

# Lecture 12: Mixed-Design ANOVA

# Combining between-group & within-subject

In the simplest 2 x 2 design you would have subjects randomly assigned to one of two groups (two study systems, Kaplan vs Princeton), but each group would experience 2 conditions (measurements).

	GRE - before	GRE - after
Kaplan	S <sub>1</sub>	S <sub>1</sub>
	S <sub>2</sub>	S <sub>2</sub>
	S <sub>3</sub>	S <sub>3</sub>
	S <sub>4</sub>	S <sub>4</sub>
	S <sub>5</sub>	S <sub>5</sub>
Princeton	S <sub>6</sub>	S <sub>6</sub>
	S <sub>7</sub>	S <sub>7</sub>
	S <sub>8</sub>	S <sub>8</sub>
	S <sub>9</sub>	S <sub>9</sub>
	S <sub>10</sub>	S <sub>10</sub>

# Another example

批准文号 国药准字H20190031

产品名称 甘露特钠胶囊

英文名称 Sodium Oligomannate Capsules

商品名 九期一

剂型 胶囊剂

规格 150mg

上市许可持有人 上海绿谷制药有限公司

生产单位 上海绿谷制药有限公司

生产地址 上海市青浦区汇金路886号

产品类别 化学药品

批准日期 2019-11-02

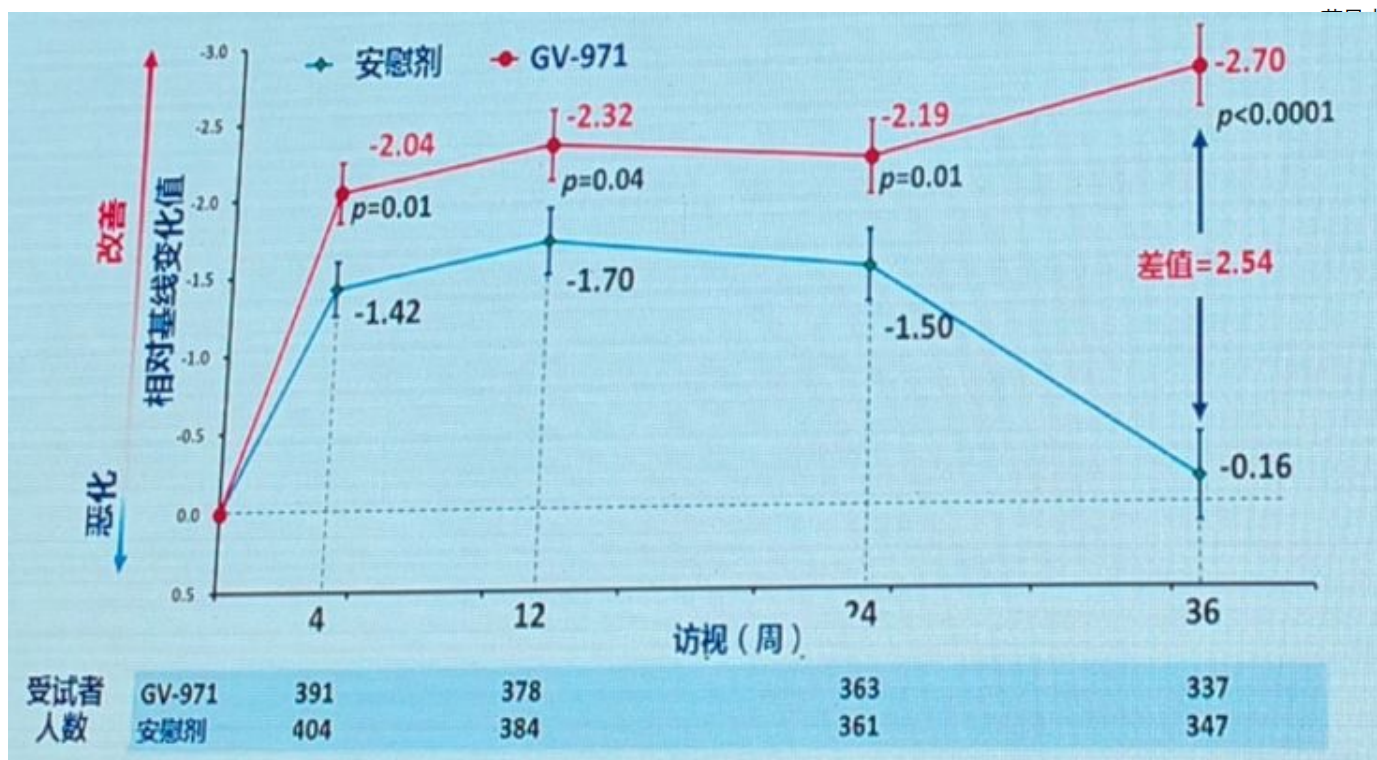
原批准文号

药品本位码 86900722000145

药品本位码备注

药品广告

据库查询 中药保护品种库



# Analysis

		$b_1$	$b_2$	$b_3$
Randomized Groups	$a_1$	$S_1$	$S_1$	$S_1$
		$S_2$	$S_2$	$S_2$
		$S_3$	$S_3$	$S_3$
		$S_4$	$S_4$	$S_4$
		$S_5$	$S_5$	$S_5$
	$a_2$	$S_6$	$S_6$	$S_6$
		$S_7$	$S_7$	$S_7$
		$S_8$	$S_8$	$S_8$
		$S_9$	$S_9$	$S_9$
		$S_{10}$	$S_{10}$	$S_{10}$
	$a_3$	$S_{11}$	$S_{11}$	$S_{11}$
		$S_{12}$	$S_{12}$	$S_{12}$
		$S_{13}$	$S_{13}$	$S_{13}$
		$S_{14}$	$S_{14}$	$S_{14}$
		$S_{15}$	$S_{15}$	$S_{15}$

# Sources of Variance

- $SS_T = SS_{BG} + SS_{WS}$
- What are the sources of variance?
  - **A** (between-group factor)
  - **S/A** (deviation of each subject from its A group mean)
  - **B** (repeated-measure factor)
  - **AB** (interaction of two factors)
  - **BxS/A** (deviation of each subject from its cell mean, minus S/A)
  - **T** (total variance).
- Degrees of freedom?

# Degree of freedom

- $A = a - 1$
- $S/A = a(s - 1) = as - a$   
*(s is the number of subjects in each A condition)*
- $B = b - 1$
- $AB = (a - 1)(b - 1)$
- $B \times S/A = a(b - 1)(s - 1)$
- $T = abs - 1$  or  $N - 1$

# Example – Books by Month

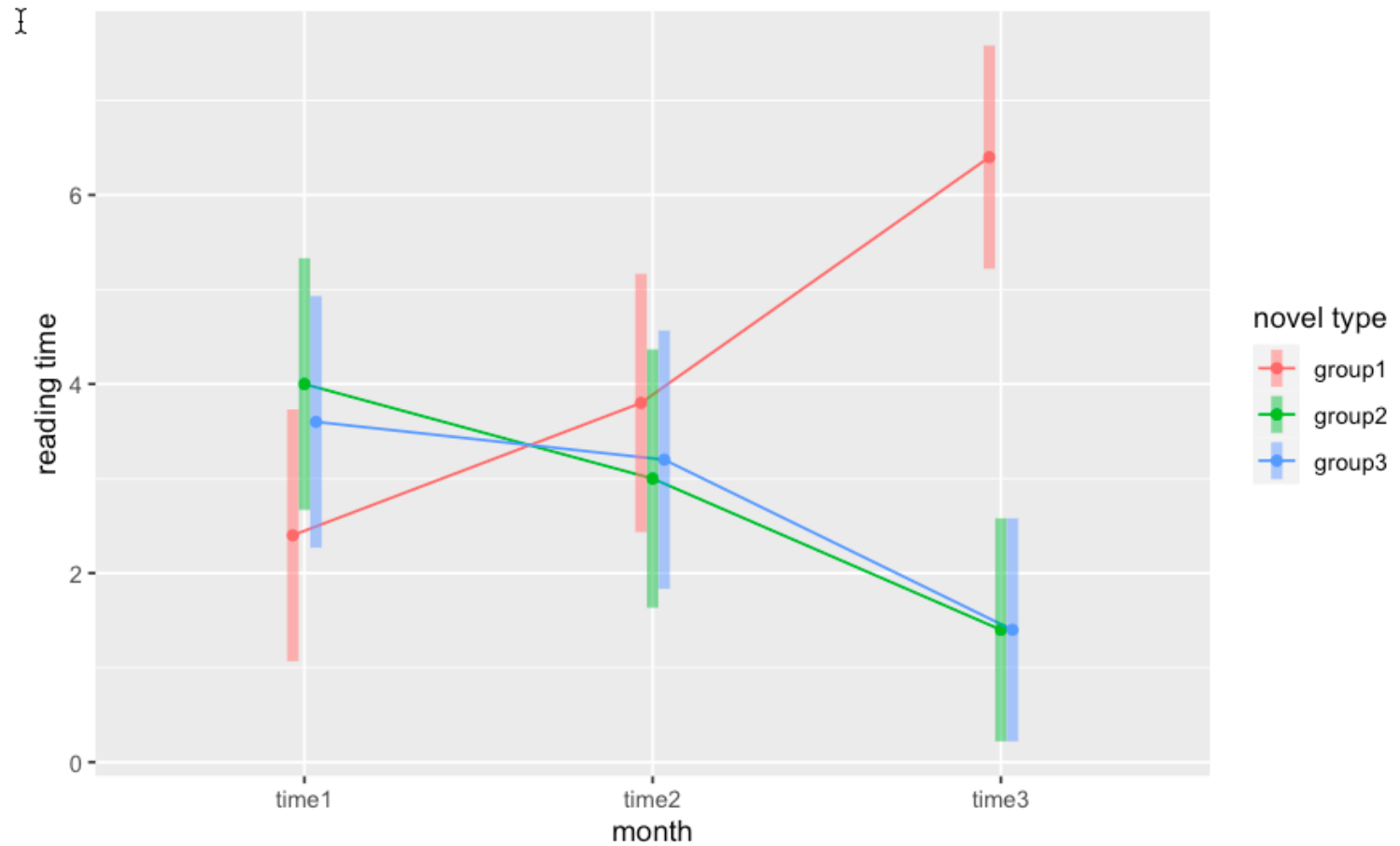
## Example:

- Imagine if we designed a study concerning reading different novels over time
- we randomly assign subjects to three types of books and have them read for three months
- Book type is the group factor, month is the repeated-measure factor.
- How many they read are measured.

			B: <i>Month</i>			Case Means
			$b_1$ : <i>Month 1</i>	$b_2$ : <i>Month 2</i>	$b_3$ : <i>Month 3</i>	
A: <i>Type of Novel</i>	: <i>Science Fiction</i>	$S_1$	1	3	6	$S_1 = 3.333$
		$S_2$	1	4	8	$S_2 = 4.333$
		$S_3$	3	3	6	$S_3 = 4$
		$S_4$	5	5	7	$S_4 = 5.667$
		$S_5$	2	4	5	$S_5 = 3.667$
			$a_1 b_1 = 2.4$	$a_1 b_2 = 3.8$	$a_1 b_3 = 6.4$	$a_1 = 4.2$
	$a_2$ : <i>Mystery</i>	$S_6$	3	1	0	$S_6 = 1.333$
		$S_7$	4	4	2	$S_7 = 3.333$
		$S_8$	5	3	2	$S_8 = 3.333$
		$S_9$	4	2	0	$S_9 = 2$
		$S_{10}$	4	5	3	$S_{10} = 4$
		$a_2$	$a_2 b_1 = 4$	$a_2 b_2 = 3$	$a_2 b_3 = 1.4$	$a_2 = 2.8$
	$a_3$ : <i>Romance</i>	$S_{11}$	4	2	0	$S_{11} = 2$
		$S_{12}$	2	6	1	$S_{12} = 3$
		$S_{13}$	3	3	3	$S_{13} = 3$
		$S_{14}$	6	2	1	$S_{14} = 3$
		$S_{15}$	3	3	2	$S_{15} = 2.667$
			$a_3 b_1 = 3.6$	$a_3 b_2 = 3.2$	$a_3 b_3 = 1.4$	$a_3 = 2.733$
			$b_1 = 3.333$	$b_2 = 3.333$	$b_3 = 3.067$	GM = 3.244



# Eyeball it first



# Sums of Squares

i: subject, 1~5

j: group factor A (novel), 1~3

k: within-subject factor B (month), 1~3

The total variability can be partitioned into A, B, AB, S/A, and B\*S/A

$$SS_{Total} = SS_A + SS_B + SS_{AB} + SS_{S/A} + SS_{B*S/A}$$

$$\begin{aligned} \sum (Y_{ijk} - \bar{Y}_{...})^2 &= \sum n_j (\bar{Y}_{.j.} - \bar{Y}_{...})^2 + \sum n_k (\bar{Y}_{..k} - \bar{Y}_{...})^2 + \\ &+ \left[ \sum n_{jk} (\bar{Y}_{.jk} - \bar{Y}_{...})^2 - \sum n_j (\bar{Y}_{.j.} - \bar{Y}_{...})^2 - \sum n_k (\bar{Y}_{..k} - \bar{Y}_{...})^2 \right] \\ &+ k \sum (\bar{Y}_{i..} - \bar{Y}_{.j.})^2 + \left[ \sum (Y_{ijk} - \bar{Y}_{.jk})^2 - k \sum (\bar{Y}_{i..} - \bar{Y}_{.j.})^2 \right] \end{aligned}$$

$$SS_A = \sum n_j (\bar{Y}_{.j} - \bar{Y}_{...})^2 = 15 * [(4.2 - 3.244)^2 + (2.8 - 3.244)^2 + (2.733 - 3.244)^2] = 20.583$$

$$SS_B = \sum n_k (\bar{Y}_{..k} - \bar{Y}_{...})^2 = 15 * [(3.333 - 3.244)^2 + (3.333 - 3.244)^2 + (3.067 - 3.244)^2] = .708$$

$$SS_{AB} = \left[ \sum n_{jk} (\bar{Y}_{.jk} - \bar{Y}_{...})^2 - \sum n_j (\bar{Y}_{.j} - \bar{Y}_{...})^2 - \sum n_k (\bar{Y}_{..k} - \bar{Y}_{...})^2 \right] =$$

$$\begin{aligned} \sum n_{jk} (\bar{Y}_{.jk} - \bar{Y}_{...})^2 = & 5 * [(2.4 - 3.244)^2 + (3.8 - 3.244)^2 + (6.4 - 3.244)^2 + \\ & + (4 - 3.244)^2 + (3 - 3.244)^2 + (1.4 - 3.244)^2 + \\ & + (3.6 - 3.244)^2 + (3.2 - 3.244)^2 + (1.4 - 3.244)^2] = 92.711 \end{aligned}$$

$$SS_{AB} = 92.711 - 20.583 - .708 = 71.420$$

This k is necessary; 3 months.

The deviation of each subject from his group mean

$$SS_{S/A} = k \sum (\bar{Y}_{i..} - \bar{Y}_{.j.})^2 = 3 * [(3.333 - 4.2)^2 + (4.333 - 4.2)^2 + (4 - 4.2)^2 + \\ + (5.667 - 4.2)^2 + (3.667 - 4.2)^2 + (3.333 - 2.8)^2 + (1.333 - 2.8)^2 + (3.333 - 2.8)^2 + \\ + (3.333 - 2.8)^2 + (2 - 2.8)^2 + (2 - 2.733)^2 + (3 - 2.733)^2 + (3 - 2.733)^2 + \\ + (2.667 - 2.733)^2] = 26.400$$

The deviation of each subject (within a cell) from its cell mean

$$\begin{aligned}
 SS_{B*S/A} &= \left[ \sum (Y_{ijk} - \bar{Y}_{.jk})^2 - k \sum (\bar{Y}_{i..} - \bar{Y}_{.j.})^2 \right] = \\
 &\sum (Y_{ijk} - \bar{Y}_{.jk})^2 = (1-2.4)^2 + (1-2.4)^2 + (3-2.4)^2 + (5-2.4)^2 + (2-2.4)^2 + \\
 &+ (3-3.8)^2 + (4-3.8)^2 + (3-3.8)^2 + (5-3.8)^2 + (4-3.8)^2 + \\
 &+ (6-6.4)^2 + (8-6.4)^2 + (6-6.4)^2 + (7-6.4)^2 + (5-6.4)^2 + \\
 &+ (3-4)^2 + (4-4)^2 + (5-4)^2 + (4-4)^2 + (4-4)^2 + \\
 &+ (1-3)^2 + (4-3)^2 + (3-3)^2 + (2-3)^2 + (5-3)^2 + \\
 &+ (0-1.4)^2 + (2-1.4)^2 + (2-1.4)^2 + (0-1.4)^2 + (3-1.4)^2 + \\
 &+ (4-3.6)^2 + (2-3.6)^2 + (3-3.6)^2 + (6-3.6)^2 + (3-3.6)^2 + \\
 &+ (2-3.2)^2 + (6-3.2)^2 + (3-3.2)^2 + (2-3.2)^2 + (3-3.2)^2 + \\
 &+ (0-1.4)^2 + (1-1.4)^2 + (3-1.4)^2 + (1-1.4)^2 + (2-1.4)^2 = 63.6 \\
 SS_{B*S/A} &= 63.6 - 26.4 = 37.2
 \end{aligned}$$

The deviation of each data point (within a cell) from the global mean

$$SS_{Total} = \sum (Y_{ijk} - \bar{Y}_{...})^2 =$$

$$\begin{aligned} SS_{Total} = & (1-3.244)^2 + (1-3.244)^2 + (3-3.244)^2 + (5-3.244)^2 + (2-3.244)^2 + \\ & + (3-3.244)^2 + (4-3.244)^2 + (3-3.244)^2 + (5-3.244)^2 + (4-3.244)^2 + \\ & + (6-3.244)^2 + (8-3.244)^2 + (6-3.244)^2 + (7-3.244)^2 + (5-3.244)^2 + \\ & + (3-3.244)^2 + (4-3.244)^2 + (5-3.244)^2 + (4-3.244)^2 + (4-3.244)^2 + \\ & + (1-3.244)^2 + (4-3.244)^2 + (3-3.244)^2 + (2-3.244)^2 + (5-3.244)^2 + \\ & + (0-3.244)^2 + (2-3.244)^2 + (2-3.244)^2 + (0-3.244)^2 + (3-3.244)^2 + \\ & + (4-3.244)^2 + (2-3.244)^2 + (3-3.244)^2 + (6-3.244)^2 + (3-3.244)^2 + \\ & + (2-3.244)^2 + (6-3.244)^2 + (3-3.244)^2 + (2-3.244)^2 + (3-3.244)^2 + \\ & + (0-3.244)^2 + (1-3.244)^2 + (3-3.244)^2 + (1-3.244)^2 + (2-3.244)^2 = 156.311 \end{aligned}$$

			B: Month			
			$b_1$ : Month 1	$b_2$ : Month 2	$b_3$ : Month 3	Case Total
A: Type of Novel	$a_1$ : Science Fiction	$S_1$	1	3	6	$S_1 = 10$
		$S_2$	1	4	8	$S_2 = 13$
		$S_3$	3	3	6	$S_3 = 12$
		$S_4$	5	5	7	$S_4 = 17$
		$S_5$	2	4	5	$S_5 = 11$
			$a_1 b_1 = 12$	$a_1 b_2 = 19$	$a_1 b_3 = 32$	$a_1 = 63$
	$a_2$ : Mystery	$S_6$	3	1	0	$S_6 = 4$
		$S_7$	4	4	2	$S_7 = 10$
		$S_8$	5	3	2	$S_8 = 10$
		$S_9$	4	2	0	$S_9 = 6$
		$S_{10}$	4	5	3	$S_{10} = 12$
			$a_2 b_1 = 20$	$a_2 b_2 = 15$	$a_2 b_3 = 7$	$a_2 = 42$
	$a_3$ : Romance	$S_{11}$	4	2	0	$S_{11} = 6$
		$S_{12}$	2	6	1	$S_{12} = 9$
		$S_{13}$	3	3	3	$S_{13} = 9$
		$S_{14}$	6	2	1	$S_{14} = 9$
		$S_{15}$	3	3	2	$S_{15} = 8$
			$a_3 b_1 = 18$	$a_3 b_2 = 16$	$a_3 b_3 = 7$	$a_3 = 41$
			$b_1 = 50$	$b_2 = 50$	$b_3 = 46$	Total = 146

# The *simplified* calculation formula

$$SS_A = \frac{\sum A^2}{bs} - \frac{T^2}{abs} = \frac{63^2 + 42^2 + 41^2}{3(5)} - \frac{146^2}{3(3)(5)}$$

$$SS_{S/A} = \frac{\sum (AS)^2}{b} - \frac{\sum A^2}{bs} = \frac{10^2 + 13^2 + 12^2 + \dots + 8^2}{3} - \frac{63^2 + 42^2 + 41^2}{3(5)}$$

$$SS_B = \frac{\sum B^2}{as} - \frac{T^2}{abs} = \frac{50^2 + 50^2 + 46^2}{3(5)} - \frac{146^2}{3(3)(5)}$$

$$SS_{AB} = \frac{\sum (AB)^2}{s} - \frac{\sum A^2}{bs} - \frac{\sum B^2}{as} + \frac{T^2}{abs} = \frac{12^2 + 19^2 + 32^2 + 20^2 + 15^2 + 7^2 + 18^2 + 16^2 + 7^2}{5} - \frac{63^2 + 42^2 + 41^2}{3(5)} - \frac{50^2 + 50^2 + 46^2}{3(5)} + \frac{146^2}{3(3)(5)}$$

$$SS_{B \times S/A} = \sum Y^2 - \frac{\sum (AB)^2}{s} - \frac{\sum (AS)^2}{b} + \frac{\sum A^2}{bs} = 1^2 + 1^2 + 3^2 + 5^2 + 2^2 + 3^2 + 4^2 + 5^2 + \dots + 1^2 + 2^2 - \frac{12^2 + 19^2 + 32^2 + 20^2 + 15^2 + 7^2 + 18^2 + 16^2 + 7^2}{5} - \frac{10^2 + 13^2 + \dots + 8^2}{3} + \frac{63^2 + 42^2 + 41^2}{3(5)}$$

$$SS_T = \sum Y^2 - \frac{T^2}{abs} = 1^2 + 1^2 + 3^2 + 5^2 + 2^2 + 3^2 + \dots + 1^2 + 2^2 - \frac{146^2}{3(3)(5)}$$



$$SS_A = 494.27 - 473.69 = 20.58$$

$$SS_{S/A} = 520.67 - 494.27 = 26.40$$

$$SS_B = 474.40 - 473.69 = 0.71$$

$$SS_{AB} = 566.40 - 494.27 - 474.40 + 473.69 = 71.42$$

$$SS_{B \times S/A} = 630 - 566.40 - 520.67 + 494.27 = 37.20$$

$$SS_T = 630 - 473.69 = 156.31$$

$$df_A = a - 1 = 3 - 1 = 2$$

$$df_{S/A} = a(s - 1) = 3(5 - 1) = 12$$

$$df_B = b - 1 = 3 - 1 = 2$$

$$df_{AB} = (a - 1)(b - 1) = (3 - 1)(3 - 1) = 4$$

$$df_{B \times S / A} = a(b - 1)(s - 1) = 3(3 - 1)(5 - 1) = 24$$

$$df_T = abs - 1 = N - 1 = 3(3)(5) - 1 = 44$$

# Results – ANOVA summary table

Source	SS	df	MS	<i>F</i>
Randomized Groups				
<i>A</i>	20.58	2	10.29	$\frac{10.29}{2.20} = 4.68$
<i>S/A</i>	26.40	12	2.20	
Repeated Measures				
<i>B</i>	0.71	2	0.36	$\frac{0.36}{1.55} = 0.23$
<i>A × B</i>	71.42	4	17.86	$\frac{17.86}{1.55} = 11.52$
<i>B × S/A</i>	37.20	24	1.55	
<i>T</i>	156.31	44		

# Post-hoc tests after obtaining significant effects

## **Between Groups**

- If you have a significant BG main effect(s) they need to be broken down to find which levels are different
- The comparisons are done the same way as complete BG comparisons (remember one-way or two-way independent-measures ANOVA?)

## **Within Groups** comparisons

- If a WG main effect is significant, it also needs to be followed by comparisons
- In the same way as in one-way repeated-measures ANOVA

# Assumptions

- **Normality** of the distribution of residuals:
  - Applies to each condition (cell)
  - Possible outliers might lead to violations
  - Practically, only requires approximately normal
  - Shapiro-Wilk test (or Q-Q plot)
- **Homogeneity** of Variance
  - Each group (combination or cell) has the same variance
  - Should hold for each level of the within-group factor
  - Levene's F test

# Assumptions

- **Sphericity** for within-group differences
  - Variance for the difference score between within-group conditions (factor B in our example) should be equal.
  - This is also an assumption for any repeated-measure ANOVA, including the one-way repeated-measure ANOVA we covered before.
  - This time, this equal variance of difference score is only tested for the repeated measures of the design, including interaction terms with repeated measures.
  - Using Mauchly's sphericity, as we covered before.

# R outputs for the example question

Levene's Test for Homogeneity of Variance:

	Levene's F	df1	df2	p
DV: time1	1.848	2	12	.200
DV: time2	0.681	2	12	.525
DV: time3	0.321	2	12	.731

Mauchly's Test of Sphericity:

	Mauchly's W	p
time	0.7890	.272
group x time	0.7890	.272

	MS	MSE	df1	df2	F	p		$\eta^2p$ [90% CI of $\eta^2p$ ]	$\eta^2G$
group	10.289	2.200	2.000	12.000	4.677	.031	*	.438 [.034, .659]	.244
time	0.431	1.877	1.651	19.818	0.229	.755		.019 [.000, .152]	.011
group x time	21.624	1.877	3.303	19.818	11.520	<.001	***	.658 [.388, .775]	.529

# What if the interaction is significant?

- **Only** reporting significant main effect is misleading (remember we cannot take the face value of main effect?).
- We will need to look at the profile plot and interpret the difference between groups (between-subject factor) accordingly.
- Use **simple main effect** to check group difference on each level (i.e., each month here).



# Post-hoc tests for main effect

*Note here main effect should be interpreted with caution since the interaction is significant.*

† Pairwise Comparisons of "group":

Contrast	Estimate	S.E.	df	t	p		Cohen's d [95% CI of d]
group2 - group1	-1.400	(0.542)	12	-2.585	.072	.	-0.859 [-1.782, 0.065]
group3 - group1	-1.467	(0.542)	12	-2.708	.057	.	-0.900 [-1.823, 0.024]
group3 - group2	-0.067	(0.542)	12	-0.123	1.000		-0.041 [-0.964, 0.883]

Pooled SD for computing Cohen's d: 1.630

Results are averaged over the levels of: time

P-value adjustment: Bonferroni method for 3 tests.

# Simple main effect on **time**

----- EMMEANS (effect = "time") -----

Joint Tests of "time":

Effect	group	df1	df2	F	p	$\eta^2p$ [90% CI of $\eta^2p$ ]
time	group1	2	12	18.219	<.001 ***	.752 [.465, .857]
time	group2	2	12	7.356	.008 **	.551 [.148, .734]
time	group3	2	12	6.941	.010 **	.536 [.131, .725]

## Post-hoc tests for simple main effect on **time**

Pairwise Comparisons of "time":

Contrast	group	Estimate	S.E.	df	t	p	Cohen's d [95% CI of d]
time2 - time1	group1	1.400	(0.917)	12	1.528	.458	0.859 [-0.704, 2.422]
time3 - time1	group1	4.000	(0.816)	12	4.899	.001 **	2.454 [ 1.062, 3.846]
time3 - time2	group1	2.600	(0.594)	12	4.374	.003 **	1.595 [ 0.581, 2.609]
time2 - time1	group2	-1.000	(0.917)	12	-1.091	.890	-0.613 [-2.176, 0.949]
time3 - time1	group2	-2.600	(0.816)	12	-3.184	.024 *	-1.595 [-2.987, -0.203]
time3 - time2	group2	-1.600	(0.594)	12	-2.692	.059 .	-0.982 [-1.995, 0.032]
time2 - time1	group3	-0.400	(0.917)	12	-0.436	1.000	-0.245 [-1.808, 1.317]
time3 - time1	group3	-2.200	(0.816)	12	-2.694	.059 .	-1.350 [-2.742, 0.043]
time3 - time2	group3	-1.800	(0.594)	12	-3.028	.032 *	-1.104 [-2.118, -0.091]

# Simple main effect on **group**

----- EMMEANS (effect = "group") -----

Joint Tests of "group":

Effect	time	df1	df2	F	p	$\eta^2p$ [90% CI of $\eta^2p$ ]
group	time1	2	12	1.857	.198	.236 [.000, .500]
group	time2	2	12	0.441	.654	.068 [.000, .291]
group	time3	2	12	28.409	<.001 ***	.826 [.612, .900]

# Post-hoc tests for simple main effect on **group**

Pairwise Comparisons of "group":

Contrast	time	Estimate	S.E.	df	t	p	Cohen's d [95% CI of d]
group2 - group1	time1	1.600	(0.864)	12	1.852	.266	0.982 [-0.492, 2.455]
group3 - group1	time1	1.200	(0.864)	12	1.389	.570	0.736 [-0.737, 2.210]
group3 - group2	time1	-0.400	(0.864)	12	-0.463	1.000	-0.245 [-1.719, 1.228]
group2 - group1	time2	-0.800	(0.887)	12	-0.902	1.000	-0.491 [-2.003, 1.022]
group3 - group1	time2	-0.600	(0.887)	12	-0.676	1.000	-0.368 [-1.880, 1.144]
group3 - group2	time2	0.200	(0.887)	12	0.225	1.000	0.123 [-1.390, 1.635]
group2 - group1	time3	-5.000	(0.766)	12	-6.528	<.001 ***	-3.067 [-4.373, -1.761]
group3 - group1	time3	-5.000	(0.766)	12	-6.528	<.001 ***	-3.067 [-4.373, -1.761]
group3 - group2	time3	-0.000	(0.766)	12	-0.000	1.000	-0.000 [-1.306, 1.306]

# Higher order mixed designs

between within

Design	Randomized-Groups IVs	Repeated-Measures IVs	Sources of Variability		
			Randomized Groups	Repeated Measures	Error Terms
Two-way mixed	A	B	A		$S/A$
				$B, A \times B$	$B \times S/A$
Three-way mixed	A, B	C	A, B, $A \times B$		$S/AB$
				$C, A \times C, B \times C, A \times B \times C$	$C \times S/AB$
	A	B, C	A		$S/A$
				$B, A \times B$	$B \times S/A$
				$C, A \times C$	$C \times S/A$
				$B \times C, A \times B \times C$	$B \times C \times S/A$
Four-way mixed	A, B	C, D	A, B, $A \times B$		$S/AB$
				$C, A \times C, B \times C, A \times B \times C$	$C \times S/AB$
				$D, A \times D, B \times D, A \times B \times D$	$D \times S/AB$
				$C \times D, A \times C \times D, B \times C \times D, A \times B \times C \times D$	$C \times D \times S/AB$

# Another example for R coding

	no	group	day1	day2	day3	day4	day5
	1	1	26	20	18	11	10
	2	1	34	35	29	22	23
	3	1	41	37	25	18	15
	4	1	29	28	22	15	13
	5	1	35	34	27	21	17
	6	1	28	22	17	14	10
	7	1	38	34	28	25	22
	8	1	43	37	30	27	25
	9	2	42	38	26	20	15
	10	2	31	27	21	18	13
	11	2	45	40	33	25	18
	12	2	29	25	17	13	8
	13	2	29	32	28	22	18
	14	2	33	30	24	18	7
	15	2	34	30	25	24	23
	16	2	37	31	25	22	20

**How learning a memory technique improves memory:**

- Two memory techniques are compared
- 16 subjects, divided into two groups
- Measured 5 times in 5 days