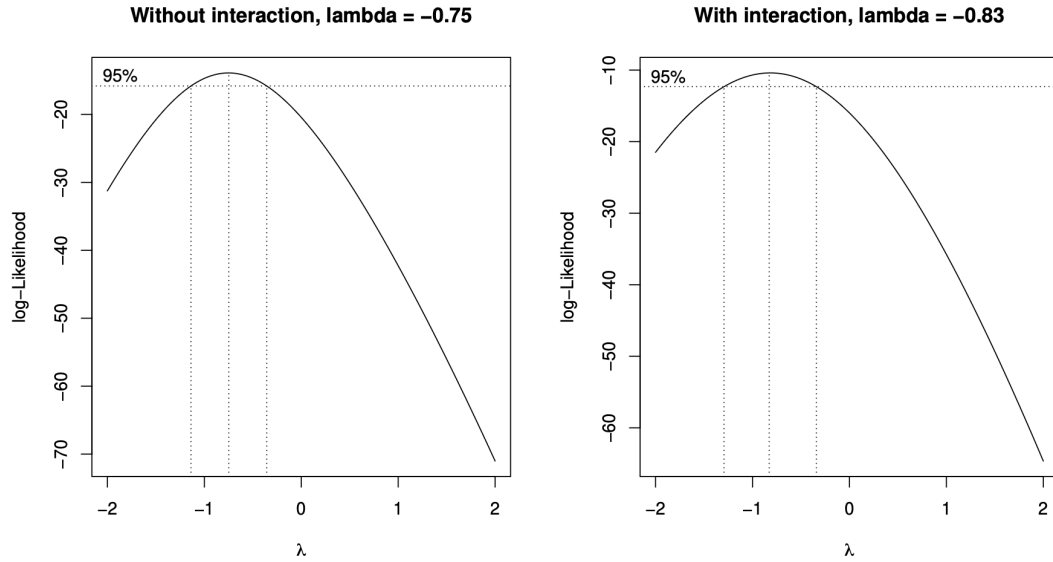


Figure 3. Likelihood of Box-Cox Transformation

Example 4. For the Poisson data, we present the ANOVA table after the two transformations.

	Df	Sum Sq	Mean Sq	F value	P-value
Poison	2	34.877	17.4386	70.6302	5.17e-13
Treatment	3	20.414	6.8048	27.5610	2.48e-09
Interaction	6	1.571	0.2618	1.0603	0.4046
Residuals	35	8.643	0.2469		
Poison	2	11.926	5.9631	66.5525	1.18e-12
Treatment	3	7.158	2.3860	26.6295	3.76e-09
Interaction	6	0.486	0.0810	0.9040	0.5032
Residuals	35	3.136	0.0896		

Table 6. ANOVA table after simple Taylor power transformation ($y^{(\lambda)} = y^\lambda$ with $\lambda = -1$) (above) and Box-Cox transformation (below).

Remark 4. The residual Df is reduced by 1 to compensate for the estimation of λ .

2.7 Confidence Intervals

2.7.1 When interactions are not significant

Let $u_j = \mu + p_j$ and $v_k = \mu + q_k$. Let the unbiased estimators be $\hat{u}_j = y_{j..}$ and $\hat{v}_k = y_{.k}$. with $\text{Var}(\hat{u}_j) = \sigma^2/(pr)$, $\text{Var}(\hat{v}_k) = \sigma^2/(qr)$. The sample standard deviation is

$$s(\hat{u}_j) = S_R/\sqrt{pr}, s(\hat{v}_k) = S_R/\sqrt{qr}.$$

Then $100(1 - \alpha)\%$ confidence interval for u_j is $\hat{u}_j \pm t_{\nu_R; \alpha/2} \times \text{se}(\hat{u}_j)$, where $\nu_R = pq(r - 1)$ is the degree of freedom for S_R .

For simultaneous confidence intervals, we define the contrast of interest be

$$L = \sum_{j=1}^p c_j u_j, \quad \text{where} \quad \sum_{j=1}^p c_j = 0$$

with its estimator

$$\hat{L} = \sum_{j=1}^p c_j y_{j..} \quad \text{with} \quad s(\hat{L}) = S_R \sqrt{(qr)^{-1} \sum_{j=1}^p c_j^2}$$

Then a $100(1 - \alpha)\%$ simultaneous confidence interval for L takes the form:

$$\hat{L} \pm T s(\hat{L}),$$

where T is a multiplier that depends on the type of the inference method used.

1. **Tukey's Method (Pairwise Comparisons):** Tukey's method is designed for simultaneous confidence intervals when comparing all possible pairwise differences between group means. The multiplier is:

$$T = \frac{q(p, \nu_R; \alpha)}{\sqrt{2}}$$

where $q(p, \nu_R; \alpha)$ is the studentized range statistic for p groups and residual degrees of freedom ν_R .

When comparing all pairs, each pairwise difference is a contrast (with coefficients $c_j = 1$ for one group and $c_j = -1$ for the other, and 0 elsewhere). Each interval is given by:

$$\text{CI for } (u_j - u_k): A C \quad (y_{j..} - y_{k..}) \pm \frac{q(p, \nu_R; \alpha)}{\sqrt{2}} s_R \sqrt{\frac{1^2 + (-1)^2}{qr}}$$

Here, the multiplier $\frac{q(p, \nu_R; \alpha)}{\sqrt{2}}$ is applied to each pairwise contrast.

2. **Scheffé's Method (All Contrasts):** Scheffé's method is more conservative and applies to all possible contrasts, not just pairwise comparisons. The multiplier is:

$$T = \sqrt{(p-1) F_{(p-1), \nu_R; \alpha}}$$

where $F_{(p-1), \nu_R; \alpha}$ is the critical value from the F-distribution with $p-1$ and ν_R degrees of freedom.

For any contrast $L = \sum_j c_j u_j$, the simultaneous confidence interval is:

$$\hat{L} \pm \sqrt{(p-1) F_{(p-1), \nu_R; \alpha}} S_R \sqrt{\frac{1}{qr} \sum_j c_j^2}$$

This interval applies to every possible contrast you might form.

3. **Bonferroni's Method (For g Comparisons):** The Bonferroni method controls the familywise error rate by adjusting the significance level for multiple comparisons. For any set of g comparisons, the multiplier is:

$$T = t_{\nu_R; \alpha/(2g)}$$

where $t_{\nu_R; \alpha/(2g)}$ is the t-distribution critical value with residual degrees of freedom ν_R and a Bonferroni-adjusted significance level of $\alpha/(2g)$.

If you have a specific set of g comparisons (contrasts) you plan to test, each interval is:

$$\hat{L} \pm t_{\nu_R; \alpha/(2g)} S_R \sqrt{\frac{1}{qr} \sum_j c_j^2}.$$

Each of the g contrasts gets its own interval, with the critical value adjusted by dividing α by $2g$.

2.7.2 When interactions are significant

In this case, each combination of factor levels has its own mean:

$$\mu_{jk} = \mu + p_j + q_k + (pq)_{jk}$$

If we are interested in comparing the means of two specific treatment combinations, say $\mu_{j_1 k_1}$ and $\mu_{j_2 k_2}$, we are comparing two of the pq treatments.

1. **Tukey's Method:** for all possible pairs of treatment means,

$$(\hat{\mu}_{j_1 k_1} - \hat{\mu}_{j_2 k_2}) \pm \frac{q(pq, \nu_R; \alpha)}{\sqrt{2}} S_R \sqrt{\frac{2}{r}}$$

where $q(pq, \nu_R; \alpha)$ is the quantile of the studentized range statistic for pq treatments and ν_R is the residual degrees of freedom.

2. **Bonferroni's Method:** for g pairs of comparisons,

$$(\hat{\mu}_{j_1 k_1} - \hat{\mu}_{j_2 k_2}) \pm t_{\nu_R; \alpha/(2g)} S_R \sqrt{\frac{2}{r}},$$

where $t_{\nu_R; \alpha/(2g)}$ is the quantile from t -distribution.

3. **Scheffé's Method for General Contrasts:** the contrast takes the form:

$$L = \sum_{j=1}^p \sum_{k=1}^q c_{jk} \mu_{jk}, \quad \text{with} \quad \sum_{j,k} c_{jk} = 0$$

The confidence interval for the contrast L is given by:

$$\hat{L} \pm \sqrt{(pq-1) F_{(pq-1), \nu_R; \alpha}} s(\hat{L})$$

where:

- $\hat{L} = \sum_{j,k} c_{jk} \hat{\mu}_{jk}$,
- $s(\hat{L}) = s_R \sqrt{\sum_{j,k} \frac{c_{jk}^2}{r}}$ (assuming balanced replication),
- $F_{(pq-1), \nu_R; \alpha}$ is the critical value from the F -distribution with $pq - 1$ and ν_R degrees of freedom.

2.8 Two-way Factorial with Blocks

2.8.1 No Replicates Within Blocks

Model:

$$y_{ijk} = \mu + b_i + t_{jk} + \varepsilon_{ijk} = \mu + b_i + p_j + q_k + (pq)_{jk} + \varepsilon_{ijk}$$

for $i = 1, \dots, b$, $j = 1, \dots, p$, and $k = 1, \dots, q$, with sum-to-zero constraints:

$$\sum_i b_i = \sum_j p_j = \sum_k q_k = 0.$$

Data decomposition:

$$\begin{aligned} y_{ijk} &= y_{...} + (y_{i..} - y_{...}) + (y_{.jk} - y_{...}) + (y_{ijk} - y_{i..} - y_{.jk} + y_{...}) \\ &= y_{...} + (y_{i..} - y_{...}) + (y_{.j.} - y_{...}) + (y_{..k} - y_{...}) \\ &\quad + (y_{.jk} - y_{.j.} - y_{..k} + y_{...}) + (y_{ijk} - y_{i..} - y_{.jk} + y_{...}) \end{aligned}$$

Source	SS	df
Blocks	$p q \sum_i (y_{i..} - y_{...})^2$	$b - 1$
P	$b q \sum_j (y_{.j.} - y_{...})^2$	$p - 1$
Q	$b p \sum_k (y_{..k} - y_{...})^2$	$q - 1$
PQ	$b \sum_j \sum_k (y_{.jk} - y_{.j.} - y_{..k} + y_{...})^2$	$(p - 1)(q - 1)$
Residual	$\sum_i \sum_j \sum_k (y_{ijk} - y_{i..} - y_{.jk} + y_{...})^2$	$(b - 1)(pq - 1)$
Total	$\sum_i \sum_j \sum_k (y_{ijk} - y_{...})^2$	$b p q - 1$

Table 7. ANOVA table for two-way factorial with blocks and without replicates

2.8.2 With Replicates Within Blocks

Model:

$$y_{ijkl} = \mu + b_i + t_{jk} + \varepsilon_{ijk} = \mu + b_i + p_j + q_k + (pq)_{jk} + \varepsilon_{ijkl}$$

or $i = 1, \dots, b$, $j = 1, \dots, p$, $l = 1, \dots, r$, and $k = 1, \dots, q$, with sum-to-zero constraints:

$$\sum_i b_i = \sum_j p_j = \sum_k q_k = 0.$$

Data decomposition:

$$\begin{aligned} y_{ijkl} &= y_{....} + (y_{i...} - y_{....}) + (y_{.j..} - y_{....}) + (y_{..k.} - y_{....}) \\ &\quad + (y_{.jk.} - y_{.j..} - y_{..k.} + y_{....}) + (y_{ijkl} - y_{i...} - y_{.jk.} + y_{....}) \end{aligned}$$

Source	SS	df
Blocks	$pqr \sum_i (y_{i...} - y_{....})^2$	$b - 1$
P	$bqr \sum_j (y_{.j..} - y_{....})^2$	$p - 1$
Q	$bpr \sum_k (y_{..k.} - y_{....})^2$	$q - 1$
PQ	$b \sum_j \sum_k (y_{.jk.} - y_{.j..} - y_{..k.} + y_{....})^2$	$(p - 1)(q - 1)$
Residual	$\sum_i \sum_j \sum_k \sum_l (y_{ijkl} - y_{i...} - y_{.jk.} + y_{....})^2$	$b p q r - b - p q + 1$
Total	$\sum_i \sum_j \sum_k \sum_l (y_{ijkl} - y_{....})^2$	$b p q r - 1$

Table 8.

2.8.3 Advantages of Two-level Factorial Designs

1. **Efficiency in runs:** They require only a few experimental runs per factor, making them cost-effective and time-saving even when several factors are under study.

Two-level factorial designs use only two settings per factor, which are chosen to maximize the contrast between high and low values. This strong contrast helps in estimating the main effects with greater clarity.

2. **Identifying Major Trends:** While they don't cover the entire range of possible factor values, these designs help reveal the main effects and trends, pointing toward areas that merit further investigation.
3. **Flexible Augmentation:** If more detailed local analysis is needed later, two-level designs can be expanded or combined with additional points (composite designs) to explore the factor space more thoroughly.
4. **Modular Design Building:** They serve as building blocks. By combining them appropriately, you can tailor the complexity of the experimental design to match the sophistication of the problem at hand.
5. **Precision in Effect Estimation:** With fewer runs and a clear structure, the estimates of the main effects of factors are generally precise.
6. They form the basis for two-level fractional factorial designs.

3 Experimental Design

3.1 Comparasion of designs

3.1.1 OFAT v.s. Factorial

OFAT Experiments: An experimental approach in which only **one factor (independent variable)** is varied at a time while keeping all other factors constant. The goal is to observe the effect of that single factor on the response variable.

2² Factorial Design: an experimental design where there are **two factors**, each with **two levels** (e.g., "low" and "high"). This results in **2² = 4 experimental runs**, covering all possible combinations of factor levels.

Example 5. OFAT Design: An engineer designed an experiment to compare a standard and a new gas anneal process by varying pressure and temperature across three runs:

1. Standard pressure & standard temperature
2. Standard pressure & new temperature

3. New pressure & new temperature

A single lot of 48 wafers was used, with 16 wafers assigned to each run.

Pressure	Temperature	
	Standard	New
Standard	16 wafers	16 wafers
New	-	16 wafers

Table 9. OFAT Design

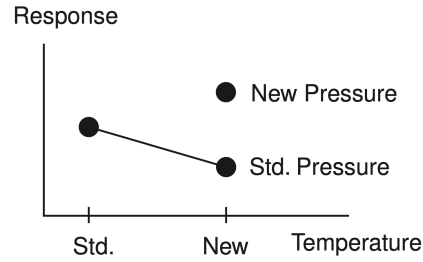


Figure 4. OFAT Design

2² Factorial Design:

Pressure	Temperature	
	Standard	New
Standard	12 wafers	12 wafers
New	12 wafers	12 wafers

Table 10. 2² Factorial Design

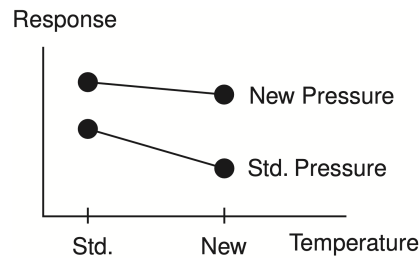


Figure 5. 2² Factorial Design

Advantages of 2² Factorial Design:

1. Estimates of the factor effects are more precise (all 48 wafers are used, while only 32 wafers in the OFAT design)
 - a. Variance of each effect is $\sigma^2/12$ for the 2² design.
 - b. Variance of each factor effect for the OFAT is $\sigma^2/8$ (50% more).
2. Interaction effect can be estimated.
3. 2² design provides data over a broader factor space.

3.1.2 Central Composite Design

Example 6. Two engineers planned an experiment for a rapid thermal anneal process. They wanted to study the sensitivity of the response sheet resistance to two factors—time and temperature. The followings are 3 different designs

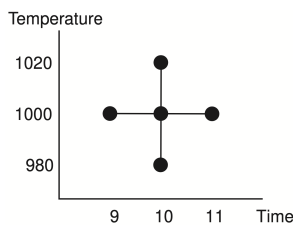


Figure 6. OFAT Design

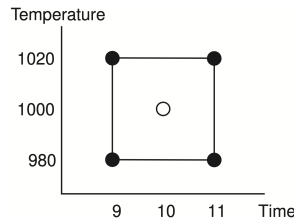


Figure 7. 2² Design

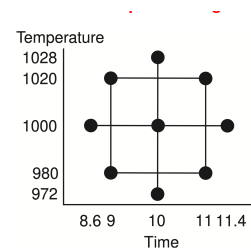


Figure 8. Central Composite

Central Composite Design (CCD): is a type of **Response Surface Methodology (RSM)** used for building a quadratic model in experimental designs. It helps **optimize processes** by exploring both linear and interaction effects, as well as curvature in factor responses. It contains the following components:

- **Factorial Points**: A full or fractional factorial design that represents the main effects and interactions.
- **Axial (Star) Points**: Points placed outside the factorial design to capture curvature and quadratic effects.
- **Center Points**: Replicated runs at the center of the design space to improve model accuracy and detect pure error.

Type	X_1	X_2
Factorial Points	$-1, +1$	$-1, +1$
Axial Points	$\pm\alpha$	$\pm\alpha$
Center Points	0	0

Table 11. Example of CCD with two factors

Advantages of Central Composite Design:

1. Estimates of factor effects are more precise
2. Interactions can be estimated
3. Curvature can be estimated in the entire space
4. Optimization is possible for the entire space
5. If center point is replicated 4 times, the design is rotatable, i.e., equal precision in estimation at all points equidistant from the center

3.1.3 2^3 Design

Example 7. 3-factor OFAT design & 2^3 experiment

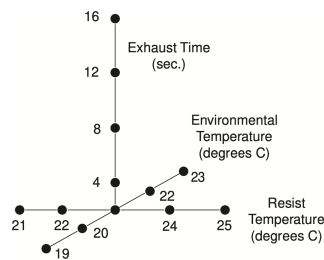


Figure 9. 3-factor OFAT in 15 runs

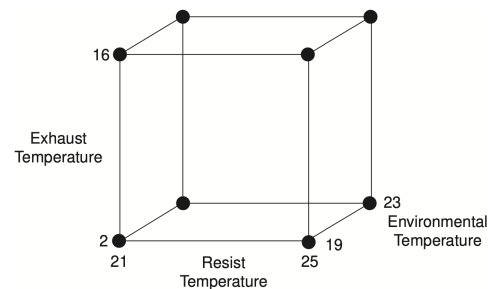


Figure 10. 2^3 experiment in 8 runs

Advantages of the 2^3 design:

1. Requires less resources: 8 runs instead of 15.
2. Estimates of factor effects are more accurate: all 8 runs are used in each estimate. But in OFAT, only 5 runs are used in each estimate.
3. Interaction effects can be estimated.
4. 2^3 design yields information over a larger region of the factor space

3.1.4 Box-Behnken design

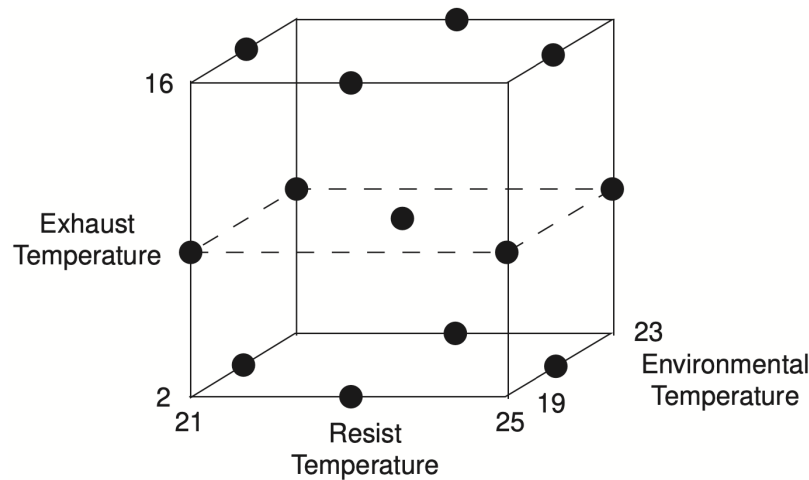


Figure 11. Box-Behnken design in 15 runs

Advantages of Box-Behnken Design:

1. Interactions can be estimated in Box-Behnken design.
2. Runs are more evenly spread out—hence more accurate predictions while OFAT can estimate curvature along 3 lines only.
3. Box-Behnken can estimate curvature in entire experimental region.
4. Box-Behnken design can be optimized over entire factor space while OFAT can be optimized along 3 lines only.

3.2 Analysis of Factorial Design

3.2.1 2^2 Design

We use the following notations to represent factors and interactions in a 2^2 design:

- Two factors A and B, each at two levels.
- 4 sets of experimental conditions. “Low” indicated by “−” and “high” indicated by “+”.
- Use notation (1), a , b , ab to denote the mean responses at the treatment combinations listed below:

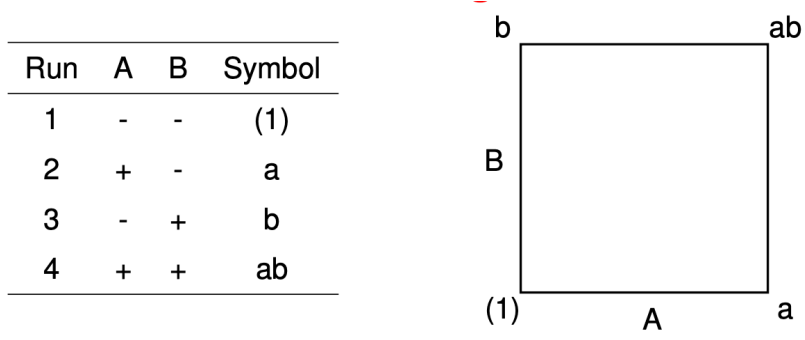


Figure 12. Notations in a 2^2 design

Model:

$$y_{ijk} = \eta + \alpha_i + \beta_j + (\alpha\beta)_{ij} + \varepsilon_{ijk}$$

with sum-to-zero constraints:

$$\alpha_1 + \alpha_2 = \beta_1 + \beta_2 = (\alpha\beta)_{11} + (\alpha\beta)_{12} = (\alpha\beta)_{21} + (\alpha\beta)_{22} = (\alpha\beta)_{11} + (\alpha\beta)_{21} = 0.$$

Define $\alpha_2 = \alpha$, $\beta_2 = \beta$, then we can write

$$\alpha_1 = -\alpha, \beta_1 = -\beta, (\alpha\beta)_{21} = -(\alpha\beta), (\alpha\beta)_{11} = (\alpha\beta), (\alpha\beta)_{12} = -(\alpha\beta)$$

i	j	A	B	Observation	Without constraints	With constraints
1	1	-	-	(1)	$\eta + \alpha_1 + \beta_1 + (\alpha\beta)_{11}$	$\eta - \alpha - \beta + (\alpha\beta)$
2	1	+	-	a	$\eta + \alpha_2 + \beta_1 + (\alpha\beta)_{21}$	$\eta + \alpha - \beta - (\alpha\beta)$
1	2	-	+	b	$\eta + \alpha_1 + \beta_2 + (\alpha\beta)_{12}$	$\eta - \alpha + \beta - (\alpha\beta)$
2	2	+	+	ab	$\eta + \alpha_2 + \beta_2 + (\alpha\beta)_{22}$	$\eta + \alpha + \beta + (\alpha\beta)$

Table 12. Summary of 2^2 design model

Then we have the least square estimates of the parameters:

$$\begin{aligned}\hat{\eta} &= [ab + a + b + (1)] / 4 = y... \\ \hat{\alpha} &= [ab + a - b - (1)] / 4 \\ \hat{\beta} &= [ab - a + b - (1)] / 4 \\ \widehat{(\alpha\beta)} &= [ab - a - b + (1)] / 4\end{aligned}$$

The **main effect of A** is defined as:

$$\begin{aligned}A &= 2\hat{\alpha} = [ab + a - b - (1)] / 2 \\ &= [ab + a] / 2 - [b + (1)] / 2 \\ &= (1/2) \{ [ab - b] + [a - (1)] \}\end{aligned}$$

The **interaction effect** is defined as:

$$\begin{aligned}AB &= 2\widehat{(\alpha\beta)} \\ &= [ab - a - b + (1)] / 2 \\ &= (1/2) \{ [ab - b] - [a - (1)] \} \\ &= (1/2) \{ [ab - a] - [b - (1)] \}\end{aligned}$$

3.2.2 Generalization to 2^3 Design

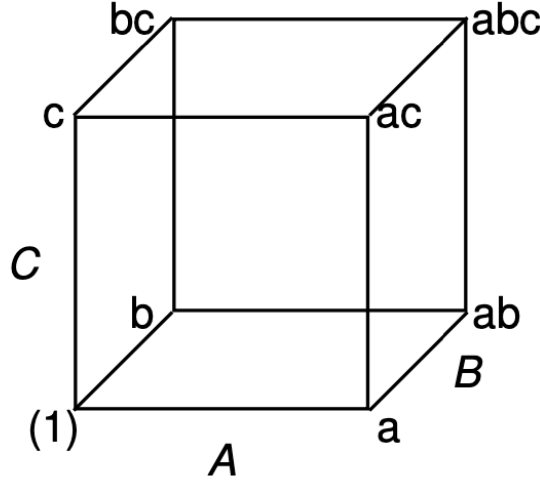


Figure 13. 2^3 design

Mean effect of A:

$$\begin{aligned}
 A &= (a + ab + ac + abc) / 4 - ((1) + b + c + bc) / 4 \\
 &= \text{mean of high A} - \text{mean of low A} \\
 &= (1/4) \{ (a - (1)) + (ab - b) + (ac - c) + (abc - bc) \} \\
 &= \text{mean of 4 univariate A effects} \\
 &= (1/2) [\{abc + ac - bc - c\} / 2 + \{ab + a - b - (1)\} / 2] \\
 &= (1/2) \{ A \text{ effect in A-B design at high C} + A \text{ effect in A-B design at low C} \}
 \end{aligned}$$

Interaction Effects:

- The 2^3 consists of two 2^2 designs: one at high C and one at low C
- AB interaction at low C is $AB_1 = (ab + (1) - a - b) / 2$
- AB interaction at high C is $AB_2 = (abc + c - ac - bc) / 2$
- AB and ABC effects for 2^3 are defined as

$$\begin{aligned}
 AB &= (1/2) (AB_2 + AB_1) = (1/4) (abc + c - ac - bc + ab + (1) - a - b) \\
 ABC &= (1/2) (AB_2 - AB_1) = (1/4) (abc + c - ac - bc - ab - (1) + a + b)
 \end{aligned}$$

Alternative Expressions:

The LSEs

$$\begin{aligned}
 \hat{\eta} &= [abc + ab + ac + bc + a + b + c + (1)] / 8 \\
 A &= [abc + ab + ac + a - bc - b - c - (1)] / 4 \\
 AB &= [abc + ab - ac - bc - a - b + c + (1)] / 4 \\
 ABC &= [abc - bc - ac - ab + a + b + c - (1)] / 4
 \end{aligned}$$

Combining the expressions, we have:

$$\begin{aligned}\hat{\eta} &= (a+1)(b+1)(c+1)/8 \\ A &= (a-1)(b+1)(c+1)/4 \\ AB &= (a-1)(b-1)(c+1)/4 \\ ABC &= (a-1)(b-1)(c-1)/4\end{aligned}$$

- Variance of any effect estimation is $\text{var}(\text{effect}) = 4 N^{-1} \sigma^2$, where N is total number of observations and $\sigma^2 = \text{var}(\epsilon)$
- Let s_i^2 be the estimate of σ^2 at the i th treatment combination ($i = 1, 2, \dots, g$). Let $\nu_i = r_i - 1$ be the degrees of freedom for s_i^2 , where r_i is the number of replicates in i th combination. Then the pooled estimate of σ^2 is:

$$s^2 = \frac{\nu_1 s_1^2 + \dots + \nu_g s_g^2}{\nu_1 + \dots + \nu_g}$$

- Confidence intervals for estimated effects may be obtained using the Bonferroni method.
- Hidden replication property: When estimating the effect of a single factor (say, A) in a 2^3 design (with factors A , B , and C), the precision of the estimate for A is the same as it would be if you had conducted an experiment with only A at two levels with the same number of replications (for example, 4 runs at low A and 4 runs at high A).

We may use a **table of contrast** to summary the result of a 2^3 design. The “Dot product” in this table refers to the inner product between each column (e.g. A) and the response vector \mathbf{y} , and Dividing it by $N/2$ gives the estimate of the corresponding effect.

	A	B	C	AB	AC	BC	ABC	y
	-1	-1	-1	1	1	1	-1	60
	1	-1	-1	-1	-1	1	1	72
	-1	1	-1	-1	1	-1	1	54
	1	1	-1	1	-1	-1	-1	68
	-1	-1	1	1	-1	-1	1	52
	1	-1	1	-1	1	-1	-1	83
	-1	1	1	-1	-1	1	-1	45
	1	1	1	1	1	1	1	80
Dot product	92	-20	6.0	6.0	40	0	2.0	
Division by $N/2$	23	-5	1.5	1.5	10	0	0.5	

Table 13. Table of contrast example

3.2.3 Regression Formulation

The model of 2^3 design can be written as:

$$\mathbb{E}(y_{ijk}) = \eta + \alpha_i + \beta_j + \gamma_k + (\alpha\beta)_{ij} + (\alpha\gamma)_{ik} + (\beta\gamma)_{jk} + (\alpha\beta\gamma)_{ijk}$$

For an observation y_l , Let $x_{il} = 1$ indicate the i th factor is at its high level and $x_{il} = -1$ indicate the i th factor is at its low level. Then we can write

$$\mathbb{E}(y_l) = \beta_0 + \beta_1 x_{1l} + \beta_2 x_{2l} + \beta_3 x_{3l} + \beta_{12} x_{1l} x_{2l} + \beta_{13} x_{1l} x_{3l} + \beta_{23} x_{2l} x_{3l} + \beta_{123} x_{1l} x_{2l} x_{3l}.$$

The LSEs are:

$$\begin{aligned}\hat{\eta} = \hat{\beta}_0 &= [a b c + a b + a c + b c + a + b + c + (1)] / 8 \\ A &= 2 \hat{\beta}_1 = [a b c + a b + a c + a - b c - b - c - (1)] / 4 \\ A B &= 2 \hat{\beta}_{12} = [a b c + a b - a c - b c - a - b + c + (1)] / 4 \\ A B C &= 2 \hat{\beta}_{123} = [a b c - b c - a c - a b + a + b + c - (1)] / 4, \text{ etc.}\end{aligned}$$

The general 2^k design model can be written in a matrix form:

$$\mathbf{y} = \mathbf{X}\boldsymbol{\beta} + \boldsymbol{\epsilon},$$

where \mathbf{y} is the vector of observations and when $k = 3$:

$$\mathbf{X} = \begin{pmatrix} 1 & -1 & -1 & -1 & 1 & 1 & 1 & -1 \\ 1 & 1 & -1 & -1 & -1 & -1 & 1 & 1 \\ 1 & -1 & 1 & -1 & -1 & 1 & -1 & 1 \\ 1 & 1 & 1 & -1 & 1 & -1 & -1 & -1 \\ 1 & -1 & -1 & 1 & 1 & -1 & -1 & 1 \\ 1 & 1 & -1 & 1 & -1 & 1 & -1 & -1 \\ 1 & -1 & 1 & 1 & -1 & -1 & 1 & -1 \\ 1 & 1 & 1 & 1 & 1 & 1 & 1 & 1 \end{pmatrix}, \quad \boldsymbol{\beta} = \begin{pmatrix} \eta \\ \alpha \\ \beta \\ \gamma \\ (\alpha\beta) \\ (\alpha\gamma) \\ (\beta\gamma) \\ (\alpha\beta\gamma) \end{pmatrix}$$

Note that \mathbf{X} is an orthogonal matrix, then $\mathbf{X}^T \mathbf{X} = 2^k \mathbf{I}$ is an diagonal matrix, and we have

$$\hat{\boldsymbol{\beta}} = 2^{-k} \mathbf{X}^T \mathbf{y}.$$

3.3 Yates Algorithm for 2^k Experiments

Yates algorithm is a systematic computational procedure used to analyze data from a full 2^k factorial experiment. In these experiments, you have k factors, each at two levels (often labeled as "low" and "high"). The algorithm helps you calculate the main effects and interaction effects quickly and efficiently.

1. Order the Data:

List the experimental data in the standard order (the order shown in the 1st column of table 14).

2. Construct k New Columns:

For each new column (starting from the first column of responses), perform the following:

- **Addition (First Half):**

For the first 2^{k-1} entries, form each new entry by adding pairs of consecutive entries from the previous column. That is, for row i , compute:

$$\text{New entry} = 2i - 1 + 2i.$$

- **Subtraction (Second Half):**

For the remaining 2^{k-1} entries, compute each new entry by subtracting the entry in row $2i - 1$ from the entry in row $2i$ in the previous column:

$$\text{New entry} = 2i - 2i - 1.$$

3. Create a Scaling Column:

Add an additional column (the $(k+1)$ th column) where all entries are 2^{k-1} , except the first entry, which is 2^k .

4. **Compute the Estimates:**

Obtain the effect estimates by dividing the entries in the k th column by the corresponding entries in the $(k + 1)$ th (scaling) column.

5. **Identify the Effects:**

Determine the identity of each effect by inspecting the signs of the factors along each row in the table.

A	B	C	y	(1)	(2)	(3)	Div.	Est.	Effect
-1	-1	-1	60	132	254	514	8	64.25	\bar{y}
1	-1	-1	72	122	260	92	4	23.00	A
-1	1	-1	54	135	26	-20	4	-5.00	B
1	1	-1	68	125	66	6	4	1.50	AB
-1	-1	1	52	12	-10	6	4	1.50	C
1	-1	1	83	14	-10	40	4	10.00	AC
-1	1	1	45	31	2	0	4	0.00	BC
1	1	1	80	35	4	2	4	0.50	ABC

Table 14. Example of data table created by Yates algorithm

3.4 Factorial Design with blocks

3.4.1 2^2 Design in 2 Blocks of Size 2

Let $x_1, x_2, z \in \{-1, 1\}$, where x_1, x_2 indicates levels of A and B and z indicates the block variables. The model is defined as

$$y = \beta_0 + \beta_1 x_1 + \beta_2 x_2 + \beta_{12} x_1 x_2 + \delta z + \varepsilon$$

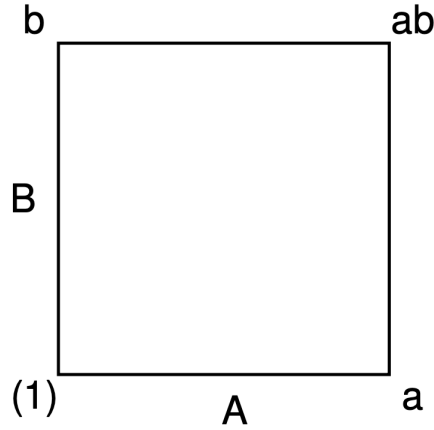


Figure 14. 2^2 design with blocks

Consider 3 possible designs:

- D1. (1), a in block 1; b, ab in block 2
- D2. (1), b in block 1; a, ab in block 2
- D3. (1), ab in block 1; a, b in block 2

In D1, B is confounded by the block effect because $z = x_2$ leads to:

$$\mathbb{E}(ab + b - a - (1)) = 4\beta_2 + 4\delta,$$

while A and AB are not counfounded.

3.4.2 2^3 in 2 Blocks of Size 4

OFAT Block Design:

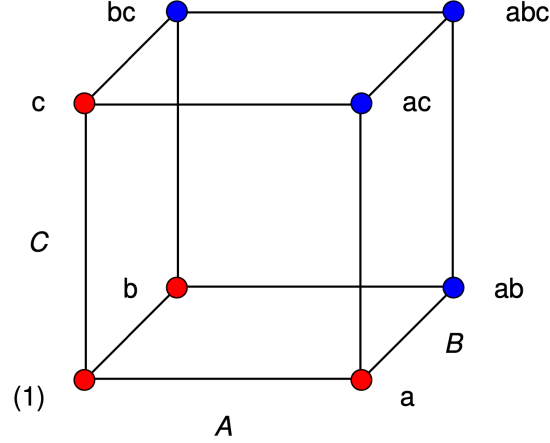


Figure 15. OFAT block design. Different colors indicate the experiment is conducted in different blocks.

	A	B	C	AB	AC	BC	ABC	Block
(1)	-	-	-	+	+	+	-	-
a	+	-	-	-	-	+	+	-
b	-	+	-	-	+	-	+	-
ab	+	+	-	+	-	-	-	+
c	-	-	+	+	-	-	+	-
ac	+	-	+	-	+	-	-	+
bc	-	+	+	-	-	+	-	+
abc	+	+	+	+	+	+	+	+

Table 15. Treatment table of OFAT block design

The corresponding model is:

$$y = \beta_0 + \beta_1 x_1 + \beta_2 x_2 + \beta_3 x_3 + \beta_{12} x_1 x_2 + \beta_{13} x_1 x_3 + \beta_{23} x_2 x_3 + \beta_{123} x_1 x_2 x_3 + \delta z + \epsilon \quad (3)$$

A is confounded in the design shown in Figure 15 and Table 15 because the expression

$$A = [abc + ab + ac + a - bc - b - c - (1)] / 4$$

cannot eliminate δ if we plug in (3).

We may change the design of the experiment to make the parameters of interest unfounded while some other parameters confounded. As shown in tabe 16, we can meke the assignment of blocks exactly the same as the assignment of the 3-way interaction ABC. By doing this, only ABC is confounded in our data. However, we can never make all the paramters unconfounded if we have to impose 2 blocks of size 4.

	A	B	C	AB	AC	BC	ABC	Block
(1)	-	-	-	+	+	+	-	-
a	+	-	-	-	-	+	+	+
b	-	+	-	-	+	-	+	+
ab	+	+	-	+	-	-	-	-
c	-	-	+	+	-	-	+	+
ac	+	-	+	-	+	-	-	-
bc	-	+	+	-	-	+	-	-
abc	+	+	+	+	+	+	+	+

Table 16. Experimental design such that only ABC is confounded.

Remark 5. In a 2^k design in 2 blocks, we usually use the largest-order interaction column to define the assignment of blocks such that only the highest order interaction is confounded.

3.4.3 2^3 in 4 Blocks of Size 2

If we have 4 blocks, we need two block variables $X, Y \in \{-1, 1\}$ to represent the block combinations in the table.

	A	B	C	AB	AC	BC	ABC	X	Y
(1)	-	-	-	+	+	+	-	+	+
a	+	-	-	-	-	+	+	-	+
b	-	+	-	-	+	-	+	-	-
ab	+	+	-	+	-	-	-	+	-
c	-	-	+	+	-	-	+	+	-
ac	+	-	+	-	+	-	-	-	-
bc	-	+	+	-	-	+	-	-	+
abc	+	+	+	+	+	+	+	+	+

Table 17. 2^3 design with blocks of size 2

If we put the columns $X = AB, Y = BC$, then all the two-way interactions are confounded while the other parameters are not.

3.4.4 Partial Confounding