

Lecture 12: Mixed-Design ANOVA

Combining between-group & within-subject

In the simplest 2×2 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_1	S_1
	S_2	S_2
	S_3	S_3
	S_4	S_4
	S_5	S_5
Princeton	S_6	S_6
	S_7	S_7
	S_8	S_8
	S_9	S_9
	S_{10}	S_{10}

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

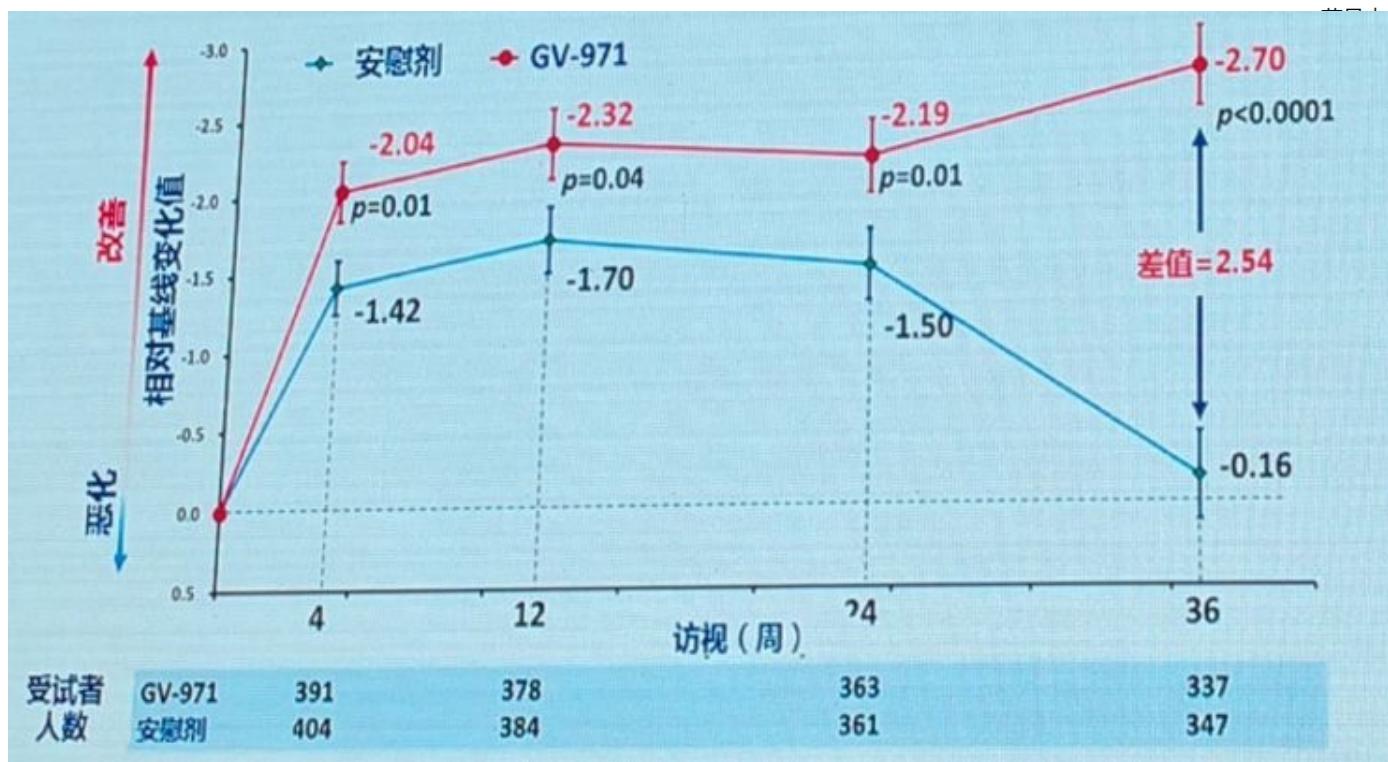
批准日期 2019-11-02

原批准文号

药品本位码 86900722000145

药品广告

居库查询 中药保护品种库



Analysis

		b_1	b_2	b_3
Randomized Groups	a_1	S_1	S_1	S_1
	a_1	S_2	S_2	S_2
	a_1	S_3	S_3	S_3
	a_1	S_4	S_4	S_4
	a_1	S_5	S_5	S_5
	a_2	S_6	S_6	S_6
	a_2	S_7	S_7	S_7
	a_2	S_8	S_8	S_8
	a_2	S_9	S_9	S_9
	a_2	S_{10}	S_{10}	S_{10}
	a_3	S_{11}	S_{11}	S_{11}
	a_3	S_{12}	S_{12}	S_{12}
	a_3	S_{13}	S_{13}	S_{13}
	a_3	S_{14}	S_{14}	S_{14}
	a_3	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)
- $B = b - 1$
- $AB = (a - 1)(b - 1)$
- $BxS/A = a(b - 1)(s - 1)$
- $T = abs - 1$ or $N - 1$

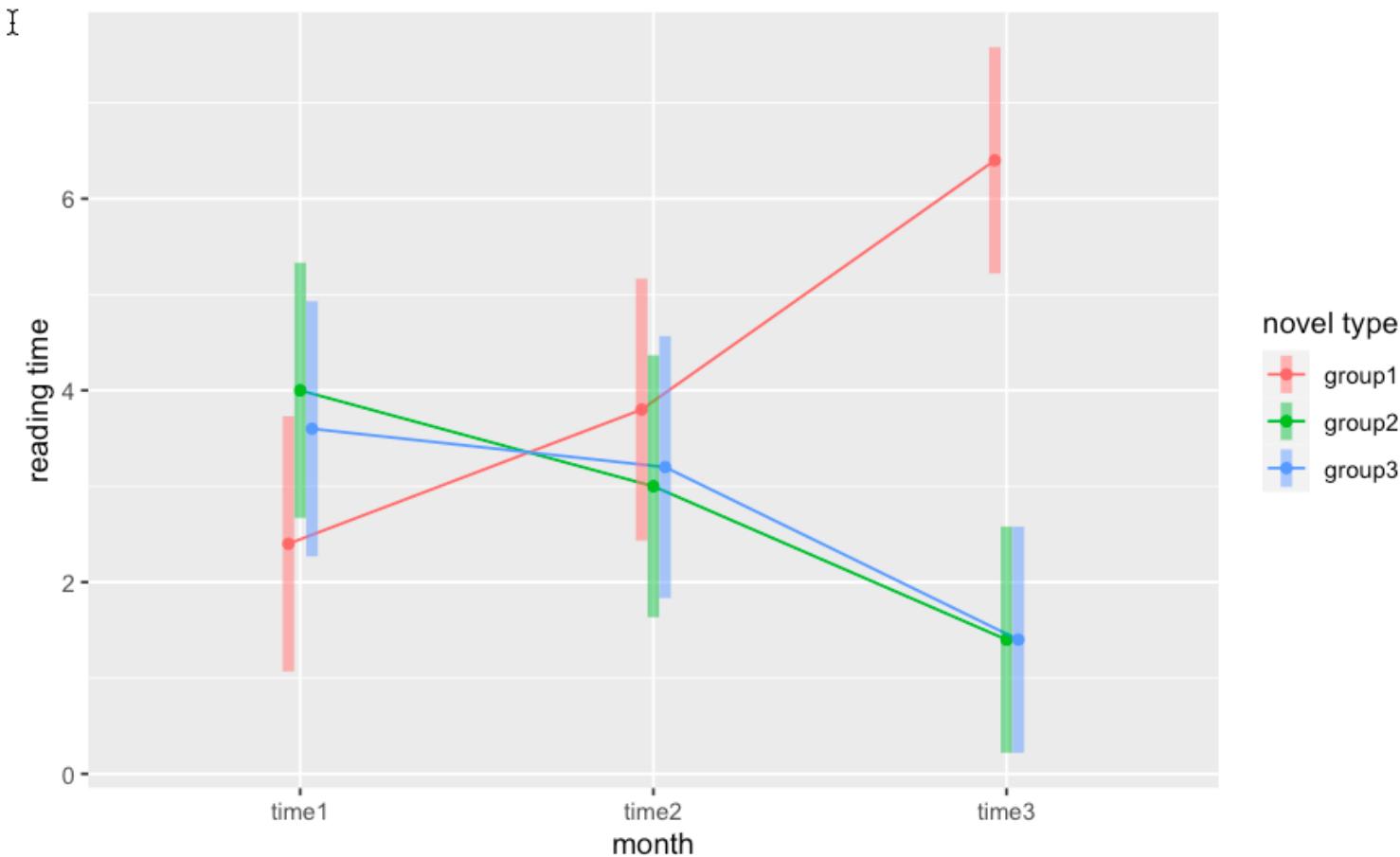
Example – Books by Month

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: Month			Case Means		
		b_1 : Month 1	b_2 : Month 2	b_3 : Month 3			
A: Type of Novel	: Science Fiction	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	$S_5 = 3.667$	
			$a_1 b_1 = 2.4$	$a_1 b_2 = 3.8$	$a_1 b_3 = 6.4$	$a_1 = 4.2$	
	$a_2 : Mystery$	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	$S_{10} = 4$	
			$a_2 b_1 = 4$	$a_2 b_2 = 3$	$a_2 b_3 = 1.4$	$a_2 = 2.8$	
	$a_3 : Romance$	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

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

$$\sum (Y_{ijk} - \bar{Y}_{...})^2 = \sum n_j (\bar{Y}_{.j.} - \bar{Y}_{...})^2 + \sum n_k (\bar{Y}_{..k} - \bar{Y}_{...})^2 +$$

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

$$+ k \sum (\bar{Y}_{i..} - \bar{Y}_{.j.})^2 + \left[\sum (Y_{ijk} - \bar{Y}_{.jk})^2 - k \sum (\bar{Y}_{i..} - \bar{Y}_{.j.})^2 \right]$$

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

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

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

$$\sum n_{jk} (\bar{Y}_{.jk} - \bar{Y}_{...})^2 = 5 * [(2.4 - 3.244)^2 + (3.8 - 3.244)^2 + (6.4 - 3.244)^2 +$$

$$+ (4 - 3.244)^2 + (3 - 3.244)^2 + (1.4 - 3.244)^2 +$$

$$+ (3.6 - 3.244)^2 + (3.2 - 3.244)^2 + (1.4 - 3.244)^2] = 92.711$$

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

This k is necessary; 3 months.

The deviation of each subject from his group mean

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

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			Case Total	
		b ₁ : Month 1	b ₂ : Month 2	b ₃ : Month 3		
A: Type of Novel	a ₁ : Science Fiction	S ₁	1	3	6	S₁ = 10
		S ₂	1	4	8	S₂ = 13
		S ₃	3	3	6	S₃ = 12
		S ₄	5	5	7	S₄ = 17
		S ₅	2	4	5	S₅ = 11
			a₁b₁ = 12	a₁b₂ = 19	a₁b₃ = 32	a₁ = 63
	a ₂ : Mystery	S ₆	3	1	0	S₆ = 4
		S ₇	4	4	2	S₇ = 10
		S ₈	5	3	2	S₈ = 10
		S ₉	4	2	0	S₉ = 6
		S ₁₀	4	5	3	S₁₀ = 12
			a₂b₁ = 20	a₂b₂ = 15	a₂b₃ = 7	a₂ = 42
	a ₃ : Romance	S ₁₁	4	2	0	S₁₁ = 6
		S ₁₂	2	6	1	S₁₂ = 9
		S ₁₃	3	3	3	S₁₃ = 9
		S ₁₄	6	2	1	S₁₄ = 9
		S ₁₅	3	3	2	S₁₅ = 8
			a₃b₁ = 18	a₃b₂ = 16	a₃b₃ = 7	a₃ = 41
			b₁ = 50	b₂ = 50	b₃ = 46	Total = 146

The *simplified* calculation formula

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

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

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

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

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

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

$$\begin{aligned} \text{SS}_A &= 494.27 - 473.69 & = & 20.58 \\ \text{SS}_{S/A} &= 520.67 - 494.27 & = & 26.40 \\ \text{SS}_B &= 474.40 - 473.69 & = & 0.71 \\ \text{SS}_{AB} &= 566.40 - 494.27 - 474.40 + 473.69 & = & 71.42 \\ \text{SS}_{B \times S/A} &= 630 - 566.40 - 520.67 + 494.27 & = & 37.20 \\ \text{SS}_T &= 630 - 473.69 & = & 156.31 \end{aligned}$$

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

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

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

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

$$df_{BxS/A} = a(b - 1)(s - 1) = 3(3 - 1)(5 - 1) = 24$$

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

Results – ANOVA summary table

Source	SS	df	MS	F
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</i> \times <i>B</i>	71.42	4	17.86	$\frac{17.86}{1.55} = 11.52$
<i>B</i> \times <i>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	η^2 p	[90% CI of η^2 p]	η^2 G
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	η^2	p [90% CI of η^2 p]
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	$\eta^2 p$ [90% CI of $\eta^2 p$]
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 × B	B × S/A		
Three-way mixed	A, B	C	A, B, A × B		S/AB		
				C, A × C, B × C, A × B × C	C × S/AB		
	A	B, C	A		S/A		
				B, A × B	B × S/A		
Four-way mixed	A, B	C, D	C, A × C		C × S/A		
				B × C, A × B × C	B × C × S/A		
			A, B, A × B		S/AB		
				C, A × C, B × C, A × B × C	C × S/AB		
				D, A × D, B × D, A × B × D	D × S/AB		
				C × D, A × C × D, B × C × D, A × B × C × D	C × D × 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