

# 相關樣本/配對資料 的統計推論

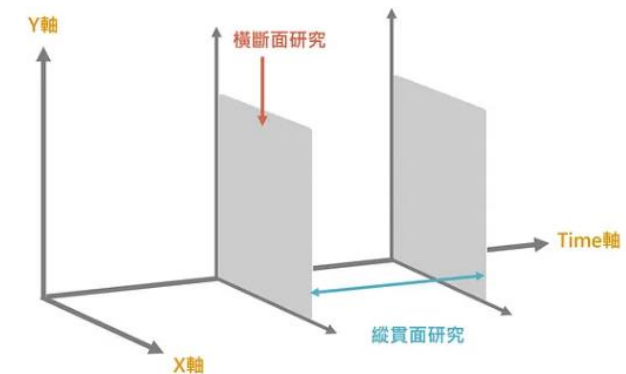
醫學研究部生統小組

陳俊朋

日期: 2024/6/12

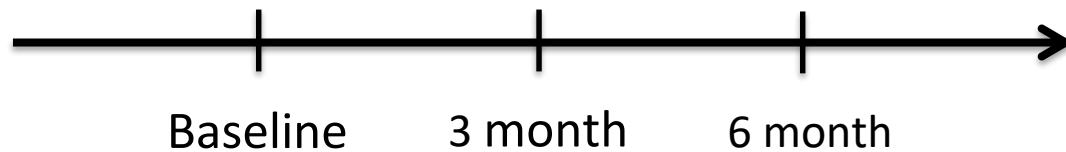
# 研究設計

| Cross-sectional                                   | Longitudinal                                                      |
|---------------------------------------------------|-------------------------------------------------------------------|
| Single point in time                              | Period of time                                                    |
| Different samples                                 | Same sample                                                       |
| Provides snapshot of society at a given point     | Follows changes in participants over time                         |
| Compare many different variables at the same time | The difference between these studies is the timeline and variable |
| Cheaper                                           | Expensive and require more resources                              |

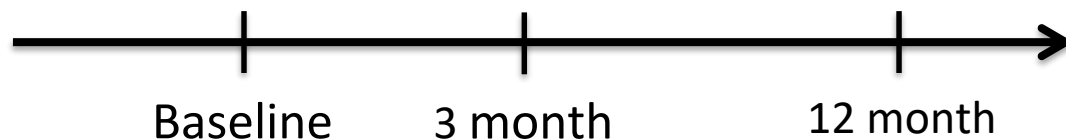


# 重複測量使用時機

- 相同受試者在不同時間點進行多次測量
  - 間隔時間相同



- 間隔時間不同



# 重複測量優缺點

- 優點

- 需求樣本數較少
- 不同時間點變化
- 不同組別，隨著時間的變化

- 缺點

- 資料完整性不佳
- 個案須定期追蹤
- 某些因素可能影響資料

# 重複測量統計

- 單組重複測量
  - 同一組內的個案，在多個時間點測量
- 多組重複測量
  - 將個案分組後，依照各組分類在多個時間點測量

# 重複測量統計方法

## 重複2次

- 類別變項
  - McNemar test (2\*2 table)
  - McNemar-Bowker test (3\*3 table)
- 連續變項
  - Paired t test
  - Wilcoxon signed-rank test
  - ANCOVA

## 重複3次以上

- 類別變項
  - Cochran's Q test
- 連續變項
  - Friedman test
  - Repeated measures ANOVA
  - Linear Mixed Model (LMM)
  - Generalized Estimating Equation (GEE)

# 重複測量建檔

- 寬資料

| no  | Group | Sex | MMSE_1 | MMSE_2 | MMSE_3 |
|-----|-------|-----|--------|--------|--------|
| N01 | 0     | 0   | 30     | 28     | 30     |
| N02 | 0     | 1   | 11     | 15     | 14     |
| N03 | 0     | 1   | 26     | 28     | 26     |
| N04 | 0     | 0   | 22     | 21     | 22     |
| N05 | 0     | 0   | 27     | 26     | 25     |

- 長資料

| no  | Group | Sex | visit | MMSE |
|-----|-------|-----|-------|------|
| N01 | 0     | 0   | 1     | 30   |
| N01 | 0     | 0   | 2     | 28   |
| N01 | 0     | 0   | 3     | 30   |
| N02 | 0     | 1   | 1     | 11   |
| N02 | 0     | 1   | 2     | 15   |
| N02 | 0     | 1   | 3     | 14   |
| N03 | 0     | 1   | 1     | 26   |
| N03 | 0     | 1   | 2     | 28   |
| N03 | 0     | 1   | 3     | 26   |
| N04 | 0     | 0   | 1     | 22   |
| N04 | 0     | 0   | 2     | 21   |
| N04 | 0     | 0   | 3     | 22   |
| N05 | 0     | 0   | 1     | 27   |
| N05 | 0     | 0   | 2     | 26   |
| N05 | 0     | 0   | 3     | 25   |

# McNemar test

- 僅適用2\*2 table
- 單組同一人前後測/兩種不同診斷工具

International Urogynecology Journal  
<https://doi.org/10.1007/s00192-021-04977-7>

ORIGINAL ARTICLE



## Transvaginal repair of anterior vaginal wall prolapse with polyvinylidene fluoride (PVDF) mesh: an alternative for previously restricted materials?

Mohammad-Javad Eslami<sup>1</sup> • Mahtab Zargham<sup>1</sup> • Farshad Gholipour<sup>2</sup> • Mohammadreza Hajian<sup>3</sup> • Katayoun Bakhtiari<sup>4</sup> • Sakineh Hajebrahami<sup>5</sup> • Melina Eghbal<sup>6</sup> • Ziba Farajzadegan<sup>7</sup>

**Table 3** Urinary and vaginal symptoms at baseline and 12-month follow-up

|                                                 | Preoperative | Postoperative<br>(12 months) | <i>P</i> value |
|-------------------------------------------------|--------------|------------------------------|----------------|
| Storage symptoms ( <i>N</i> =108)               |              |                              |                |
| SUI: <i>n</i> (%)                               | 74 (68.5%)   | 7 (6.5%)                     | <0.0001 *      |
| UUI: <i>n</i> (%)                               | 33 (30.5%)   | 9 (8.3%)                     | <0.0001 *      |
| MUI: <i>n</i> (%)                               | 29 (26.9%)   | 3 (2.8%)                     | <0.0001 *      |
| Other storage symptoms                          | 30 (27.8%)   | 8 (7.4%)                     | <0.0001 *      |
| Voiding symptoms: <i>n</i> (%) ( <i>N</i> =108) | 24 (22.2%)   | 13 (12%)                     | 0.019 *        |
| ICIQ-UI (mean±SD) ( <i>N</i> =98)               | 8.79±6.05    | 0.77±1.87                    | <0.0001**      |
| ICIQ-VS (mean±SD) ( <i>N</i> =41)               |              |                              |                |
| Vaginal symptoms                                | 43.31±6.13   | 7.49±3.75                    | <0.0001**      |
| Sexual symptoms                                 | 43.17±6.15   | 7.82±4.89                    | <0.0001**      |
| Quality of life                                 | 8.27±0.98    | 1.6±1.73                     | <0.0001**      |

\* McNemar's test

\*\* Wilcoxon signed-rank test

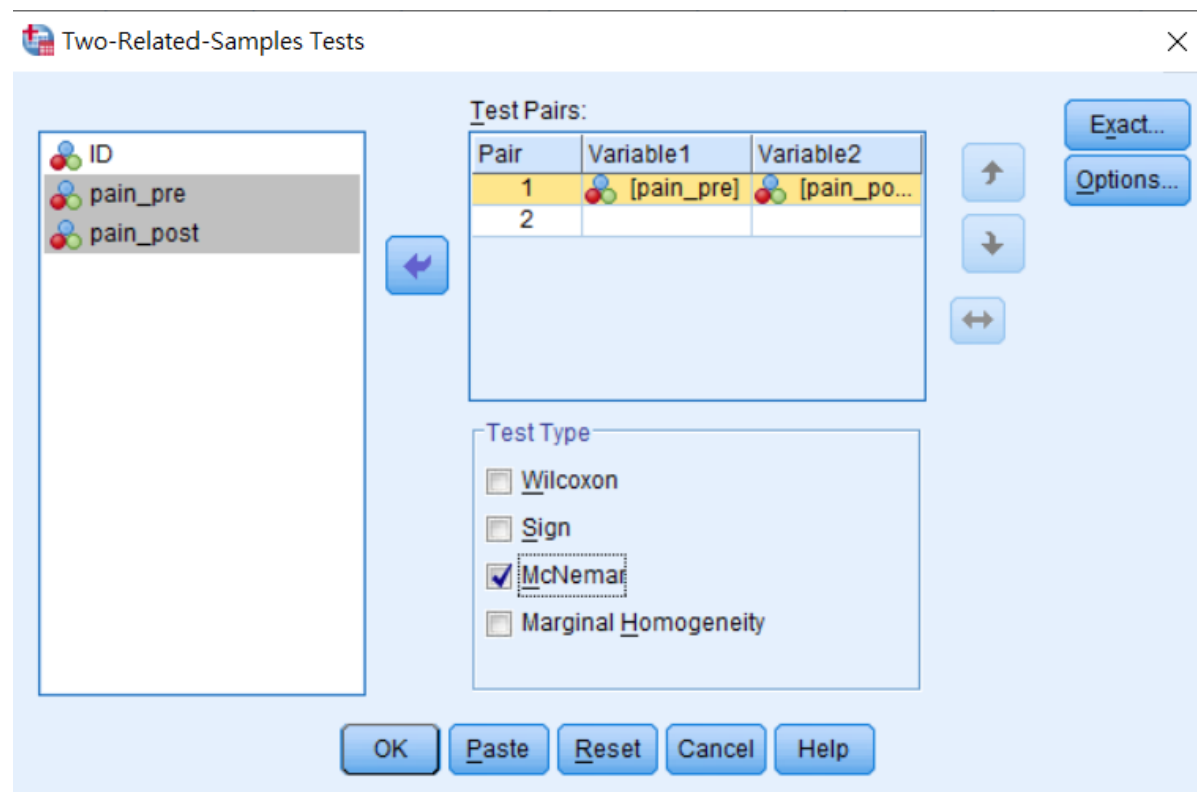
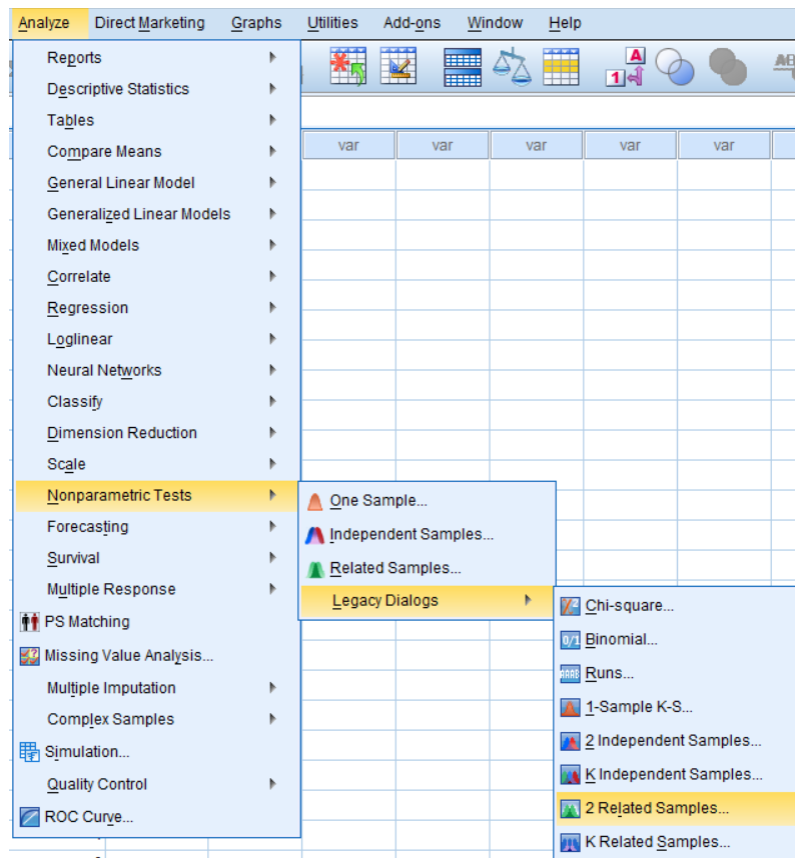
# McNemar test dataset

- 術前和術後有無疼痛比較

| ID | pain_pre | pain_post |
|----|----------|-----------|
| 1  | 1        | 0         |
| 2  | 0        | 0         |
| 3  | 0        | 0         |
| 4  | 0        | 0         |
| 5  | 0        | 0         |
| 6  | 1        | 0         |
| 7  | 0        | 0         |
| 8  | 0        | 0         |
| 9  | 0        | 0         |
| 10 | 0        | 0         |
| 11 | 0        | 0         |
| 12 | 1        | 0         |
| 13 | 0        | 0         |
| 14 | 0        | 0         |
| 15 | 0        | 0         |

# McNemar test analysis

分析>無母數檢定>歷史對話記錄>2個相關樣本



# McNemar test output

## McNemar Test

### Crosstabs

**pain\_pre & pain\_post**

| pain_pre | pain_post |   |
|----------|-----------|---|
|          | 0         | 1 |
| 0        | 21        | 0 |
| 1        | 5         | 1 |

**Test Statistics<sup>a</sup>**

|                       | pain_pre & pain_post |
|-----------------------|----------------------|
| N                     | 27                   |
| Exact Sig. (2-tailed) | .063 <sup>b</sup>    |

a. McNemar Test

b. Binomial distribution used.

**pain\_post \* pain\_pre Crosstabulation**

|             |            | pain_pre |       | Total  |
|-------------|------------|----------|-------|--------|
|             |            | 0        | 1     |        |
| pain_post 0 | Count      | 21       | 5     | 26     |
|             | % of Total | 77.8%    | 18.5% | 96.3%  |
| 1           | Count      | 0        | 1     | 1      |
|             | % of Total | 0.0%     | 3.7%  | 3.7%   |
| Total       | Count      | 21       | 6     | 27     |
|             | % of Total | 77.8%    | 22.2% | 100.0% |

Table.

|          | Pre-OP    | Post-OP 1y | <i>p</i> value |
|----------|-----------|------------|----------------|
| Symptoms |           |            |                |
| pain     | 6 (22.2%) | 1 (3.7%)   | 0.063          |

McNemar test. \* $p < 0.05$ , \*\* $p < 0.01$

# McNemar-Bowker test

- 適用3\*3 table以上
- 單組同一人前後測/兩種不同診斷工具

## Research Article

### Changes in Food Security Status During Undergraduate Enrollment

Cara L. Cuite, PhD<sup>1</sup>; Kelly E. Dietz, MPP<sup>2</sup>; La Reina J. Bates, PhD<sup>3</sup>; Stephanie A. Brescia, PhD<sup>4</sup>



**Table 2.** 2016 FSS by 2019 FSS Among Undergraduates (n = 338)

| 2016 FSS <sup>a</sup> | 2019 FSS <sup>b</sup> |           |           |           | Total      |
|-----------------------|-----------------------|-----------|-----------|-----------|------------|
|                       | High                  | Marginal  | Low       | Very Low  |            |
| High                  | 106 (31.4)            | 36 (10.7) | 38 (11.2) | 18 (5.3)  | 198 (58.6) |
| Marginal              | 21 (6.2)              | 16 (4.7)  | 16 (4.7)  | 11 (3.3)  | 64 (18.9)  |
| Low                   | 9 (2.7)               | 9 (2.7)   | 9 (2.7)   | 13 (3.8)  | 40 (11.8)  |
| Very low              | 1 (0.3)               | 1 (0.3)   | 17 (5.0)  | 17 (5.0)  | 36 (10.6)  |
| Total                 | 137 (40.4)            | 62 (18.3) | 80 (23.6) | 59 (17.4) | 338 (100)  |

FSS indicates food security status.

<sup>a</sup>Measured using the 10-item Food Security Survey Module; <sup>b</sup>Measured using the 18-item Food Security Survey Module.

Note: Values displayed as n (% of total). Statistical test conducted is the McNemar-Bowker test ( $\chi^2 = 47.878$ , degrees of freedom = 6,  $P < 0.001$ ) and paired samples Wilcoxon signed rank test ( $P < 0.001$ ). Critical  $P \leq 0.05$ .

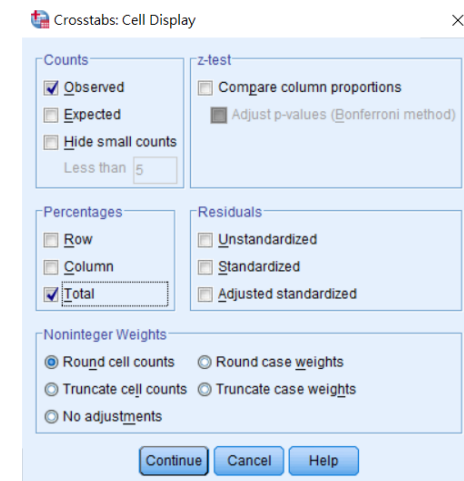
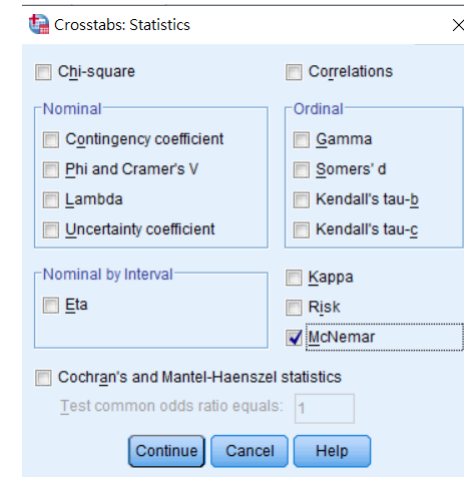
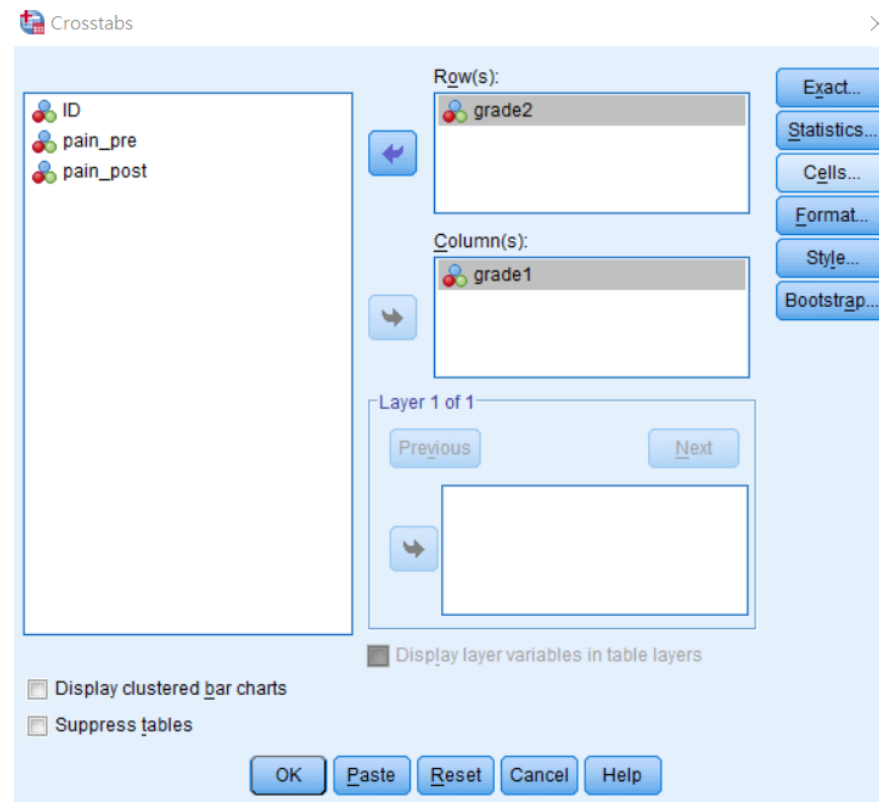
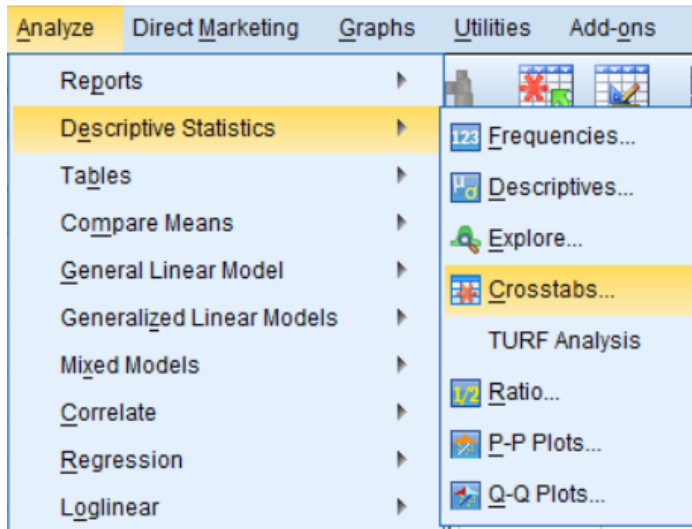
# McNemar-Bowker test dataset

- 新舊診斷工具一致性比較

| ID | grade1 | grade2 |
|----|--------|--------|
| 1  | 0      | 1      |
| 2  | 0      | 1      |
| 3  | 0      | 1      |
| 4  | 0      | 2      |
| 5  | 1      | 2      |
| 6  | 1      | 2      |
| 7  | 1      | 1      |
| 8  | 1      | 1      |
| 9  | 1      | 1      |
| 10 | 1      | 1      |
| 11 | 1      | 1      |
| 12 | 2      | 1      |
| 13 | 2      | 1      |
| 14 | 2      | 1      |
| 15 | 2      | 1      |

# McNemar-Bowker test analysis

分析 > 敘述統計 > 交叉表



# McNemar-Bowker test output

grade2 \* grade1 Crosstabulation

|        |            |            | grade1 |       |        | Total |
|--------|------------|------------|--------|-------|--------|-------|
|        |            |            | 0      | 1     | 2      |       |
| grade2 | 0          | Count      | 0      | 2     | 0      | 2     |
|        |            | % of Total | 0.0%   | 7.4%  | 0.0%   | 7.4%  |
|        | 1          | Count      | 7      | 7     | 8      | 22    |
|        |            | % of Total | 25.9%  | 25.9% | 29.6%  | 81.5% |
|        | 2          | Count      | 1      | 2     | 0      | 3     |
|        |            | % of Total | 3.7%   | 7.4%  | 0.0%   | 11.1% |
| Total  | Count      | 8          | 11     | 8     | 27     |       |
|        | % of Total | 29.6%      | 40.7%  | 29.6% | 100.0% |       |

Chi-Square Tests

|                     | Value | df | Asymp. Sig.<br>(2-sided) |
|---------------------|-------|----|--------------------------|
| McNemar-Bowker Test | 7.378 | 3  | .061                     |
| N of Valid Cases    | 27    |    |                          |

Table.

|        | Grade1   |          |          | p value |
|--------|----------|----------|----------|---------|
|        | 0        | 1        | 2        |         |
| Grade2 |          |          |          | 0.061   |
| 0      | 0(0.0%)  | 2(7.4%)  | 0(0.0%)  |         |
| 1      | 7(25.9%) | 7(25.9%) | 8(29.6%) |         |
| 2      | 1(3.7%)  | 2(7.4%)  | 0(0.0%)  |         |

McNemar test. \* $p < 0.05$ , \*\* $p < 0.01$

# Cochran's Q test

- 單組的相同個案，用同一工具測量三次/不同工具(類別型資料屬於二分類)

Open access

Original research

## BMJ Open Implementation of a comprehensive surveillance system for recording suicides and attempted suicides in rural India

Lakshmi Vijayakumar,<sup>1</sup> Soumitra Pathare,<sup>2</sup> Nikhil Jain,<sup>2</sup> Renuka Nardodkar ,<sup>2</sup> Deepa Pandit,<sup>2</sup> Sadhvi Krishnamoorthy,<sup>2</sup> Jasmine Kalha,<sup>2</sup> Laura Shields-Zeeman<sup>3</sup>

**Table 4** Comparison of the ability of community surveillance in obtaining additional data on suicide compared with the hospital and police records

|                        | Non-case | Case | Test statistics                      |
|------------------------|----------|------|--------------------------------------|
| Community surveillance | 15       | 67   | Cochran's<br>Q test 67.9<br>(p<0.01) |
| Hospital records       | 69       | 13   |                                      |
| Police records         | 65       | 17   |                                      |

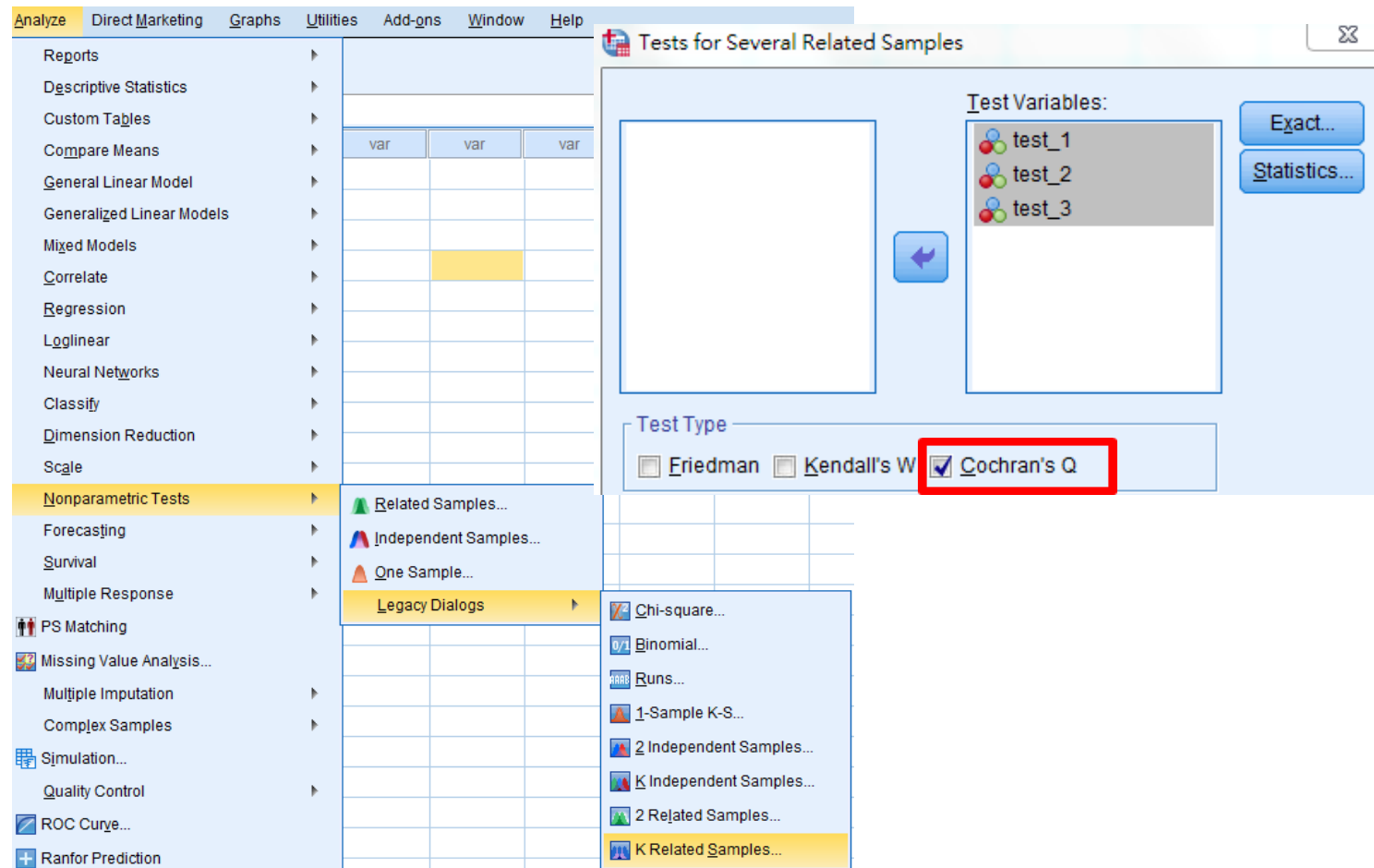
# Cochran Q test dataset

- 同一組個案，測量三次時間點(類別型資料--二分類)
- 0: 無抽菸; 1: 有抽菸

| no | test_1 | test_2 | test_3 |
|----|--------|--------|--------|
| 1  | 0      | 1      | 1      |
| 2  | 0      | 1      | 1      |
| 3  | 1      | 1      | 1      |
| 4  | 1      | 0      | 1      |
| 5  | 0      | 0      | 0      |
| 6  | 1      | 1      | 1      |
| 7  | 1      | 1      | 1      |
| 8  | 1      | 0      | 1      |
| 9  | 1      | 1      | 1      |
| 10 | 0      | 0      | 1      |
| 11 | 1      | 1      | 1      |
| 12 | 1      | 1      | 1      |
| 13 | 1      | 0      | 1      |
| 14 | 1      | 0      | 0      |
| 15 | 0      | 0      | 1      |

# Cochran Q test analysis

分析>無母數檢定>歷史對話記錄>K個相關樣本



# Cochran Q test output

## Cochran Test

### Frequencies

|        | Value |    |
|--------|-------|----|
|        | 0     | 1  |
| test_1 | 5     | 10 |
| test_2 | 7     | 8  |
| test_3 | 2     | 13 |

### Test Statistics

|             |                    |
|-------------|--------------------|
| N           | 15                 |
| Cochran's Q | 4.750 <sup>a</sup> |
| df          | 2                  |
| Asymp. Sig. | .093               |

a. 0 is treated as a success.

Table.

|         | Test1 (n=15) | Test2 (n=15) | Test3 (n=15) | <i>p</i> value |
|---------|--------------|--------------|--------------|----------------|
| Smoking |              |              |              | 0.093          |
| No      | 5 (33.3%)    | 7 (46.7%)    | 2 (13.3%)    |                |
| Yes     | 10 (66.7%)   | 8 (53.3%)    | 13 (86.7%)   |                |

Cochran Q test.

# Paired t test

- 針對常態分布的連續數值或是大樣本的資料做平均值的比較
- 用於前後測比較。



## Subtypes of Premorbid Metabolic Syndrome and Associated Clinical Outcomes in Older Adults

Chu-Sheng Lin<sup>1,2,3,4</sup>, Wei-Ju Lee<sup>4,5,6</sup>, Shih-Yi Lin<sup>2</sup>, Hui-Ping Lin<sup>7</sup>, Ran-Chou Chen<sup>5,7</sup>, Chi-Hung Lin<sup>4,5,8</sup> and Liang-Kung Chen<sup>4,5,9,10\*</sup>

**TABLE 1** | Demographic data of study participants in 2014 and 2016.

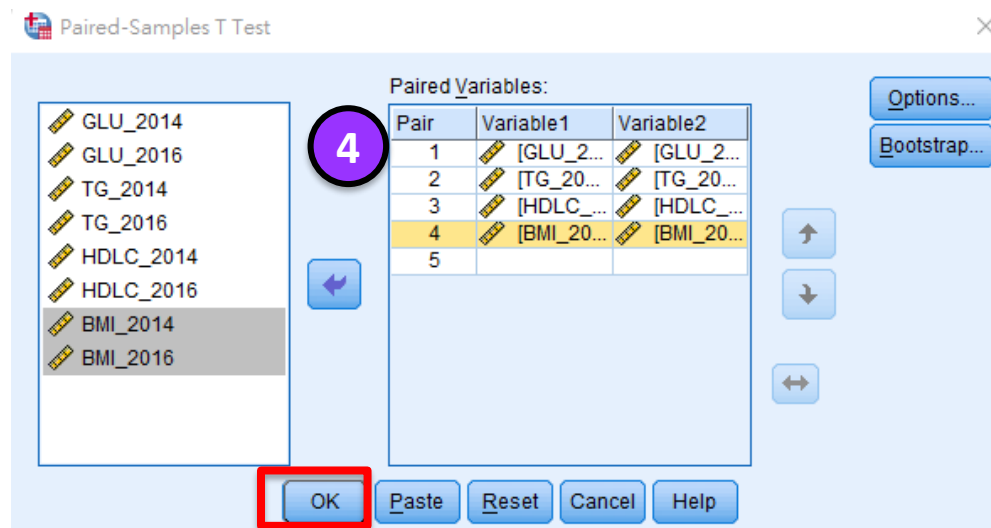
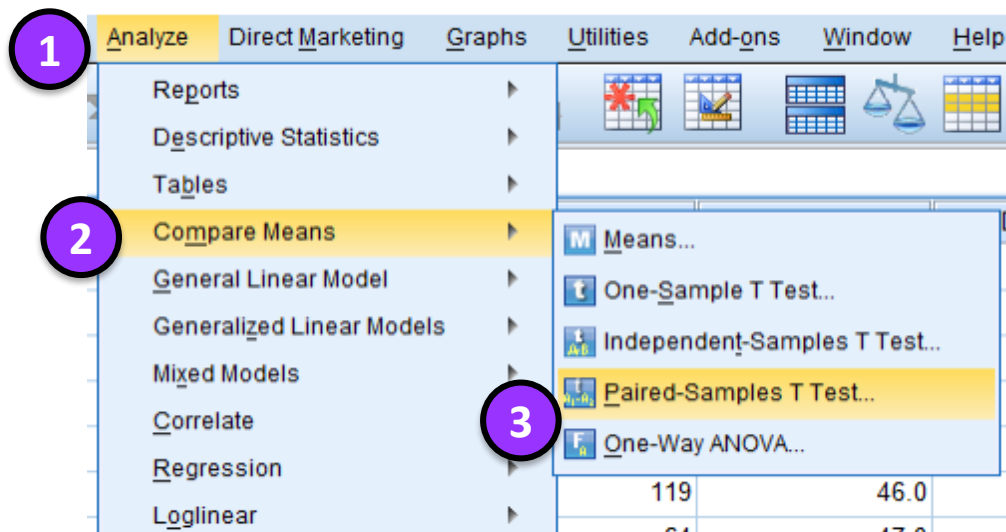
|                                | 2014 (n = 4,537) |         | 2016 (n = 4,537) |         | p-value            |
|--------------------------------|------------------|---------|------------------|---------|--------------------|
|                                | n                | %       | n                | %       |                    |
| Age                            | 71.75            | ±5.93   | 73.75            | ±5.93   | –                  |
| Gender                         |                  |         |                  |         | 1.000              |
| Male                           | 2,207            | (48.6%) | 2,207            | (48.6%) |                    |
| Female                         | 2,330            | (51.4%) | 2,330            | (51.4%) |                    |
| Smoking                        | 309              | (6.8%)  | 297              | (6.5%)  | 0.119              |
| Drinking                       | 547              | (12.1%) | 581              | (12.8%) | 0.862              |
| Exercise                       | 3,237            | (71.3%) | 3,462            | (76.3%) | 0.930              |
| BMI                            | 24.49            | ±3.47   | 24.41            | ±3.42   | <b>0.004**</b>     |
| Height (cm)                    | 157.67           | ±8.15   | 157.59           | ±8.12   | 0.063              |
| Weight (kg)                    | 60.99            | ±10.38  | 60.78            | ±10.47  | <b>&lt;0.001**</b> |
| Waist (cm)                     | 83.20            | ±9.72   | 84.59            | ±9.66   | <b>&lt;0.001**</b> |
| SBP (mmHg)                     | 133.91           | ±18.19  | 134.63           | ±18.52  | <b>0.007**</b>     |
| DBP (mmHg)                     | 78.14            | ±10.91  | 77.21            | ±11.26  | <b>&lt;0.001**</b> |
| Pulse pressure (mmHg)          | 55.77            | ±13.82  | 57.42            | ±13.74  | <b>&lt;0.001**</b> |
| Fasting plasma glucose (mg/dl) | 104.02           | ±21.92  | 104.89           | ±25.17  | <b>0.009**</b>     |
| Triglycerides (mg/dl)          | 117.03           | ±66.71  | 114.85           | ±67.83  | <b>0.024*</b>      |
| HDL cholesterol (mg/dl)        | 55.61            | ±15.66  | 55.60            | ±15.78  | 0.922              |

# Paired t test dataset

| caseno | GLU_2014 | GLU_2016 | TG_2014 | TG_2016 | HDLC_2014 | HDLC_2016 | BMI_2014 | BMI_2016 |
|--------|----------|----------|---------|---------|-----------|-----------|----------|----------|
| 1      | 91       | 91       | 245     | 180     | 34.0      | 42.0      | 27.68    | 27.27    |
| 2      | 103      | 224      | 89      | 104     | 45.0      | 35.0      | 19.15    | 20.16    |
| 3      | 92       | 96       | 96      | 56      | 54.0      | 47.0      | 20.94    | 23.57    |
| 4      | 97       | 96       | 123     | 92      | 57.0      | 57.0      | 25.54    | 25.57    |
| 5      | 88       | 307      | 125     | 106     | 42.0      | 87.0      | 18.97    | 17.10    |
| 6      | 94       | 94       | 101     | 119     | 46.0      | 47.0      | 21.50    | 21.79    |
| 7      | 92       | 91       | 49      | 64      | 47.0      | 50.0      | 25.67    | 26.58    |
| 8      | 151      | 89       | 83      | 69      | 39.0      | 47.0      | 24.61    | 26.37    |
| 9      | 97       | 97       | 120     | 59      | 42.0      | 44.0      | 23.95    | 24.31    |
| 10     | 149      | 156      | 142     | 128     | 50.0      | 50.0      | 25.54    | 23.84    |
| 11     | 113      | 124      | 337     | 104     | 58.0      | 80.0      | 23.28    | 23.74    |
| 12     | 99       | 92       | 114     | 76      | 67.0      | 60.0      | 20.20    | 20.57    |

# Paired t test analysis

分析 > 比較平均數法 > 成對樣本T檢定



# Paired T-test SPSS output

**Paired Samples Statistics**

|        |           | Mean    | N    | Std. Deviation | Std. Error Mean |
|--------|-----------|---------|------|----------------|-----------------|
| Pair 1 | GLU_2014  | 104.02  | 4537 | 21.923         | .325            |
|        | GLU_2016  | 104.89  | 4537 | 25.166         | .374            |
| Pair 2 | TG_2014   | 117.03  | 4537 | 66.712         | .990            |
|        | TG_2016   | 114.85  | 4537 | 67.833         | 1.007           |
| Pair 3 | HDLc_2014 | 55.615  | 4537 | 15.6628        | .2325           |
|        | HDLc_2016 | 55.598  | 4537 | 15.7802        | .2343           |
| Pair 4 | BMI_2014  | 24.4877 | 4537 | 3.46738        | .05148          |
|        | BMI_2016  | 24.4141 | 4537 | 3.41873        | .05076          |

在Glucose的部分，  
2014和2016有統計差異  
2016平均高於2014 ( $p=0.009$ )

**Paired Samples Test**

|        |                          | Paired Differences |                |                    |                                              |        | t      | df   | Sig. (2-tailed) |
|--------|--------------------------|--------------------|----------------|--------------------|----------------------------------------------|--------|--------|------|-----------------|
|        |                          | Mean               | Std. Deviation | Std. Error<br>Mean | 95% Confidence Interval of the<br>Difference |        |        |      |                 |
|        |                          |                    |                |                    | Lower                                        | Upper  |        |      |                 |
| Pair 1 | GLU_2014 - GLU_2016      | -.869              | 22.503         | .334               | -1.524                                       | -.214  | -2.600 | 4536 | .009            |
| Pair 2 | TG_2014 - TG_2016        | 2.186              | 65.013         | .965               | .294                                         | 4.078  | 2.265  | 4536 | .024            |
| Pair 3 | HDLC_2014 -<br>HDLC_2016 | .0164              | 11.3191        | .1680              | -.3131                                       | .3458  | .098   | 4536 | .922            |
| Pair 4 | BMI_2014 - BMI_2016      | .07361             | 1.73773        | .02580             | .02303                                       | .12418 | 2.853  | 4536 | .004            |

# Wilcoxon signed-rank test

- 針對非常態分布的連續數值或小樣本，檢定兩組資料的中位數
- 用於前後測比較

Clinical Rheumatology  
<https://doi.org/10.1007/s10067-023-06735-0>

## ORIGINAL ARTICLE

### Potential alleviation of bone mineral density loss with Janus kinase inhibitors in rheumatoid arthritis

Yun-Wen Chen<sup>1,2</sup> · Hsin-Hua Chen<sup>1,2,3,4,5,6,7</sup> · Wen-Nan Huang<sup>1,3,4,8</sup> · Jun-Peng Chen<sup>9,10</sup> · Yi-Hsing Chen<sup>1,4</sup> · Yi-Ming Chen<sup>1,3,4,10,11</sup> 



**Table 2** Comparisons of the changes in BMD and T-score from the pretreatment to posttreatment assessment based on the RA treatment

| Before                      |      |        |       | After |        |       | <i>p</i> -value |
|-----------------------------|------|--------|-------|-------|--------|-------|-----------------|
| JAKi                        |      |        |       |       |        |       |                 |
| BMD, L spine                | 1.02 | (0.92, | 1.20) | 1.03  | (0.88, | 1.21) | 0.597           |
| BMD, left femoral neck      | 0.84 | (0.76, | 0.90) | 0.84  | (0.76, | 0.92) | 0.168           |
| BMD, right femoral neck     | 0.87 | (0.77, | 0.92) | 0.87  | (0.75, | 0.94) | 0.829           |
| T-score, L spine            | −1.5 | (−2.5, | −0.6) | −1.6  | (−2.5, | −1.0) | 0.836           |
| T-score, left femoral neck  | −1.5 | (−2.0, | −1.0) | −1.7  | (−2.6, | −1.1) | 0.073           |
| T-score, right femoral neck | −1.4 | (−1.9, | −0.8) | −1.8  | (−2.7, | −1.0) | 0.032*          |

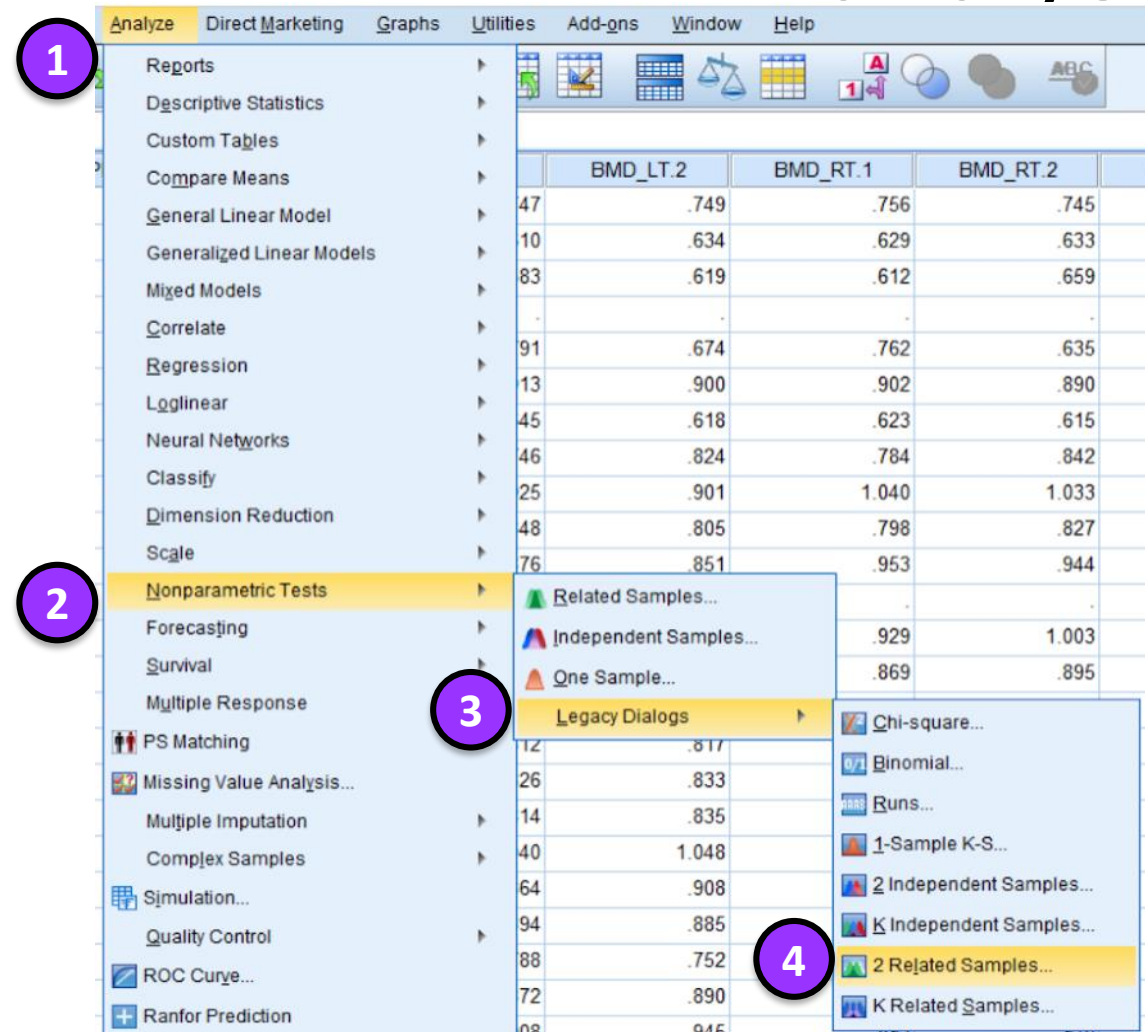
Wilcoxon signed-ranks test. \* $p < 0.05$ , \*\*  $p < 0.01$

# Wilcoxon signed-rank test dataset

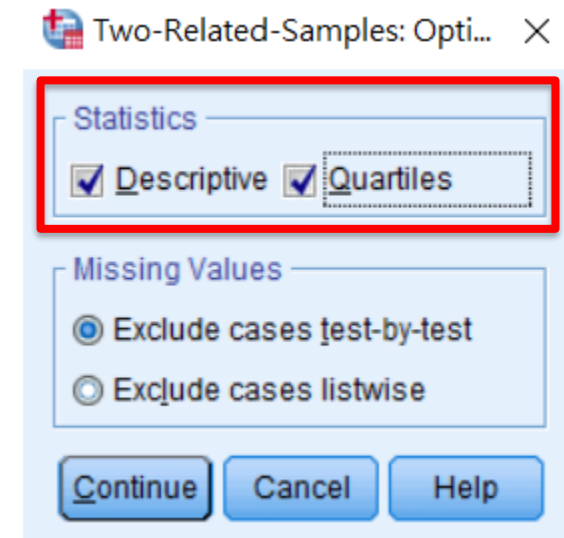
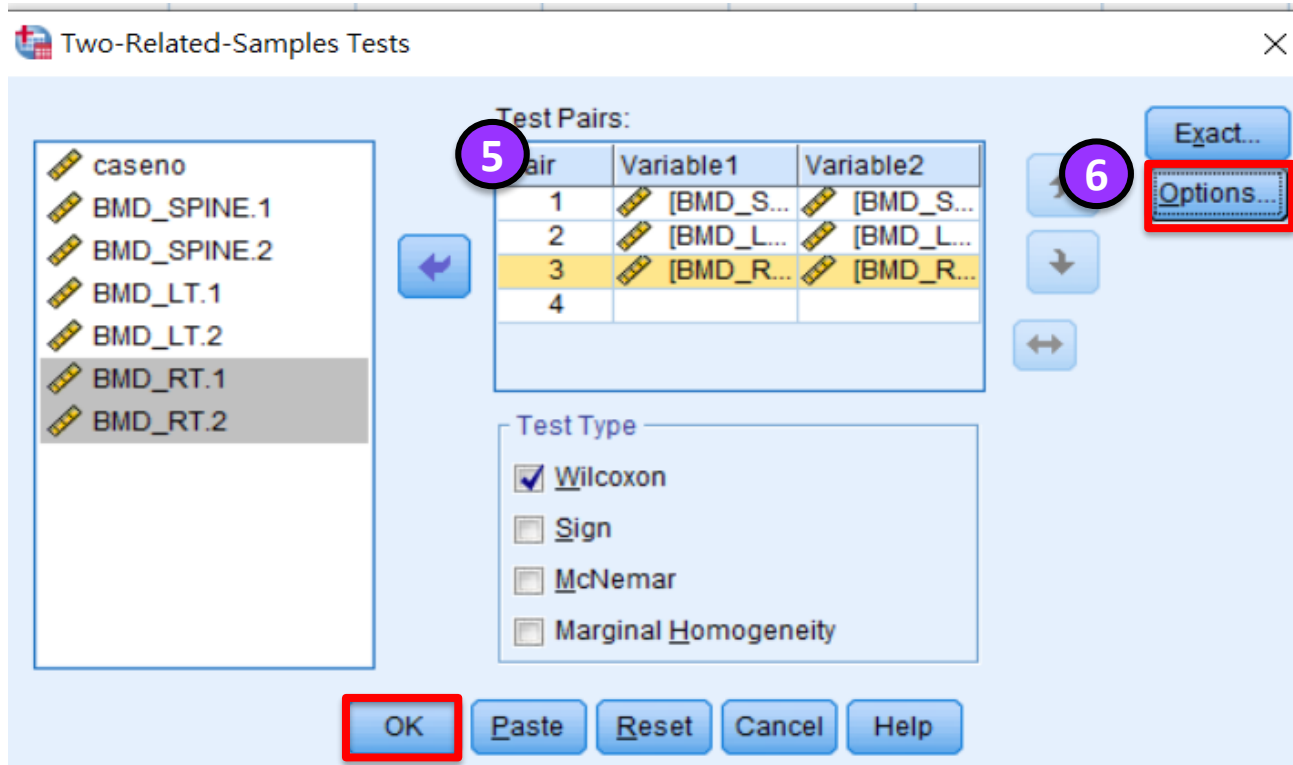
| caseno | BMD_SPINE.1 | BMD_SPINE.2 | BMD_LT.1 | BMD_LT.2 | BMD_RT.1 | BMD_RT.2 |
|--------|-------------|-------------|----------|----------|----------|----------|
| 1      | .805        | .788        | .747     | .749     | .756     | .745     |
| 2      | .866        | .900        | .610     | .634     | .629     | .633     |
| 3      | .804        | .868        | .583     | .619     | .612     | .659     |
| 4      | .714        | .839        | .        | .        | .        | .        |
| 5      | .973        | .813        | .791     | .674     | .762     | .635     |
| 6      | .           | .           | .913     | .900     | .902     | .890     |
| 7      | .823        | .798        | .645     | .618     | .623     | .615     |
| 8      | 1.151       | 1.232       | .746     | .824     | .784     | .842     |
| 9      | 1.086       | .           | .925     | .901     | 1.040    | 1.033    |
| 10     | .989        | .866        | .848     | .805     | .798     | .827     |
| 11     | 1.169       | 1.078       | .876     | .851     | .953     | .944     |
| 12     | .899        | .881        | .        | .        | .        | .        |
| 13     | 1.232       | 1.199       | .888     | .982     | .929     | 1.003    |
| 14     | 1.020       | .974        | .881     | .918     | .869     | .895     |
| 15     | 1.081       | 1.030       | .772     | .783     | .796     | .782     |

# Wilcoxon signed-rank SPSS analysis-1

分析>無母數檢定>歷史對話記錄  
>2個相關樣本



# Wilcoxon signed-rank SPSS analysis-2



# Wilcoxon signed-rank SPSS output

Descriptive Statistics

|             | N  | Mean    | Std. Deviation | Minimum | Maximum | Percentiles |               |         |
|-------------|----|---------|----------------|---------|---------|-------------|---------------|---------|
|             |    |         |                |         |         | 25th        | 50th (Median) | 75th    |
| BMD_SPINE.1 | 28 | 1.05589 | .185310        | .714    | 1.478   | .93050      | 1.01900       | 1.19175 |
| BMD_LT.1    | 28 | .82861  | .131799        | .536    | 1.097   | .76075      | .83700        | .90450  |
| BMD_RT.1    | 28 | .84746  | .141434        | .579    | 1.187   | .76650      | .87050        | .92300  |
| BMD_SPINE.2 | 28 | 1.04739 | .196029        | .725    | 1.463   | .87125      | 1.02250       | 1.20650 |
| BMD_LT.2    | 28 | .83682  | .135854        | .553    | 1.110   | .75550      | .84100        | .91550  |
| BMD_RT.2    | 28 | .84875  | .146636        | .560    | 1.194   | .75375      | .87250        | .93675  |

Test Statistics<sup>a</sup>

|                        | BMD_SPINE.<br>2 -<br>BMD_SPINE.<br>1 | BMD_LT.2 -<br>BMD_LT.1 | BMD_RT.2 -<br>BMD_RT.1 |
|------------------------|--------------------------------------|------------------------|------------------------|
| Z                      | -.529 <sup>b</sup>                   | -1.378 <sup>b</sup>    | -.216 <sup>b</sup>     |
| Asymp. Sig. (2-tailed) | .597                                 | .168                   | .829                   |

a. Wilcoxon Signed Ranks Test

b. Based on negative ranks.

在BMD spine的部分，使用JAK  
Before和After無統計差異  
After的中位數高於Before  
(p=0.597)

# Friedman test

- 相同個案，測量三次時間點(數值型資料)
- 整體顯著可再做事後檢定(Dunn-Bonferroni)

BURNS 46 (2020) 1942–1951

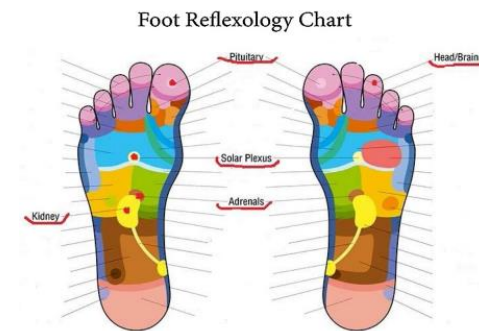
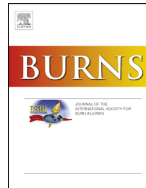


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**The effect of foot reflexology massage on burn-specific pain anxiety and sleep quality and quantity of patients hospitalized in the burn intensive care unit (ICU)**

Reza Alinia-najjar<sup>a</sup>, Masoumeh Bagheri-Nesami<sup>b,c,\*</sup>,  
Seyed Afshin Shorofi<sup>b,d</sup>, Seyed Nouraddin Mousavinasab<sup>e</sup>,  
Kiarash Saatchi<sup>f</sup>

**Table 3 – Comparison of median of duration of last night's sleep on the third, fourth, fifth and sixth days of hospitalization in two groups.**

| Duration of last night's sleep      |        | Third day<br>(before the<br>intervention) | Fourth day<br>(before the<br>intervention) | Fifth day<br>(before the<br>intervention) | Sixth day<br>(before the<br>intervention) | Friedman statistic<br>and p-value |
|-------------------------------------|--------|-------------------------------------------|--------------------------------------------|-------------------------------------------|-------------------------------------------|-----------------------------------|
| Intervention                        | Median | 4.5 (3–7.5) <sup>a</sup>                  | 6 (3–7.3)                                  | 6 (3–7.5)                                 | 8 (5.5–9.75)                              | $X^2=60.184$<br>$p<0.001$         |
| Control                             | Median | 4.5 (3–7.5)                               | 4.30 (3–7.3)                               | 4.5 (3–7.5)                               | 4.5 (3–7)                                 | $X^2=12.183$<br>$p=0.007$         |
| Mann–Whitney U test<br>and p-values |        | Z=0.000<br>$p=1.000$                      | Z=2.28<br>$p=0.023$                        | Z=3.24<br>$p<0.001$                       | Z=6.12<br>$p<0.001$                       |                                   |
| <sup>a</sup> (Min–Max).             |        |                                           |                                            |                                           |                                           |                                   |

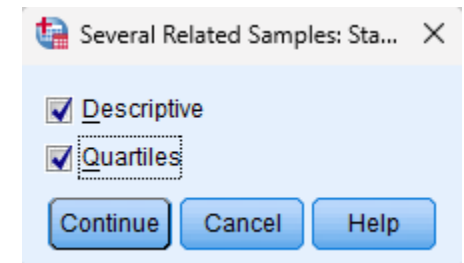
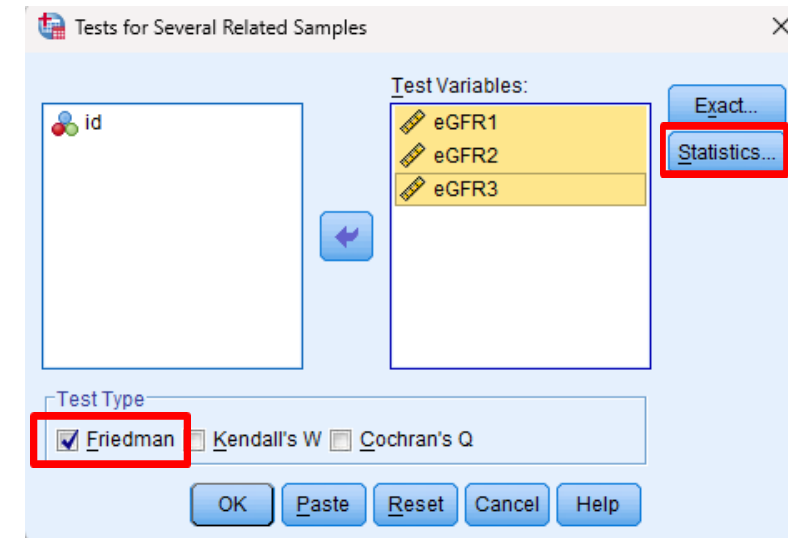
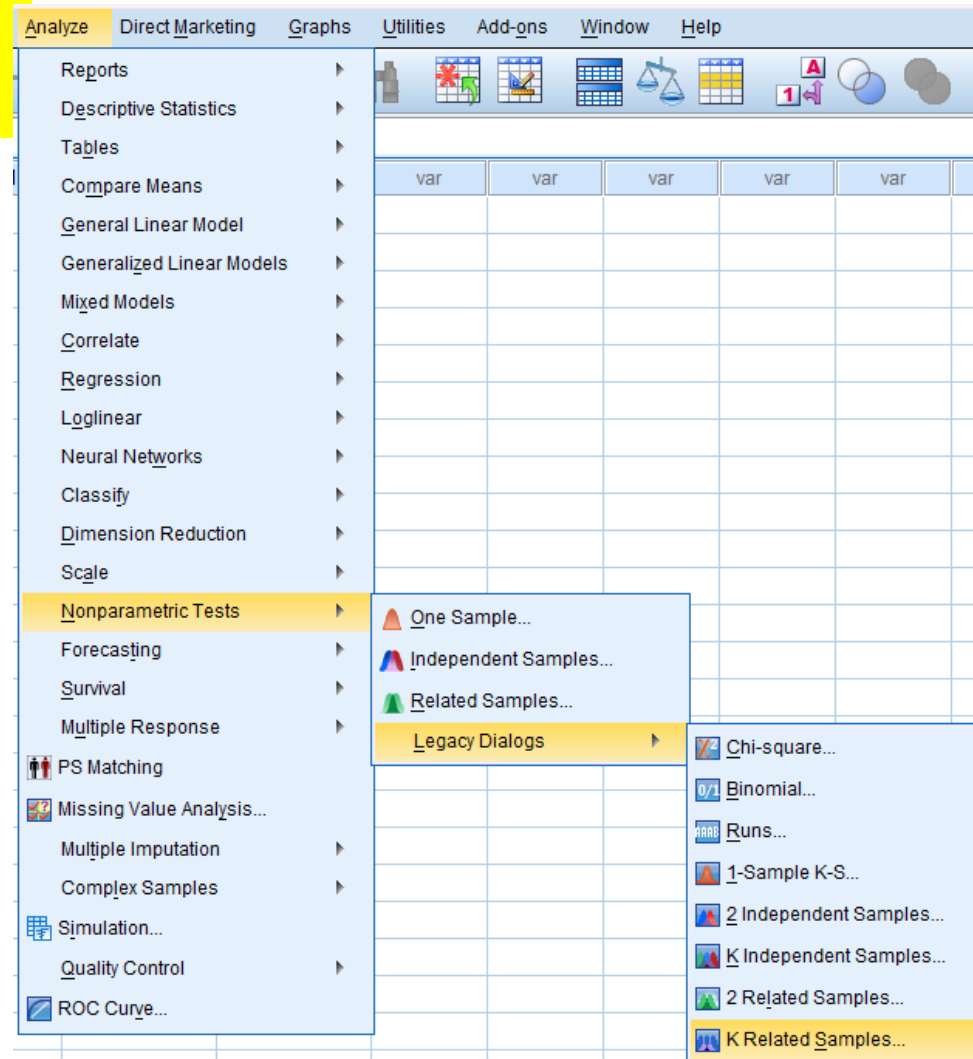
# Friedman test dataset

- 同一組個案，測量三次時間點(數值型資料)
- 整體顯著可再做事後檢定(Dunn-Bonferroni)

| id | eGFR1 | eGFR2 | eGFR3 |
|----|-------|-------|-------|
| 1  | 60.6  | 80.2  | 13.6  |
| 2  | 85.9  | 14.4  | 48.1  |
| 3  | 56.7  | 60.6  | 78.9  |
| 4  | 47.3  | 32.7  | 89.1  |
| 5  | 42.7  | 59.7  | 97.9  |
| 6  | 100.4 | 71.1  | 60.7  |
| 7  | 42.8  | 80.7  | 38.9  |
| 8  | 80.2  | 75.2  | 32.1  |
| 9  | 14.4  | 73.4  | 48.1  |
| 10 | 60.6  | 21.2  | 80.3  |
| 11 | 38.9  | 50.6  | 30.7  |
| 12 | 32.1  | 62.6  | 59.1  |
| 13 | 48.1  | 56.8  | 15.1  |
| 14 | 80.3  | 58.2  | 36.5  |
| 15 | 30.7  | 55.4  | 74.5  |

# Friedman test analysis

分析>無母數檢定>歷史對話記錄>K個相關樣本



# Friedman test output

**Descriptive Statistics**

|       | N  | Mean   | Std. Deviation | Minimum | Maximum | Percentiles |               |        |
|-------|----|--------|----------------|---------|---------|-------------|---------------|--------|
|       |    |        |                |         |         | 25th        | 50th (Median) | 75th   |
| eGFR1 | 57 | 50.565 | 20.7201        | 14.4    | 100.4   | 32.590      | 48.090        | 67.365 |
| eGFR2 | 57 | 59.949 | 17.1628        | 14.4    | 99.8    | 52.150      | 60.570        | 71.830 |
| eGFR3 | 57 | 49.082 | 21.6933        | 13.6    | 97.9    | 32.070      | 42.910        | 69.035 |

## Friedman Test

**Ranks**

|       | Mean Rank |
|-------|-----------|
| eGFR1 | 1.84      |
| eGFR2 | 2.33      |
| eGFR3 | 1.82      |

**Test Statistics<sup>a</sup>**

|             |       |
|-------------|-------|
| N           | 57    |
| Chi-Square  | 9.509 |
| df          | 2     |
| Asymp. Sig. | .009  |

a. Friedman Test

Table. (N=57)

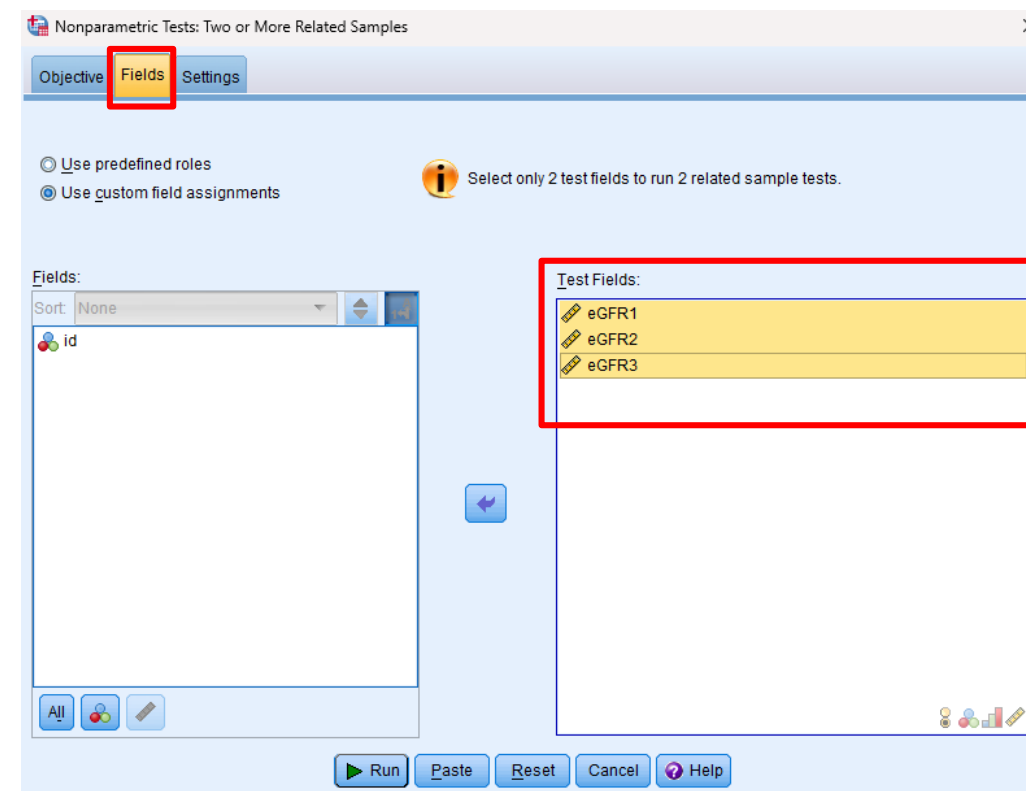
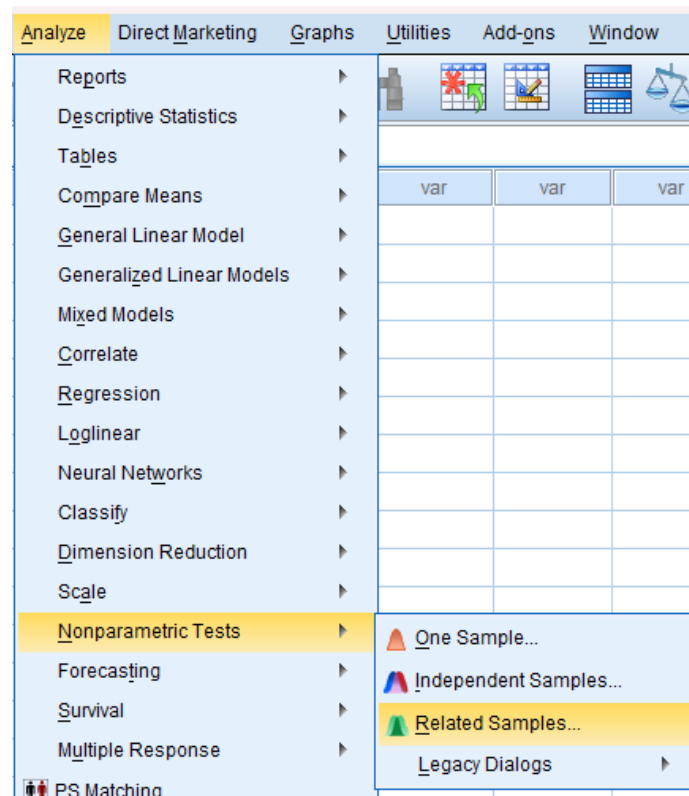
|      | Baseline         | 3 month          | 6 month          | <i>P value</i> |
|------|------------------|------------------|------------------|----------------|
| eGFR | 48.1 (32.6-67.4) | 60.6 (52.2-71.8) | 42.9 (32.1-69.0) | 0.009          |

Friedman test. Median (IQR)

# Friedman test (Post-hoc)

| Name  | Type    | Width | Decimals | Label | Values | Missing | Columns | Align | Measure |
|-------|---------|-------|----------|-------|--------|---------|---------|-------|---------|
| id    | Numeric | 2     | 0        |       | None   | None    | 10      | Right | Nominal |
| eGFR1 | Numeric | 2     | 1        |       | None   | None    | 10      | Right | Scale   |
| eGFR2 | Numeric | 2     | 1        |       | None   | None    | 10      | Right | Scale   |
| eGFR3 | Numeric | 2     | 1        |       | None   | None    | 10      | Right | Scale   |

分析 > 無母數檢定 > 相關樣本

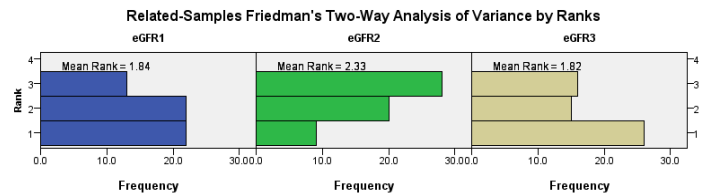


# Friedman test (Post-hoc output)

Hypothesis Test Summary

|   | Null Hypothesis                                           | Test                                                             | Sig. | Decision                    |
|---|-----------------------------------------------------------|------------------------------------------------------------------|------|-----------------------------|
| 1 | The distributions of eGFR1, eGFR2 and eGFR3 are the same. | Related-Samples Friedman's Two-Way Analysis of Variance by Ranks | .009 | Reject the null hypothesis. |

Asymptotic significances are displayed. The significance level is .05.



|                                |       |
|--------------------------------|-------|
| Total N                        | 57    |
| Test Statistic                 | 9.509 |
| Degrees of Freedom             | 2     |
| Asymptotic Sig. (2-sided test) | .009  |

Related Samples Test View  
Continuous Field Information  
Pairwise Comparisons  
Related Samples Test View

Test: Friedman Field(s): eGFR1, eGFR2, eGFR3(Test 1) View: Related Samples Test View

Each node shows the sample average rank.

| Sample1-Sample2 | Test Statistic | Std. Error | Std. Test Statistic | Sig. | Adj.Sig. |
|-----------------|----------------|------------|---------------------|------|----------|
| eGFR3-eGFR1     | .018           | .187       | .094                | .925 | 1.000    |
| eGFR3-eGFR2     | .509           | .187       | 2.716               | .007 | .020     |
| eGFR1-eGFR2     | -.491          | .187       | -2.622              | .009 | .026     |

Each row tests the null hypothesis that the Sample 1 and Sample 2 distributions are the same. Asymptotic significances (2-sided tests) are displayed. The significance level is .05.

# Repeated measures ANOVA

- Outcome為數值型資料且需符合常態分佈
- 寬資料，有一次資料missing則排除此個案

## Neurodegeneration

### ORIGINAL RESEARCH

Effects of vitamin D supplementation on cognitive function and blood A $\beta$ -related biomarkers in older adults with Alzheimer's disease: a randomised, double-blind, placebo-controlled trial

Jingya Jia,<sup>1</sup> Jing Hu,<sup>1</sup> Xiaoxu Huo,<sup>1</sup> Rujuan Miao,<sup>1</sup> Yanping Zhang,<sup>2</sup> Fei Ma <sup>1</sup>

**Table 2** The level of biochemical parameters at baseline, 6 months and 12 months between the two groups

| Items                | groups       | Cases (n) | Treatment time*    |                    |                    | Repeated measures†    |                |                 |
|----------------------|--------------|-----------|--------------------|--------------------|--------------------|-----------------------|----------------|-----------------|
|                      |              |           | Baseline           | 6 months           | 12 months          | Interaction effect, P | Time effect, P | Group effect, P |
| A $\beta$ 40 (pg/mL) | Intervention | 105       | 27.78 $\pm$ 20.80  | 26.88 $\pm$ 20.81  | 28.62 $\pm$ 20.79  | 0.468                 | 0.027          | 0.480           |
|                      | Control      | 105       | 33.46 $\pm$ 24.06  | 32.43 $\pm$ 24.00  | 33.22 $\pm$ 23.20  |                       |                |                 |
| A $\beta$ 42 (pg/mL) | Intervention | 105       | 34.39 $\pm$ 21.71  | 32.33 $\pm$ 21.70  | 30.50 $\pm$ 21.69  | <0.001                | <0.001         | 0.987           |
|                      | Control      | 105       | 33.28 $\pm$ 21.26  | 33.76 $\pm$ 21.24  | 33.19 $\pm$ 20.51  |                       |                |                 |
| PS1 (pg/mL)          | Intervention | 105       | 48.78 $\pm$ 44.39  | 48.89 $\pm$ 44.39  | 48.89 $\pm$ 44.39  | 0.951                 | 0.727          | 0.272           |
|                      | Control      | 105       | 44.51 $\pm$ 37.92  | 44.63 $\pm$ 37.92  | 44.74 $\pm$ 37.65  |                       |                |                 |
| PS2 (pg/mL)          | Intervention | 105       | 10.43 $\pm$ 19.02  | 10.57 $\pm$ 19.02  | 10.69 $\pm$ 19.03  | 0.142                 | <0.001         | 0.133           |
|                      | Control      | 105       | 11.01 $\pm$ 17.44  | 11.17 $\pm$ 17.44  | 11.55 $\pm$ 17.53  |                       |                |                 |
| PS1mRNA              | Intervention | 105       | 28.85 $\pm$ 2.59   | 30.19 $\pm$ 2.61   | 29.77 $\pm$ 2.59   | 0.040                 | <0.001         | 0.070           |
|                      | Control      | 105       | 28.89 $\pm$ 2.58   | 30.41 $\pm$ 2.58   | 29.86 $\pm$ 2.60   |                       |                |                 |
| PS2mRNA              | Intervention | 105       | 4.00 $\pm$ 0.43    | 4.34 $\pm$ 0.53    | 4.17 $\pm$ 0.53    | 0.121                 | <0.001         | 0.409           |
|                      | Control      | 105       | 4.18 $\pm$ 0.39    | 4.60 $\pm$ 0.61    | 4.34 $\pm$ 0.38    |                       |                |                 |
| APP                  | Intervention | 105       | 44.75 $\pm$ 39.77  | 41.69 $\pm$ 39.73  | 39.96 $\pm$ 39.59  | <0.001                | <0.001         | <0.001          |
|                      | Control      | 105       | 44.71 $\pm$ 33.20  | 42.56 $\pm$ 33.20  | 45.56 $\pm$ 33.24  |                       |                |                 |
| APPMRNA              | Intervention | 105       | 32.59 $\pm$ 1.53   | 30.28 $\pm$ 1.50   | 27.96 $\pm$ 1.54   | <0.001                | <0.001         | <0.001          |
|                      | Control      | 105       | 32.41 $\pm$ 1.41   | 31.44 $\pm$ 1.47   | 31.55 $\pm$ 1.38   |                       |                |                 |
| BACE1 ( $\mu$ g/mL)  | Intervention | 105       | 300.83 $\pm$ 11.96 | 298.14 $\pm$ 11.97 | 296.23 $\pm$ 12.01 | <0.001                | <0.001         | <0.001          |
|                      | Control      | 105       | 299.94 $\pm$ 12.05 | 297.76 $\pm$ 12.06 | 300.62 $\pm$ 11.84 |                       |                |                 |
| BACE1mRNA            | Intervention | 105       | 27.59 $\pm$ 1.29   | 25.29 $\pm$ 1.55   | 23.48 $\pm$ 1.91   | <0.001                | <0.001         | <0.001          |
|                      | Control      | 105       | 27.24 $\pm$ 1.32   | 26.09 $\pm$ 1.31   | 27.82 $\pm$ 1.27   |                       |                |                 |
| 25-D (ng/mL)         | Intervention | 105       | 18.82 $\pm$ 2.91   | 20.90 $\pm$ 3.12   | 22.77 $\pm$ 3.41   | <0.001                | <0.001         | <0.001          |
|                      | Control      | 105       | 19.44 $\pm$ 2.81   | 19.18 $\pm$ 2.81   | 19.08 $\pm$ 2.84   |                       |                |                 |
| 1,25-D (ng/mL)       | Intervention | 105       | 30.37 $\pm$ 2.60   | 31.79 $\pm$ 2.57   | 33.61 $\pm$ 2.77   | <0.001                | <0.001         | <0.001          |
|                      | Control      | 105       | 30.22 $\pm$ 2.67   | 29.01 $\pm$ 2.67   | 30.71 $\pm$ 2.64   |                       |                |                 |

\*Presented as mean $\pm$ SD.

†P value for group (intervention vs control) derived from analysis of covariance adjusted for respective baseline value and for age, gender and education.

A $\beta$ , amyloid beta; APP, A $\beta$  protein precursor; BACE1,  $\beta$ -secretase 1; 25-D, 25-hydroxy vitamin D; 1,25-D, 1,25-dihydroxy vitamin; PS, presenilin.

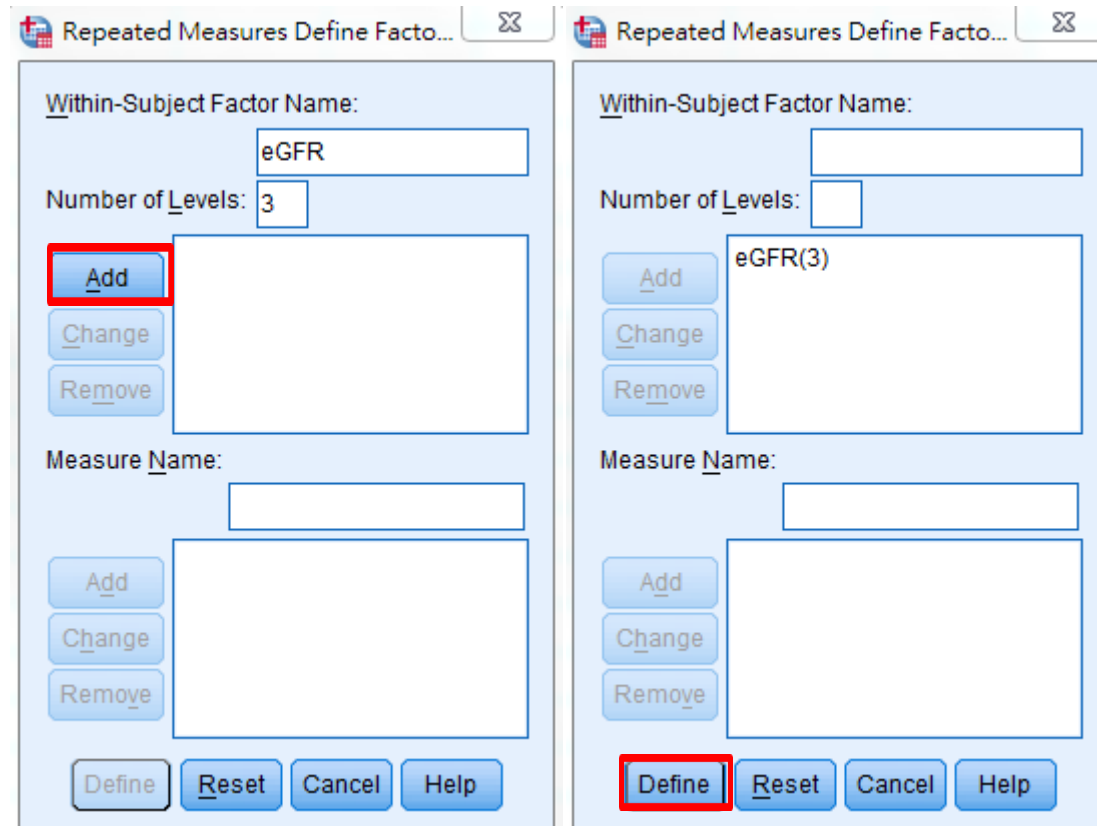
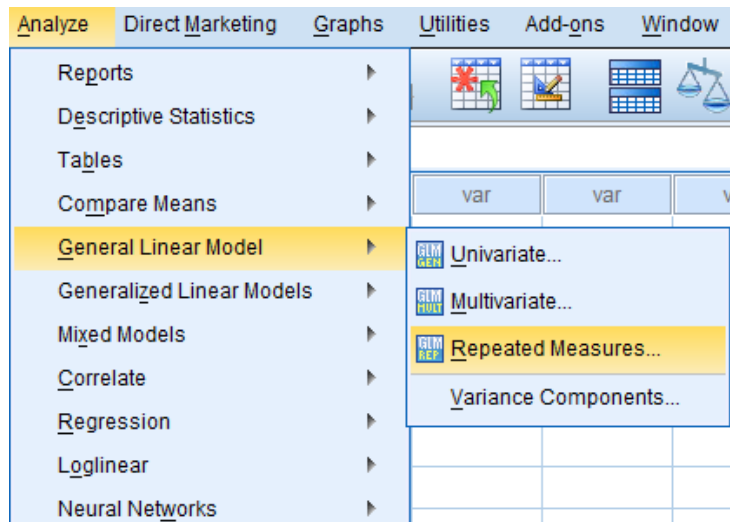
# Repeated measures ANOVA dataset

- 兩組測量三個時間點(數值型資料)
- 整體顯著可再做事後檢定

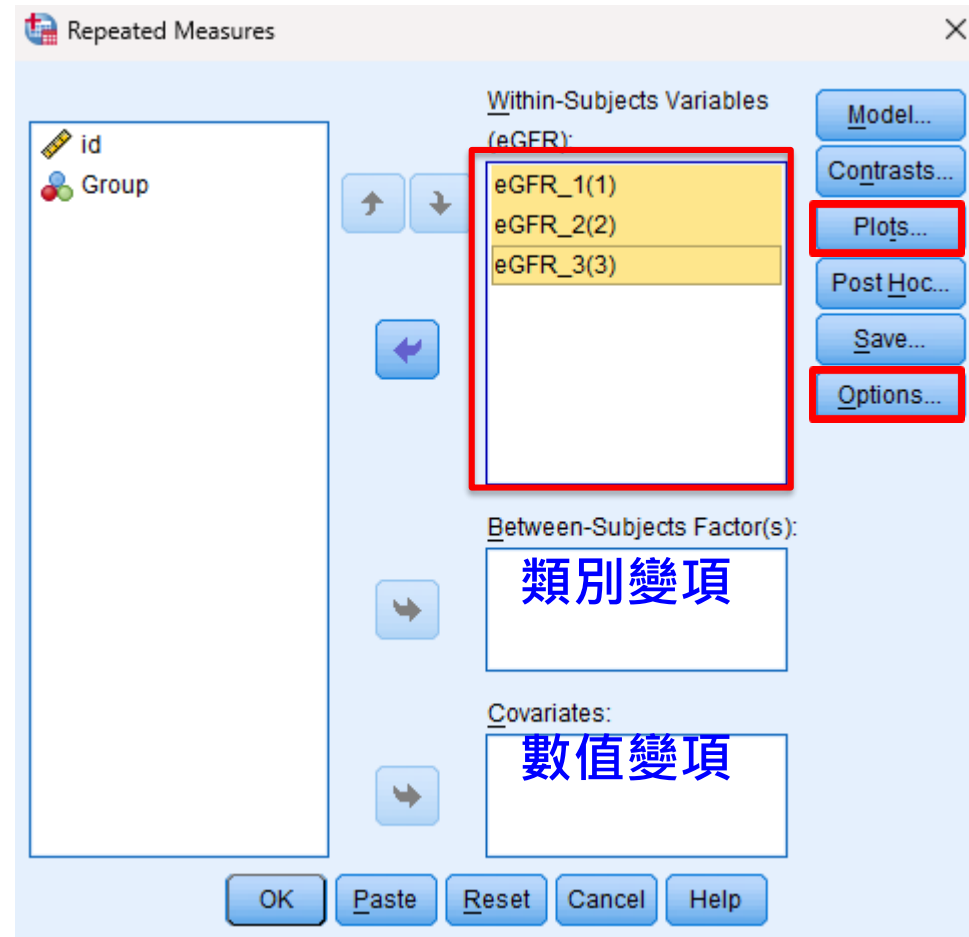
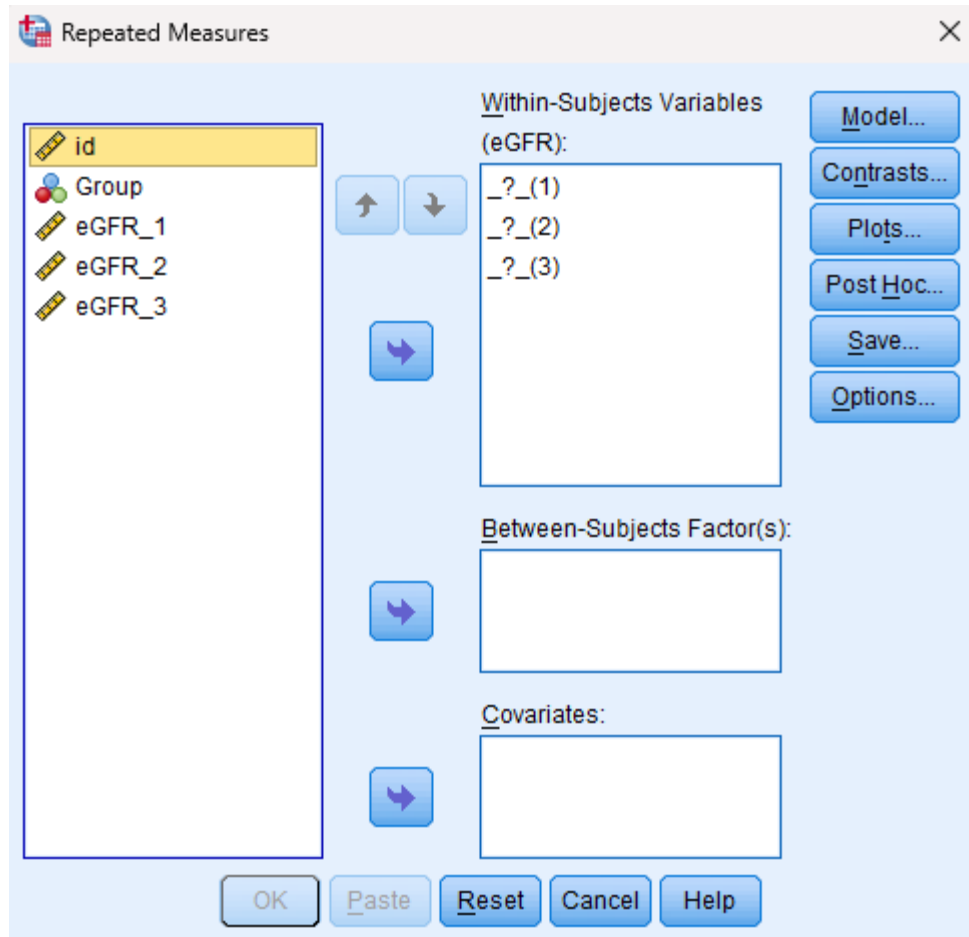
| id | Group | eGFR_1 | eGFR_2 | eGFR_3 |
|----|-------|--------|--------|--------|
| 1  | 1     | 51.58  | 53.66  | 53.36  |
| 2  | 1     | 3.25   | 3.59   | 3.29   |
| 3  | 1     | 4.63   | 3.46   | 3.16   |
| 4  | 1     | 6.35   | 9.14   | 8.84   |
| 5  | 1     | 6.94   | 15.11  | 14.81  |
| 6  | 1     | 11.74  | 35.74  | 35.44  |
| 7  | 1     | 13.61  | 14.83  | 14.53  |
| 8  | 1     | 14.03  | 51.58  | 51.28  |
| 9  | 1     | 14.38  | 22.64  | 22.34  |
| 10 | 1     | 14.70  | 12.23  | 11.93  |
| 11 | 1     | 16.97  | 25.45  | 25.15  |
| 12 | 1     | 19.99  | 38.94  | 38.64  |
| 13 | 1     | 21.23  | 31.18  | 30.88  |
| 14 | 1     | 21.99  | 32.92  | 32.62  |
| 15 | 0     | 26.60  | 30.72  | 30.42  |

# Repeated measures ANOVA analysis-1

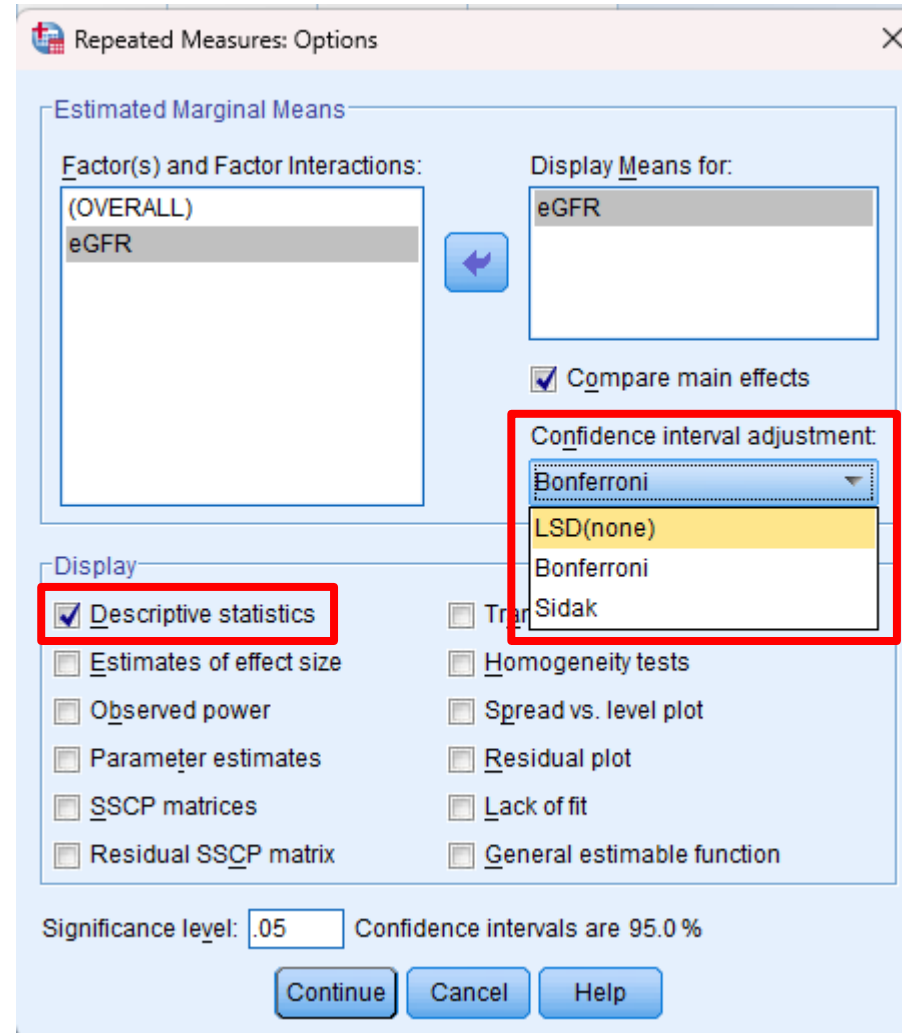
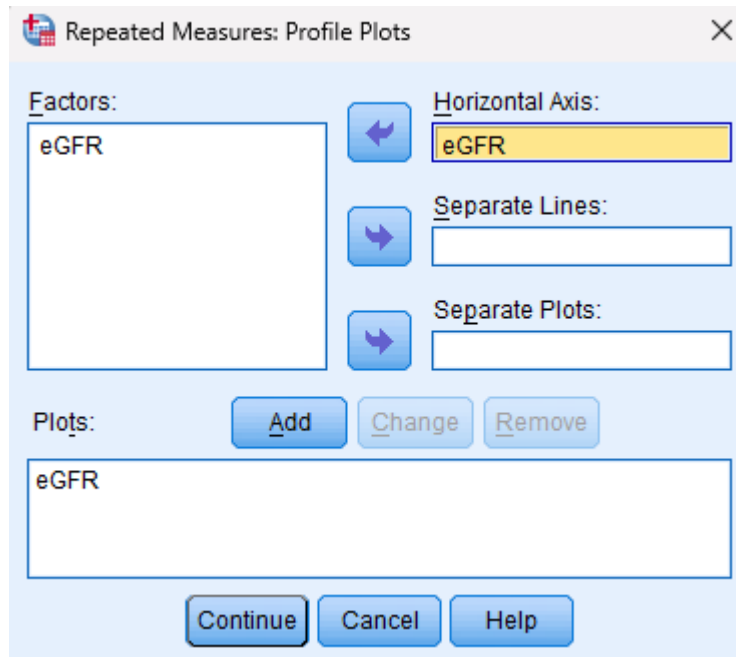
分析 > 一般線性模式 > 重複測量



# Repeated measures ANOVA analysis-2



# Repeated measures ANOVA analysis-3



# Repeated measures ANOVA output

**Descriptive Statistics**

|        | Mean    | Std. Deviation | N    |
|--------|---------|----------------|------|
| eGFR_1 | 59.3108 | 23.02112       | 1127 |
| eGFR_2 | 57.5311 | 23.78461       | 1127 |
| eGFR_3 | 57.2311 | 23.78461       | 1127 |

**Mauchly's Test of Sphericity<sup>a</sup>**

Measure: MEASURE\_1

| Within Subjects Effect | Mauchly's W | Approx. Chi-Square | df | Sig. | Epsilon <sup>b</sup> |             |             |
|------------------------|-------------|--------------------|----|------|----------------------|-------------|-------------|
|                        |             |                    |    |      | Greenhouse-Geisser   | Huynh-Feldt | Lower-bound |
| eGFR                   | .000        | 36007.760          | 2  | .000 | .500                 | .500        | .500        |

Tests the null hypothesis that the error covariance matrix of the orthonormalized transformed dependent variables is proportional to an identity matrix.

a. Design: Intercept  
Within Subjects Design: eGFR

b. May be used to adjust the degrees of freedom for the averaged tests of significance. Corrected tests are displayed in the Tests of Within-Subjects Effects table.

**Tests of Within-Subjects Effects**

Measure: MEASURE\_1

| Source      | Type III Sum of Squares | df   | Mean Square | F      | Sig. |
|-------------|-------------------------|------|-------------|--------|------|
| eGFR        | 2848.636                | 2    | 1424.318    | 20.420 | .000 |
| Error(eGFR) | 157081.715              | 2252 | 69.752      |        |      |
| Total       | 159930.351              | 2254 |             |        |      |

**$P < 0.05$  代表資料未符合球形假設**

**Pairwise Comparisons**

Measure: MEASURE\_1

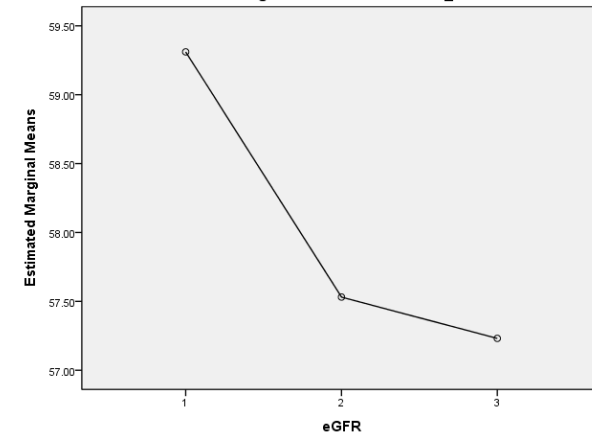
| (I) eGFR | (J) eGFR | Mean Difference (I-J) | Std. Error | Sig. <sup>b</sup> | 95% Confidence Interval for Difference <sup>b</sup> |             |
|----------|----------|-----------------------|------------|-------------------|-----------------------------------------------------|-------------|
|          |          |                       |            |                   | Lower Bound                                         | Upper Bound |
| 1        | 2        | 1.780 <sup>*</sup>    | .431       | .000              | .747                                                | 2.813       |
|          | 3        | 2.080 <sup>*</sup>    | .431       | .000              | 1.047                                               | 3.113       |
| 2        | 1        | -1.780 <sup>*</sup>   | .431       | .000              | -2.813                                              | -.747       |
|          | 3        | .300 <sup>*</sup>     | .000       | .000              | .300                                                | .300        |
| 3        | 1        | -2.080 <sup>*</sup>   | .431       | .000              | -3.113                                              | -1.047      |
|          | 2        | -.300 <sup>*</sup>    | .000       | .000              | -.300                                               | -.300       |

Based on estimated marginal means

\*. The mean difference is significant at the .05 level.

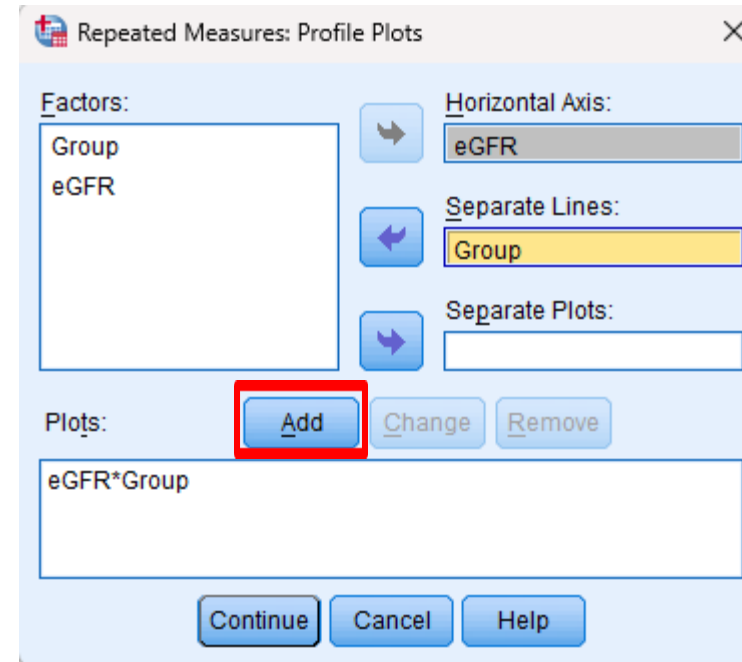
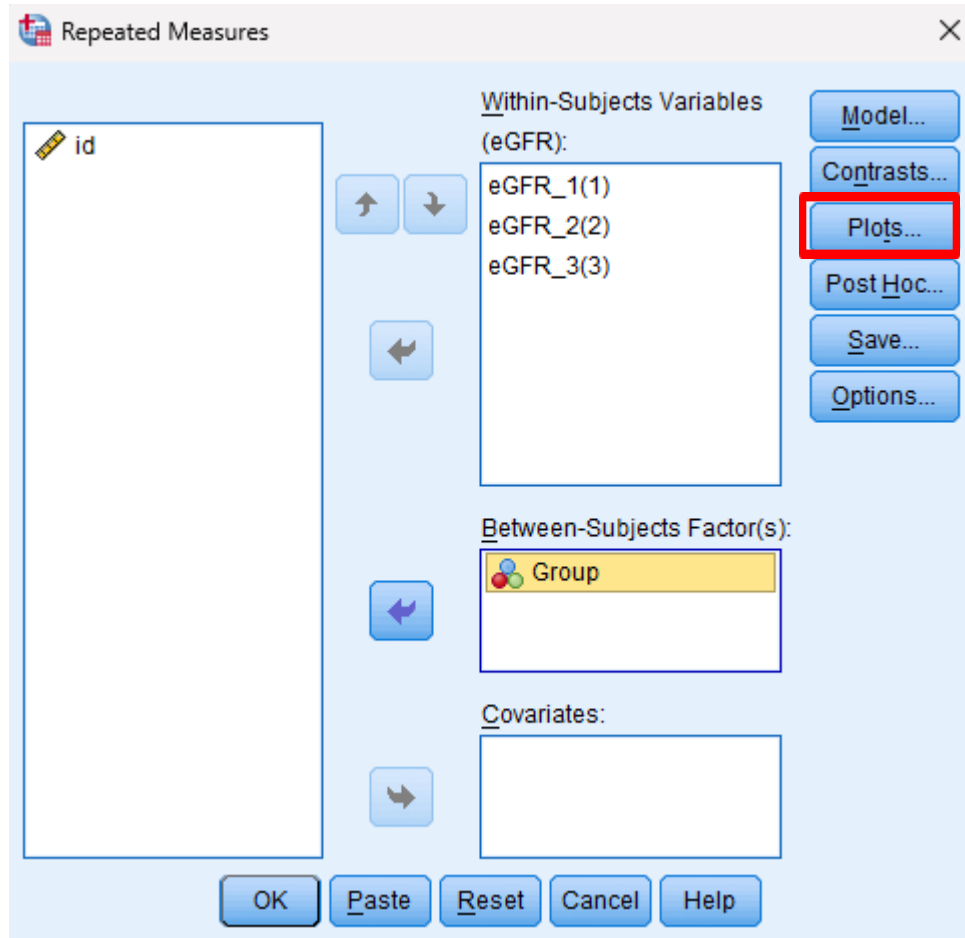
b. Adjustment for multiple comparisons: Bonferroni.

**Estimated Marginal Means of MEASURE\_1**



不符合球形假設

# Repeated measures ANOVA analysis



# Repeated measures ANOVA output

**Descriptive Statistics**

| Group    | Mean    | Std. Deviation | N    |
|----------|---------|----------------|------|
| eGFR_1 0 | 66.1851 | 19.40228       | 138  |
| 1        | 58.3516 | 23.32970       | 989  |
| Total    | 59.3108 | 23.02112       | 1127 |
| eGFR_2 0 | 63.7180 | 19.50203       | 138  |
| 1        | 56.6678 | 24.20522       | 989  |
| Total    | 57.5311 | 23.78461       | 1127 |
| eGFR_3 0 | 63.4180 | 19.50203       | 138  |
| 1        | 56.3678 | 24.20522       | 989  |
| Total    | 57.2311 | 23.78461       | 1127 |

**Mauchly's Test of Sphericity<sup>a</sup>**

Measure: MEASURE\_1

| Within Subjects Effect | Mauchly's W | Approx. Chi-Square | df | Sig. | Epsilon <sup>b</sup> |             |             |
|------------------------|-------------|--------------------|----|------|----------------------|-------------|-------------|
|                        |             |                    |    |      | Greenhouse-Geisser   | Huynh-Feldt | Lower-bound |
| eGFR                   | .000        | 36298.754          | 2  | .000 | .500                 | .500        | .500        |

Tests the null hypothesis that the error covariance matrix of the orthonormalized transformed dependent variables is proportional to an identity matrix.

a. Design: Intercept + Group  
Within Subjects Design: eGFR

b. May be used to adjust the degrees of freedom for the averaged tests of significance. Corrected tests are displayed in the Tests of Within-Subjects Effects table.

**Tests of Within-Subjects Effects**

Measure: MEASURE\_1

| Source       |                    | Type III Sum of Squares | df       | Mean Square | F      | Sig. |
|--------------|--------------------|-------------------------|----------|-------------|--------|------|
| eGFR         | Sphericity Assumed | 1621.180                | 2        | 810.590     | 11.614 | .000 |
|              | Greenhouse-Geisser | 1621.180                | 1.000    | 1621.180    | 11.614 | .001 |
|              | Huynh-Feldt        | 1621.180                | 1.001    | 1619.739    | 11.614 | .001 |
|              | Lower-bound        | 1621.180                | 1.000    | 1621.180    | 11.614 | .001 |
| eGFR * Group | Sphericity Assumed | 49.520                  | 2        | 24.760      | .355   | .701 |
|              | Greenhouse-Geisser | 49.520                  | 1.000    | 49.520      | .355   | .552 |
|              | Huynh-Feldt        | 49.520                  | 1.001    | 49.476      | .355   | .552 |
|              | Lower-bound        | 49.520                  | 1.000    | 49.520      | .355   | .552 |
| Error(eGFR)  | Sphericity Assumed | 157032.196              | 2250     | 69.792      |        |      |
|              | Greenhouse-Geisser | 157032.196              | 1125.000 | 139.584     |        |      |
|              | Huynh-Feldt        | 157032.196              | 1126.001 | 139.460     |        |      |
|              | Lower-bound        | 157032.196              | 1125.000 | 139.584     |        |      |

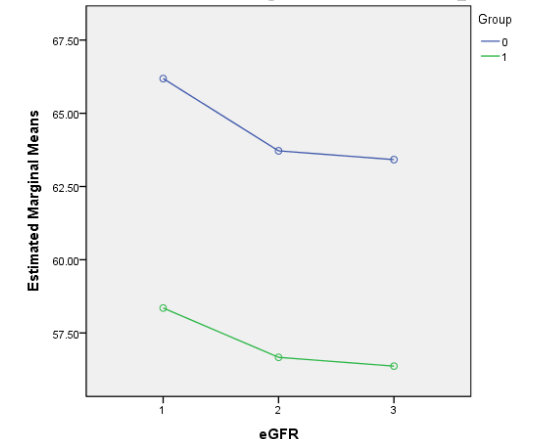
**Tests of Between-Subjects Effects**

Measure: MEASURE\_1

Transformed Variable: Average

| Source    | Type III Sum of Squares | df   | Mean Square | F        | Sig. |
|-----------|-------------------------|------|-------------|----------|------|
| Intercept | 5369346.712             | 1    | 5369346.712 | 3565.368 | .000 |
| Group     | 19420.818               | 1    | 19420.818   | 12.896   | .000 |
| Error     | 1694219.138             | 1125 | 1505.973    |          |      |

**Estimated Marginal Means of MEASURE\_1**



# Linear Mixed Model / Generalized Estimating Equation

- Linear Mixed Model
  - Outcome為數值型資料且需符合常態分佈
  - 資料型態為長資料，可容許遺漏值的存在
- GEE
  - 不需符合常態分佈假設，為半母數方法(semi-parametric)
  - Outcome可為連續、類別、順序或計數類型
  - 資料型態為長資料，可容許遺漏值的存在

# GEE矩陣

- 獨立矩陣 ( Independent )
- AR(1) 矩陣 ( Auto-regressive first order )
- 可交換矩陣 ( Exchangeable )
- 未結構化矩陣 ( Unstructured )

獨立矩陣

|       | $t_1$ | $t_2$ | $t_3$ | $t_4$ | $t_5$ |
|-------|-------|-------|-------|-------|-------|
| $t_1$ | —     |       |       |       |       |
| $t_2$ | 0     | —     |       |       |       |
| $t_3$ | 0     | 0     | —     |       |       |
| $t_4$ | 0     | 0     | 0     | —     |       |
| $t_5$ | 0     | 0     | 0     | 0     | —     |

AR(1) 矩陣

|       | $t_1$    | $t_2$    | $t_3$    | $t_4$  | $t_5$ |
|-------|----------|----------|----------|--------|-------|
| $t_1$ | —        |          |          |        |       |
| $t_2$ | $\rho$   | —        |          |        |       |
| $t_3$ | $\rho^2$ | $\rho$   | —        |        |       |
| $t_4$ | $\rho^3$ | $\rho^2$ | $\rho$   | —      |       |
| $t_5$ | $\rho^4$ | $\rho^3$ | $\rho^2$ | $\rho$ | —     |

可交換矩陣

|       | $t_1$  | $t_2$  | $t_3$  | $t_4$  | $t_5$ |
|-------|--------|--------|--------|--------|-------|
| $t_1$ | —      |        |        |        |       |
| $t_2$ | $\rho$ | —      |        |        |       |
| $t_3$ | $\rho$ | $\rho$ | —      |        |       |
| $t_4$ | $\rho$ | $\rho$ | $\rho$ | —      |       |
| $t_5$ | $\rho$ | $\rho$ | $\rho$ | $\rho$ | —     |

未結構化矩陣

|       | $t_1$    | $t_2$    | $t_3$    | $t_4$       | $t_5$ |
|-------|----------|----------|----------|-------------|-------|
| $t_1$ | —        |          |          |             |       |
| $t_2$ | $\rho_1$ | —        |          |             |       |
| $t_3$ | $\rho_2$ | $\rho_5$ | —        |             |       |
| $t_4$ | $\rho_3$ | $\rho_6$ | $\rho_8$ | —           |       |
| $t_5$ | $\rho_4$ | $\rho_7$ | $\rho_9$ | $\rho_{10}$ | —     |

# GEE

## Midlife overweight and obesity increase late-life dementia risk

A population-based twin study

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A.R. Atti, MD, PhD  
M. Gatz, PhD  
N.L. Pedersen, PhD  
B. Johansson, PhD  
L. Fratiglioni, MD, PhD

### ABSTRACT

**Objective:** The relation of overweight to dementia is controversial. We aimed to examine the association of midlife overweight and obesity with dementia, Alzheimer disease (AD), and vascular dementia (VaD) in late life, and to verify the hypothesis that genetic and early-life environmental factors contribute to the observed association.

**Methods:** From the Swedish Twin Registry, 8,534 twin individuals aged  $\geq 65$  (mean age 74.4) were assessed to detect dementia cases (DSM-IV criteria). Height and weight at midlife (mean age 43.4) were available in the Registry. Data were analyzed as follows: 1) unmatched case-

**Table 2** Adjusted odds ratio (OR) and 95% confidence interval (CI) of dementia, Alzheimer disease, and vascular dementia related to midlife BMI (results from generalized estimating equation models)

| Midlife BMI        | No. of twins | All dementia |                          |                          | Alzheimer disease |                          |                          | Vascular dementia |                          |                          |
|--------------------|--------------|--------------|--------------------------|--------------------------|-------------------|--------------------------|--------------------------|-------------------|--------------------------|--------------------------|
|                    |              | No.          | OR (95% CI) <sup>a</sup> | OR (95% CI) <sup>b</sup> | No.               | OR (95% CI) <sup>a</sup> | OR (95% CI) <sup>b</sup> | No.               | OR (95% CI) <sup>a</sup> | OR (95% CI) <sup>b</sup> |
| Continuous         | 8,534        | 464          | 1.09 (1.06-1.12)         | 1.06 (1.03-1.10)         | 232               | 1.09 (1.04-1.13)         | 1.06 (1.01-1.10)         | 74                | 1.14 (1.08-1.21)         | 1.11 (1.04-1.19)         |
| <b>Categorical</b> |              |              |                          |                          |                   |                          |                          |                   |                          |                          |
| <20                | 627          | 17           | 0.74 (0.44-1.25)         | 0.79 (0.45-1.38)         | 8                 | 0.89 (0.64-1.23)         | 0.66 (0.31-1.41)         | 0                 | —                        | —                        |
| 20-25              | 5,366        | 240          | 1 (Reference)            | 1 (Reference)            | 120               | 1 (Reference)            | 1 (Reference)            | 36                | 1 (Reference)            | 1 (Reference)            |
| >25                | 2,541        | 207          | 1.50 (1.22-1.84)         | 1.80 (1.37-2.35)         | 104               | 1.52 (1.15-2.02)         | 1.98 (1.36-2.88)         | 38                | 1.62 (1.01-2.59)         | 1.35 (0.81-2.24)         |
| 25-30              | 2,297        | 177          | 1.37 (1.11-1.70)         | 1.71 (1.30-2.25)         | 90                | 1.41 (1.05-1.89)         | 1.91 (1.30-2.80)         | 31                | 1.39 (0.85-2.29)         | 1.17 (0.69-2.00)         |
| >30                | 244          | 30           | 3.01 (1.95-4.64)         | 3.88 (2.12-7.11)         | 14                | 2.87 (1.57-5.26)         | 3.43 (1.49-7.90)         | 7                 | 4.38 (1.89-10.14)        | 3.50 (1.36-8.99)         |

Abbreviations: BMI = body mass index; CI = confidence interval; OR = odds ratio.

<sup>a</sup> Adjusted for age, sex, and education.

<sup>b</sup> Adjusted for age, sex, education, diabetes, hypertension, stroke, and heart disease.

# GEE

## Summative Effects of Vascular Risk Factors on the Progression of Alzheimer Disease

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Yung-Shuan Lin, MD,<sup>†\*\*</sup> Shuu-Jiun Wang, MD,<sup>†¶\*\*</sup> and Jong-Ling Fuh, MD<sup>†¶\*\*</sup>

**OBJECTIVES:** To investigate the summative effects of vascular risk factors (VRFs) on the progression of Alzheimer disease (AD).

**DESIGN:** Longitudinal follow-up cohort study.

**SETTING:** AD patients from two teaching hospitals in Taiwan with 3-year follow-ups.

three or fewer VRFs (MMSE,  $P = .009$ ; CDRSB,  $P = .02$ ). Subsequent analyses of APOE  $\epsilon 4$  carriers with more than three VRFs also showed their more rapid cognitive decline compared with patients without VRFs (MMSE,  $P = .02$ ; CDRSB,  $P = .001$ ) and patients with three or fewer VRFs (MMSE,  $P = .009$ ; CDRSB,  $P = .02$ ), but no significant difference was found in APOE  $\epsilon 4$  noncarriers.

| Vascular risk factors, No. (%) |            |
|--------------------------------|------------|
| Coronary heart disease         | 63 (19.1)  |
| Cardiac arrhythmia             | 30 (9.1)   |
| Cerebrovascular disease        | 12 (3.6)   |
| Hypertension                   | 217 (65.8) |
| Diabetes mellitus              | 154 (46.7) |
| Obesity                        | 18 (5.5)   |
| Smoking                        | 44 (13.3)  |
| Physical inactivity            | 121 (36.7) |

Table 2. Results of the generalized estimating equation analyzing the effect of 3-year MMSE changes in AD patients with and without APOE  $\epsilon 4$  and different VRF indexes

| Variable                            | All AD patients (n = 330) |                      | AD patients with APOE $\epsilon 4$ (n = 129) |                      | AD patients without APOE $\epsilon 4$ (n = 201) |                      |
|-------------------------------------|---------------------------|----------------------|----------------------------------------------|----------------------|-------------------------------------------------|----------------------|
|                                     | $\beta$ (95% CI)          | P value <sup>a</sup> | $\beta$ (95% CI)                             | P value <sup>a</sup> | $\beta$ (95% CI)                                | P value <sup>a</sup> |
| <b>VRF groups</b>                   |                           |                      |                                              |                      |                                                 |                      |
| >3 VRFs                             | −1.16 (−2.1 to −0.21)     | .02                  | −1.99 (−3.62 to −0.36)                       | .02                  | −0.59 (−1.7 to 0.52)                            | .3                   |
| 3 VRFs                              | −0.36 (−1.24 to 0.51)     | .42                  | −1.2 (−2.8 to 0.4)                           | .14                  | 0.11 (−0.9 to 1.1)                              | .83                  |
| 2 VRFs                              | −0.15 (−0.92 to 0.61)     | .69                  | −0.51 (−1.8 to 0.78)                         | .44                  | 0.27 (−0.64 to 1.18)                            | .56                  |
| 1 VRF                               | −0.04 (−0.83 to 0.73)     | .89                  | −0.48 (−1.76 to 0.81)                        | .47                  | 0.3 (−0.68 to 1.28)                             | .55                  |
| 0 VRFs                              | Reference                 |                      | Reference                                    |                      | Reference                                       |                      |
| <b>APOE <math>\epsilon 4</math></b> |                           |                      |                                              |                      |                                                 |                      |
| Carrier                             | −0.63 (−1.18 to −0.08)    | .03                  |                                              |                      |                                                 |                      |
| Noncarrier                          | Reference                 |                      |                                              |                      |                                                 |                      |
| <b>Sex</b>                          |                           |                      |                                              |                      |                                                 |                      |
| Female                              | 0.35 (−0.15 to 0.85)      | .17                  | 0.64 (−0.21 to 1.5)                          | .14                  | 0.19 (−0.42 to 0.79)                            | .55                  |
| Male                                | Reference                 |                      | Reference                                    |                      | Reference                                       |                      |
| Age                                 | −0.001 (−0.05 to 0.05)    | .96                  | 0.02 (−0.06 to 0.1)                          | .65                  | −0.01 (−0.07 to 0.05)                           | .7                   |
| Education                           | −0.01 (−0.07 to 0.04)     | .66                  | −0.06 (−0.15 to 0.03)                        | .2                   | <0.001 (−0.07 to 0.07)                          | .99                  |
| Disease duration                    | 0.001 (−0.004 to 0.01)    | .76                  | 0.005 (−0.003 to 0.01)                       | .2                   | −0.001 (−0.008 to 0.005)                        | .72                  |
| Baseline MMSE                       | 0.95 (0.9 to 0.99)        | <.001                | 0.97 (0.9 to 1)                              | <.001                | 0.93 (0.87 to 1)                                | <.001                |
| Time                                | −1.17 (−1.4 to −1)        | <.001                | −1.34 (−1.65 to −1.03)                       | <.001                | −1.08 (−1.3 to −0.86)                           | <.001                |
| Medication                          | −0.46 (−0.95 to 0.03)     | .07                  | −0.85 (−1.77 to 0.07)                        | .07                  | −0.18 (−0.77 to 0.4)                            | .57                  |
| Hospitalization rates               | −0.24 (−0.66 to 0.17)     | .25                  | −0.6 (−1.2 to 0.004)                         | .05                  | −0.18 (−0.71 to 0.34)                           | .49                  |

Abbreviations: AD, Alzheimer disease; APOE, apolipoprotein E; CI, confidence interval; MMSE, Mini-Mental State Examination; NMDA, N-methyl-D-aspartate; VRF, vascular risk factor.

<sup>a</sup>P values were adjusted for APOE  $\epsilon 4$  carrier status (in all AD patients), sex, age, years of education, disease duration, baseline MMSE score, time, use of medication (acetylcholinesterase inhibitors or NMDA receptor antagonists), and hospitalization rates.

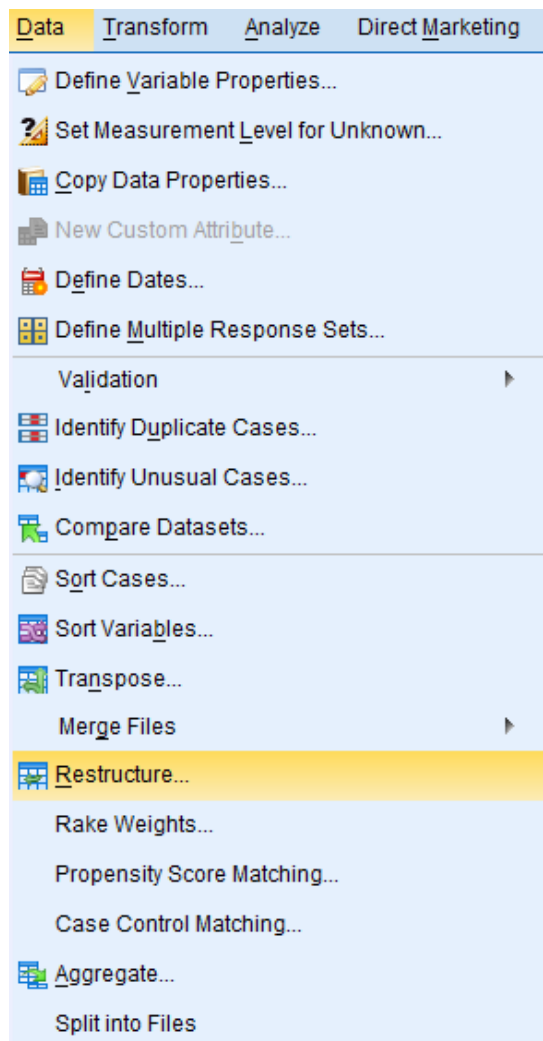
# GEE/LMM-資料轉置處理

| id | Gender | Treatment | Age | DepressionScale_1 | DepressionScale_2 | DepressionScale_3 |
|----|--------|-----------|-----|-------------------|-------------------|-------------------|
| 1  | 2      | 1         | 29  | 4.0               | 32.0              | 13.0              |
| 2  | 2      | 1         | 32  | 29.0              | 14.0              | 8.0               |
| 3  | 1      | 1         | 32  | 19.0              | 27.0              | 9.0               |
| 4  | 2      | 2         | 35  | 19.0              | 29.0              | 33.0              |
| 5  | 1      | 2         | 31  | 20.0              | 27.0              | 9.0               |
| 6  | 2      | 1         | 33  | 8.0               | 12.0              | 12.0              |
| 7  | 1      | 2         | 34  | 20.0              | 2.0               | 19.0              |
| 8  | 1      | 1         | 34  | 23.0              | 6.0               | 23.0              |
| 9  | 2      | 1         | 29  | 9.0               | .0                | .0                |
| 10 | 1      | 1         | 34  | 4.0               | 33.0              | 10.0              |
| 11 | 2      | 1         | 25  | 24.0              | 15.0              | 10.0              |
| 12 | 2      | 2         | 34  | 10.0              | 31.0              | 31.0              |
| 13 | 1      | 2         | 26  | 26.0              | 27.0              | 8.0               |

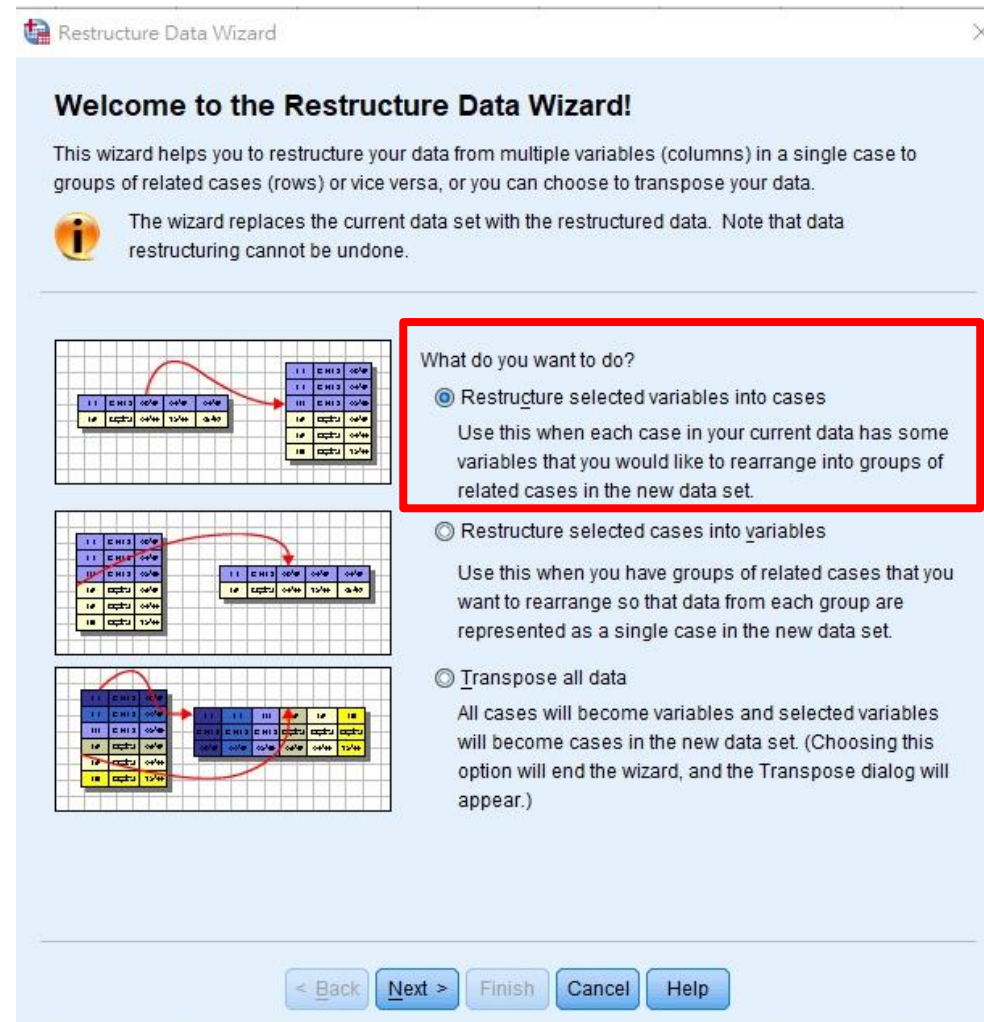


| id | Gender | Treatment | Age | Visit | DepressionScale |
|----|--------|-----------|-----|-------|-----------------|
| 1  | 2      | 1         | 29  | 1     | 4.0             |
| 1  | 2      | 1         | 29  | 2     | 32.0            |
| 1  | 2      | 1         | 29  | 3     | 13.0            |
| 1  | 2      | 1         | 29  | 4     | 19.0            |
| 1  | 2      | 1         | 29  | 5     | 11.0            |
| 2  | 2      | 1         | 32  | 1     | 29.0            |
| 2  | 2      | 1         | 32  | 2     | 14.0            |
| 2  | 2      | 1         | 32  | 3     | 8.0             |
| 2  | 2      | 1         | 32  | 4     | 5.0             |
| 2  | 2      | 1         | 32  | 5     | 4.0             |
| 3  | 1      | 1         | 32  | 1     | 19.0            |
| 3  | 1      | 1         | 32  | 2     | 27.0            |
| 3  | 1      | 1         | 32  | 3     | 9.0             |
| 3  | 1      | 1         | 32  | 4     | 15.0            |

## 資料 > 重新架構



# GEE/LMM-資料轉置處理



# GEE/LMM-資料轉置處理

Restructure Data Wizard - Step 2 of 7

## Variables to Cases: Number of Variable Groups

You have chosen to restructure selected variables into groups of related cases in the new file.

A group of related variables, called a variable group, represents measurements on one variable.

For example, the variable may be width. If it is recorded in three separate measurements, each one representing a different point in time—w1, w2, and w3, then the data are arranged in a group of variables.

If there is more than one variable in the file often it is also recorded in a variable group, for example height, recorded in h1, h2, and h3.

How many variable groups do you want to restructure?

☒ One (for example, w1, w2, and w3)

☐ More than one (for example, w1, w2, w3 and h1, h2, h3, etc.)

How Many?

< Back Next > Finish Cancel Help

Restructure Data Wizard - Step 3 of 7

## Variables to Cases: Select Variables

For each variable group you have in the current data the restructured file will have one target variable.

In this step, choose how to identify case groups in the restructured data, and choose which variables belong with each target variable.

Optionally, you can also choose variables to copy to the new file as Fixed Variables.

Variables in the Current File:

- id
- 性別 [Gender]
- 治療方式 [Treatment]
- 年齡 [Age]
- DepressionScale\_1
- DepressionScale\_2
- DepressionScale\_3
- DepressionScale\_4
- DepressionScale\_5

Case Group Identification

Use case number

Name:  Label...

Variables to be Transposed

Target Variable:  1

DepressionScale\_1 2

DepressionScale\_2

DepressionScale\_3

DepressionScale\_4

Fixed Variable(s):

id 3

性別 [Gender]

治療方式 [Treatment]

< Back Next > Finish Cancel Help

# GEE/LMM-資料轉置處理

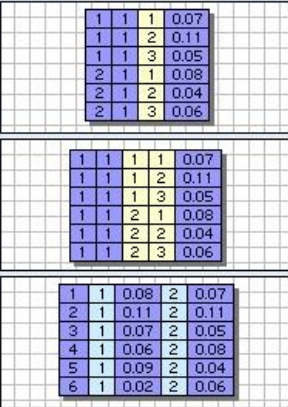
Restructure Data Wizard - Step 4 of 7

## Variables to Cases: Create Index Variables

In the current data, values for a variable group appear in a single case in multiple variables. For example, a single case contains the values for w1, w2, and w3.

In the new data, values for a variable group will appear in multiple cases in a single variable. For example, there will be three cases, one each for w1, w2, and w3.

An index is a new variable that identifies the group of new cases that was created from the original case. For example, an index named "w" would have the values 1, 2, and 3.



How many index variables do you want to create?

☒ **One**  
Use this when a variable group records the effects of a single factor, treatment or condition.

☐ **More than one**    How many?   
Use this when a variable group records the effects of more than one factor, treatment or condition.

☐ **None**  
Use this if index information is stored in one of the sets of variables to be transposed.

Restructure Data Wizard - Step 5 of 7

## Variables to Cases: Create One Index Variable

You have chosen to create one index variable. The variable's values can be sequential numbers or the names of variables in a group.

In the table you can specify the name and label for the index variable.

What kind of index values?

☒ **Sequential numbers**  
Index Values: 1, 2, 3, 4, 5

☐ **Variable names**  
Index Values: DepressionScale\_1, DepressionScale\_2, DepressionScale\_3, Depr...

Edit the Index Variable Name and Label:

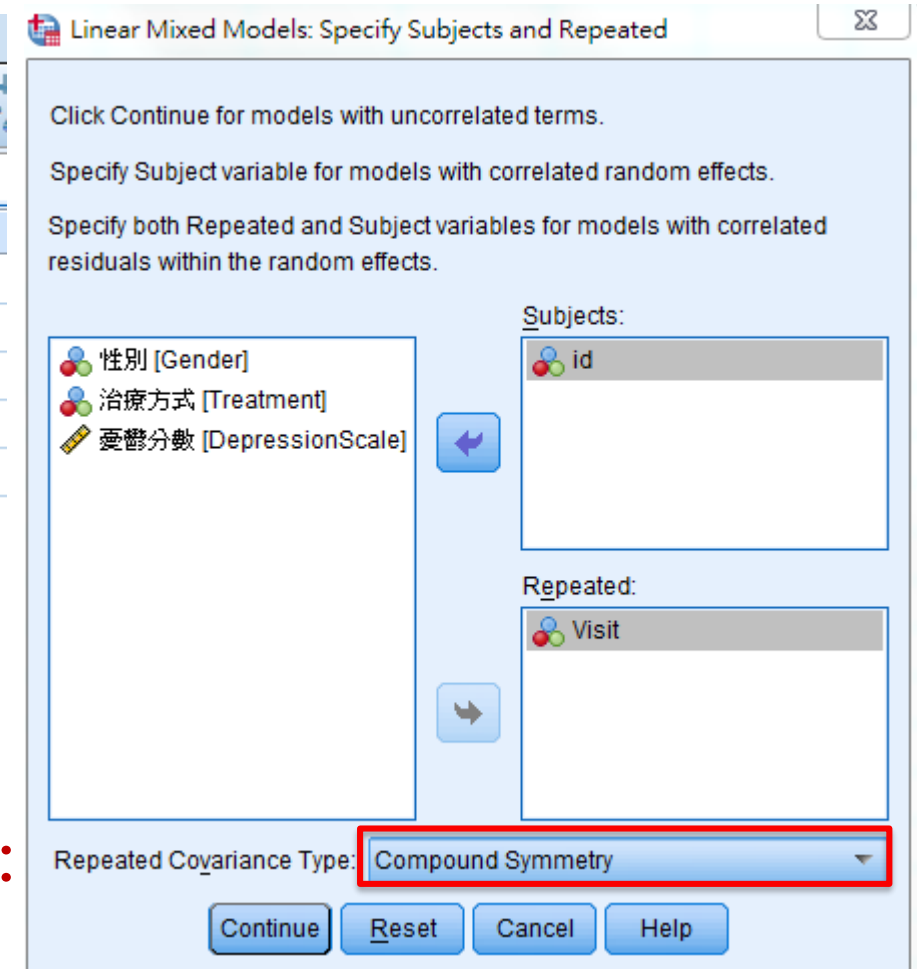
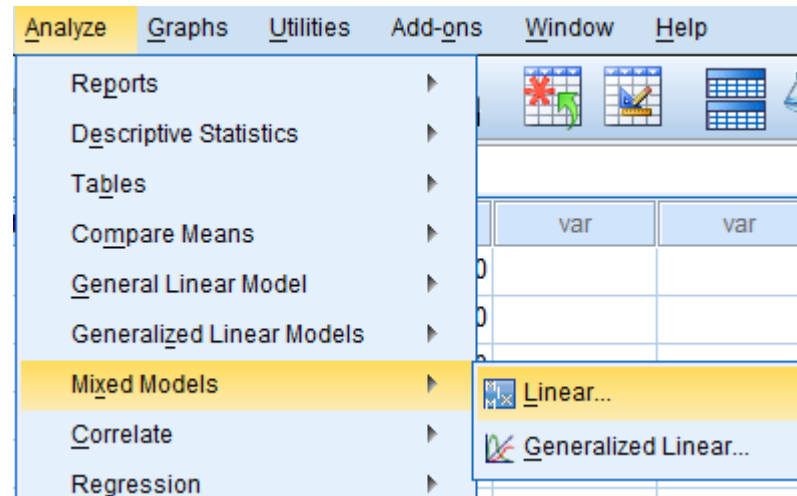
|   | Name  | Label | Levels | Index Values  |
|---|-------|-------|--------|---------------|
| 1 | Visit |       | 5      | 1, 2, 3, 4, 5 |

# Mixed model

| id | Gender | Treatment | Visit | DepressionScale |
|----|--------|-----------|-------|-----------------|
| 1  | 2      | 1         | 1     | 4.0             |
| 1  | 2      | 1         | 2     | 32.0            |
| 1  | 2      | 1         | 3     | 13.0            |
| 1  | 2      | 1         | 4     | 19.0            |
| 1  | 2      | 1         | 5     | 11.0            |
| 2  | 2      | 1         | 1     | 29.0            |
| 2  | 2      | 1         | 2     | 14.0            |
| 2  | 2      | 1         | 3     | 8.0             |
| 2  | 2      | 1         | 4     | 5.0             |
| 2  | 2      | 1         | 5     | 4.0             |
| 3  | 1      | 1         | 1     | 19.0            |
| 3  | 1      | 1         | 2     | 27.0            |
| 3  | 1      | 1         | 3     | 9.0             |
| 3  | 1      | 1         | 4     | 15.0            |
| 3  | 1      | 1         | 5     | 9.0             |

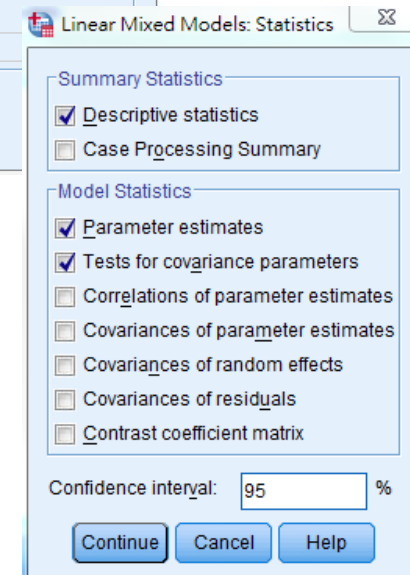
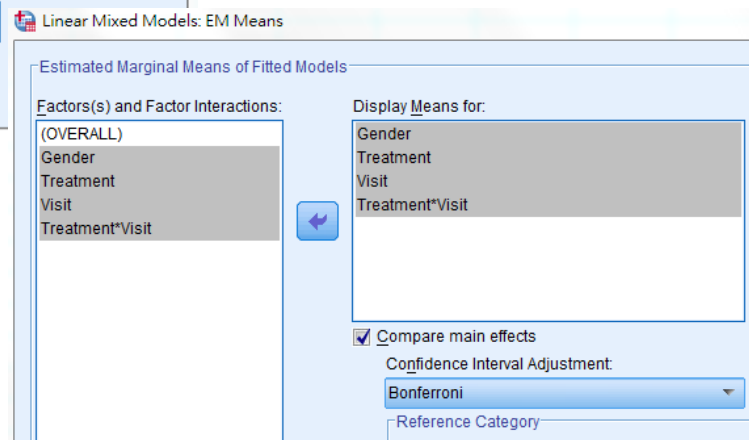
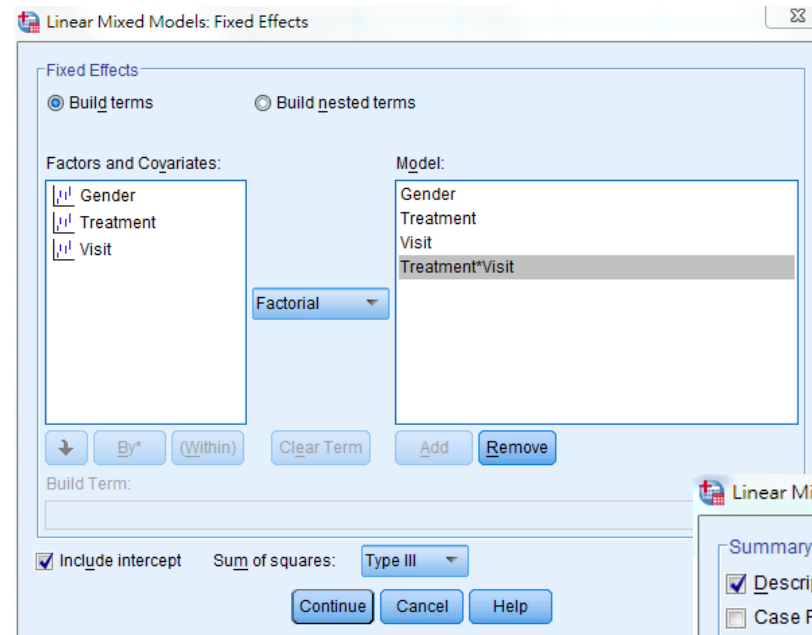
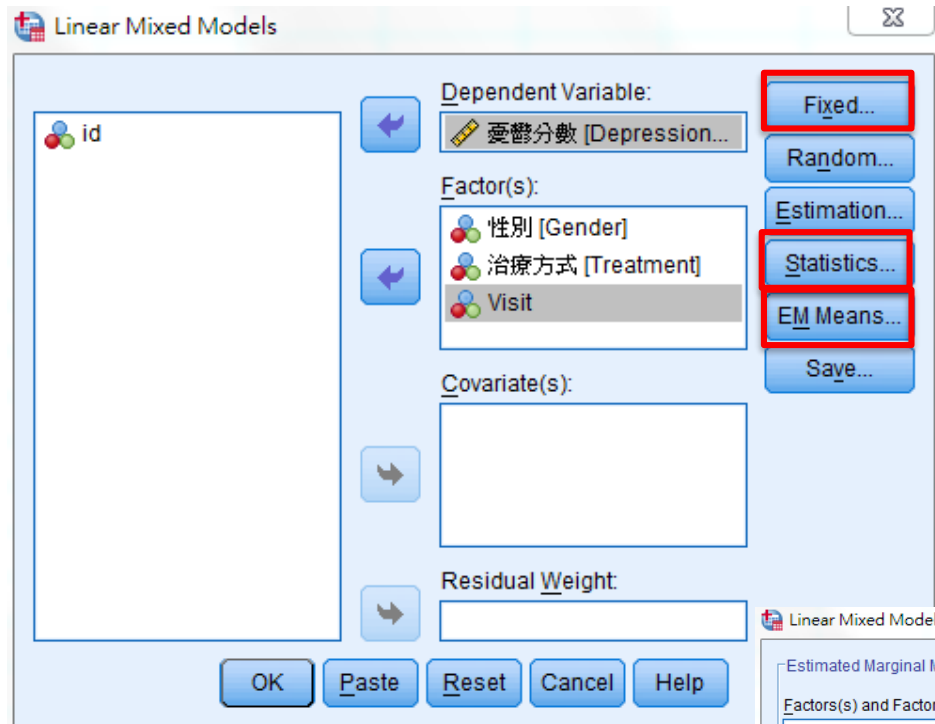
# Mixed model analysis

分析 > 混合模式 > 線性



重複共變異數類型：  
複合對稱  
(選取AIC及BIC最小)

# Mixed model analysis



Information Criteria<sup>a</sup>

|                                      |         |
|--------------------------------------|---------|
| -2 Restricted Log Likelihood         | 650.289 |
| Akaike's Information Criterion (AIC) | 654.289 |
| Hurvich and Tsai's Criterion (AICC)  | 654.433 |
| Bozdogan's Criterion (CAIC)          | 661.198 |
| Schwarz's Bayesian Criterion (BIC)   | 659.198 |

The information criteria are displayed in smaller-is-better form.

a. Dependent Variable: 憂鬱分數

Pairwise Comparisons<sup>a</sup>

| (I) Visit | (J) Visit | Mean Difference (I-J) | Std. Error | df     | Sig. <sup>b</sup> | 95% Confidence Interval for Difference <sup>b</sup> |             |
|-----------|-----------|-----------------------|------------|--------|-------------------|-----------------------------------------------------|-------------|
|           |           |                       |            |        |                   | Lower Bound                                         | Upper Bound |
| 1         | 2         | -2.712                | 2.741      | 68.882 | 1.000             | -10.662                                             | 5.238       |
|           | 3         | 2.429                 | 2.741      | 68.882 | 1.000             | -5.521                                              | 10.380      |
|           | 4         | 3.793                 | 2.741      | 68.882 | 1.000             | -4.157                                              | 11.743      |
|           | 5         | 3.567                 | 2.868      | 70.166 | 1.000             | -4.744                                              | 11.879      |
| 2         | 1         | 2.712                 | 2.741      | 68.882 | 1.000             | -5.238                                              | 10.662      |
|           | 3         | 5.141                 | 2.741      | 68.882 | .649              | -2.809                                              | 13.092      |
|           | 4         | 6.505                 | 2.741      | 68.882 | .204              | -1.445                                              | 14.455      |
|           | 5         | 6.279                 | 2.868      | 70.166 | .319              | -2.032                                              | 14.591      |
| 3         | 1         | -2.429                | 2.741      | 68.882 | 1.000             | -10.380                                             | 5.521       |
|           | 2         | -5.141                | 2.741      | 68.882 | .649              | -13.092                                             | 2.809       |
|           | 4         | 1.364                 | 2.741      | 68.882 | 1.000             | -6.587                                              | 9.314       |
|           | 5         | 1.138                 | 2.868      | 70.166 | 1.000             | -7.174                                              | 9.450       |
| 4         | 1         | -3.793                | 2.741      | 68.882 | 1.000             | -11.743                                             | 4.157       |
|           | 2         | -6.505                | 2.741      | 68.882 | .204              | -14.455                                             | 1.445       |
|           | 3         | -1.364                | 2.741      | 68.882 | 1.000             | -9.314                                              | 6.587       |
|           | 5         | -.226                 | 2.868      | 70.166 | 1.000             | -8.537                                              | 8.086       |
| 5         | 1         | -3.567                | 2.868      | 70.166 | 1.000             | -11.879                                             | 4.744       |
|           | 2         | -6.279                | 2.868      | 70.166 | .319              | -14.591                                             | 2.032       |
|           | 3         | -1.138                | 2.868      | 70.166 | 1.000             | -9.450                                              | 7.174       |
|           | 4         | .226                  | 2.868      | 70.166 | 1.000             | -8.086                                              | 8.537       |

Based on estimated marginal means

a. Dependent Variable: 憂鬱分數.

# Mixed model output

Estimates of Fixed Effects<sup>a</sup>

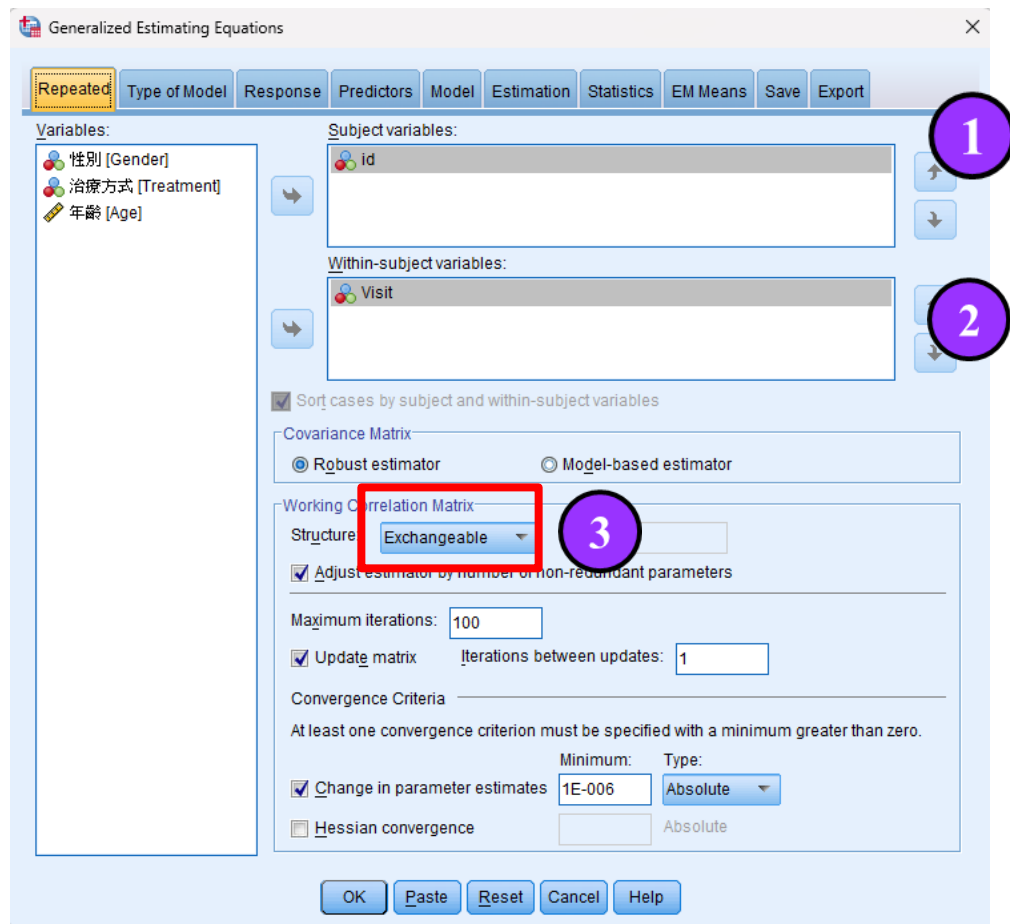
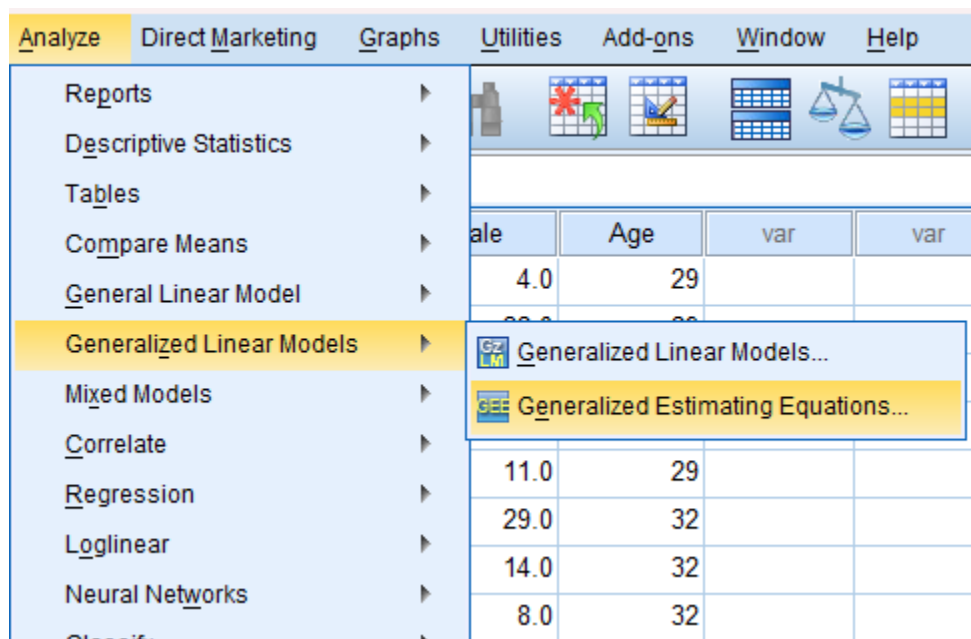
| Parameter                 | Estimate       | Std. Error | df     | t     | Sig. | 95% Confidence Interval |             |
|---------------------------|----------------|------------|--------|-------|------|-------------------------|-------------|
|                           |                |            |        |       |      | Lower Bound             | Upper Bound |
| Intercept                 | 10.457603      | 3.313212   | 74.672 | 3.156 | .002 | 3.856871                | 17.058335   |
| [Gender=1]                | 2.029279       | 2.351907   | 16.727 | .863  | .400 | -2.938983               | 6.997541    |
| [Gender=2]                | 0 <sup>b</sup> | 0          | .      | .     | .    | .                       | .           |
| [Treatment=0]             | 3.590407       | 4.499750   | 81.501 | .798  | .427 | -5.361851               | 12.542665   |
| [Treatment=1]             | 0 <sup>b</sup> | 0          | .      | .     | .    | .                       | .           |
| [Visit=1]                 | 3.529088       | 3.898315   | 70.588 | .905  | .368 | -4.244717               | 11.302894   |
| [Visit=2]                 | 3.619997       | 3.898315   | 70.588 | .929  | .356 | -4.153808               | 11.393803   |
| [Visit=3]                 | -2.107276      | 3.898315   | 70.588 | -.541 | .591 | -9.881081               | 5.666530    |
| [Visit=4]                 | .165452        | 3.898315   | 70.588 | .042  | .966 | -7.608354               | 7.939257    |
| [Visit=5]                 | 0 <sup>b</sup> | 0          | .      | .     | .    | .                       | .           |
| [Visit=1] * [Treatment=0] | .076556        | 5.731167   | 70.319 | .013  | .989 | -11.352986              | 11.506098   |
| [Visit=2] * [Treatment=0] | 5.318980       | 5.731167   | 70.319 | .928  | .357 | -6.110562               | 16.748522   |
| [Visit=3] * [Treatment=0] | 6.490697       | 5.731167   | 70.319 | 1.133 | .261 | -4.938845               | 17.920239   |
| [Visit=4] * [Treatment=0] | -.782030       | 5.731167   | 70.319 | -.136 | .892 | -12.211572              | 10.647512   |
| [Visit=5] * [Treatment=0] | 0 <sup>b</sup> | 0          | .      | .     | .    | .                       | .           |
| [Visit=1] * [Treatment=1] | 0 <sup>b</sup> | 0          | .      | .     | .    | .                       | .           |
| [Visit=2] * [Treatment=1] | 0 <sup>b</sup> | 0          | .      | .     | .    | .                       | .           |
| [Visit=3] * [Treatment=1] | 0 <sup>b</sup> | 0          | .      | .     | .    | .                       | .           |
| [Visit=4] * [Treatment=1] | 0 <sup>b</sup> | 0          | .      | .     | .    | .                       | .           |
| [Visit=5] * [Treatment=1] | 0 <sup>b</sup> | 0          | .      | .     | .    | .                       | .           |

a. Dependent Variable: 憂鬱分數.

b. This parameter is set to zero because it is redundant.

# GEE analysis

## 分析 > 廣義線性模型 > 廣義估計方程式(GEE)



# GEE analysis

Generalized Estimating Equations

Repeated Type of Model Response Predictors Model Estimation Statistics EM Means Save Export

Choose one of the model types listed below or specify a custom combination of distribution and link function.

Scale Response

☒ Linear **Continuous outcome**

☐ Gamma with log link

Ordinal Response

☐ Ordinal logistic

☐ Ordinal probit

Counts

☐ Poisson loglinear

☐ Negative binomial with log link

Mixture

☐ Tweedie with log link

☐ Tweedie with identity link

Custom

☐ Custom

Binary Response or Events/Trials Data

☐ Binary logistic **Binary outcome**

☐ Binary probit

☐ Interval censored survival

Distribution: Normal Link function: Identity

Parameter

☒ Specify value

Value: 1

☐ Estimate value

Power:

OK Paste Reset Cancel Help

Generalized Estimating Equations

Repeated Type of Model Response Predictors Model Estimation Statistics EM Means Save Export

Variables:

- id
- 性別 [Gender]
- 治療方式 [Treatment]
- Visit
- 年齢 [Age]

Dependent Variable

☒ Dependent Variable: 憂鬱分数 [DepressionScale]

Category order (multinomial only): Ascending

Type of Dependent Variable (Binomial Distribution Only)

☒ Binary

Reference Category...

☒ Number of events occurring in a set of trials

Trials

☒ Variable

Trials Variable:

☐ Fixed value

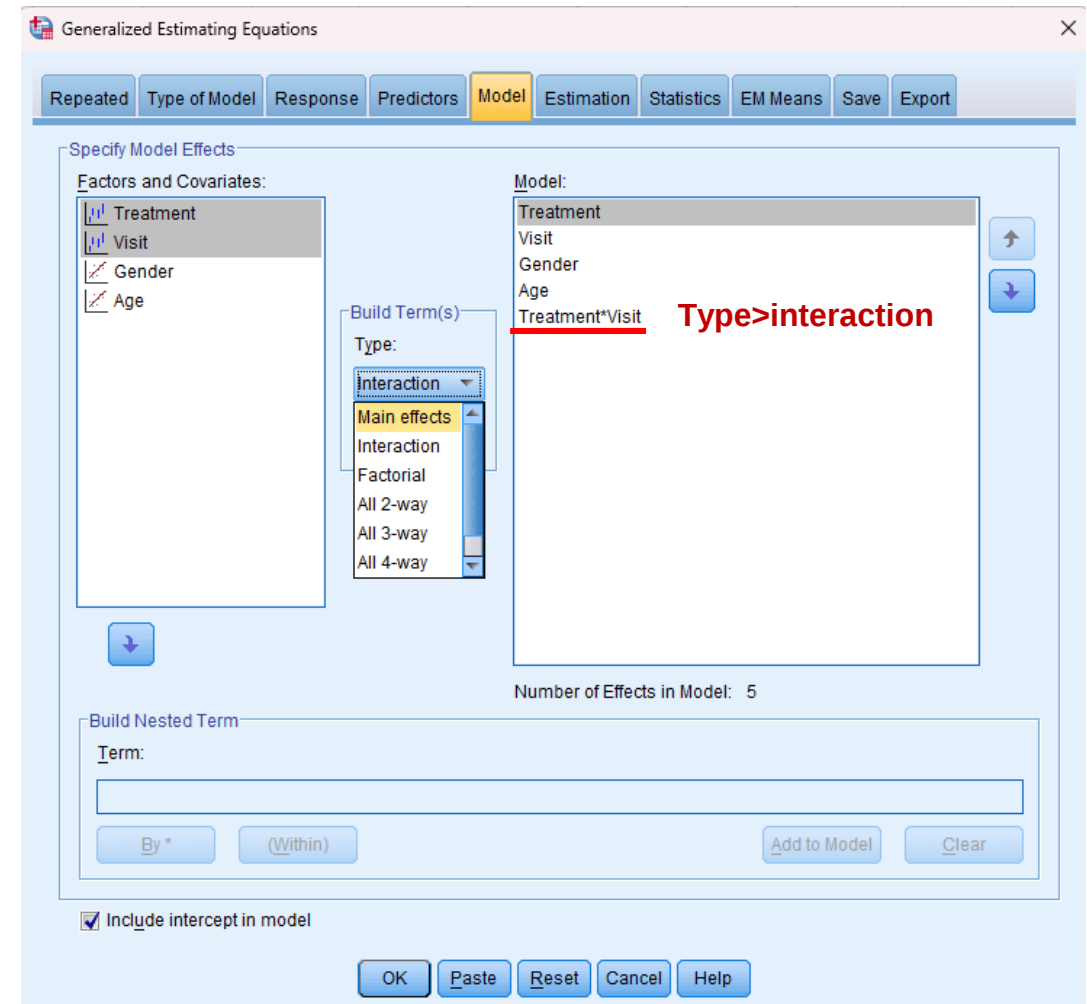
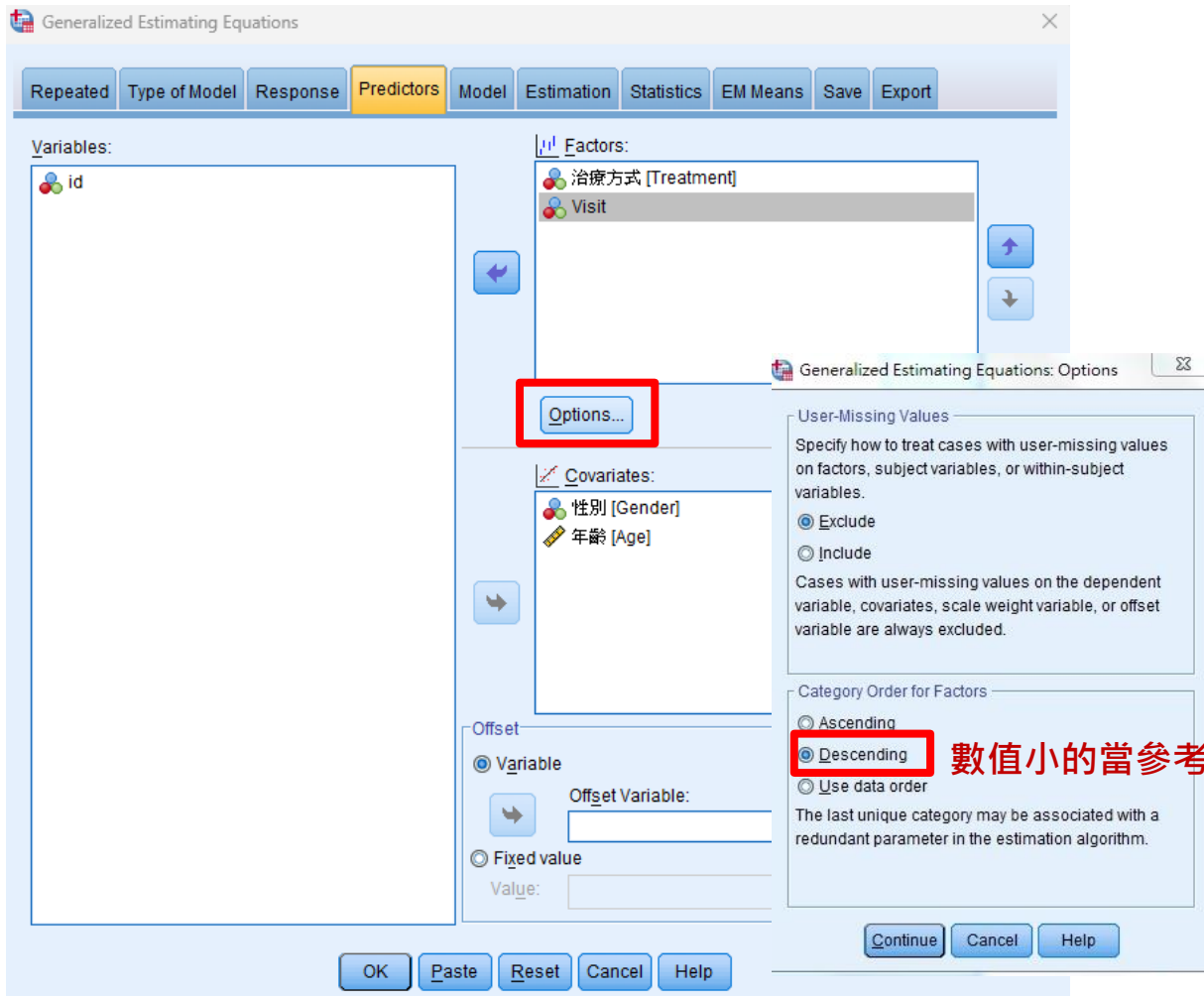
Number of Trials:

Scale Weight

☒ Scale Weight Variable:

OK Paste Reset Cancel Help

# GEE analysis



# GEE analysis

Generalized Estimating Equations

Repeated Type of Model Response Predictors Model Estimation **Statistics** EM Means Save Export

Model Effects

Analysis Type: Type III Confidence Interval Level (%): 95

Chi-square Statistics

☒ Wald  
☐ Generalized score

Log quasi-likelihood function: Full

Print

☒ Case processing summary  
☒ Descriptive statistics  
☒ Model information  
☒ Goodness of fit statistics  
☒ Model summary statistics  
☒ Parameter estimates  
☐ Include exponential parameter estimates  
☐ Covariance matrix for parameter estimates  
☐ Correlation matrix for parameter estimates  
☐ Working correlation matrix

☐ Contrast coefficient (L) matrices  
☐ General estimable functions  
☐ Iteration history  
Print Interval: 1

OK Paste Reset Cancel Help

若為binary outcome需勾選

Generalized Estimating Equations

Repeated Type of Model Response Predictors Model Estimation Statistics **EM Means** Save Export

Factors and Interactions:

| M |                                     | Term            |
|---|-------------------------------------|-----------------|
|   | <input checked="" type="checkbox"/> | Treatment       |
|   | <input checked="" type="checkbox"/> | Visit           |
|   | <input checked="" type="checkbox"/> | Treatment*Visit |

Display Means for:

| Term            | Contrast | Reference Category |
|-----------------|----------|--------------------|
| Visit           | Pairwise |                    |
| Treatment*Visit | Pairwise |                    |

By \*

Scale

☒ Compute means for response  
☐ Compute means for linear predictor

Adjustment for Multiple Comparisons:  
Least significant difference

☐ Display overall estimated mean

OK Paste Reset Cancel Help

# GEE-Output

## Model Information

|                                      |              |
|--------------------------------------|--------------|
| Dependent Variable                   | 憂鬱分數         |
| Probability Distribution             | Normal       |
| Link Function                        | Identity     |
| Subject Effect                       | 1            |
| Within-Subject Effect                | 1            |
| Working Correlation Matrix Structure | Exchangeable |

## Case Processing Summary

|          | N   | Percent |
|----------|-----|---------|
| Included | 97  | 97.0%   |
| Excluded | 3   | 3.0%    |
| Total    | 100 | 100.0%  |

## Categorical Variable Information

|        |       |           | N      | Percent |
|--------|-------|-----------|--------|---------|
| Factor | 治療方式  | Placebo   | 44     | 45.4%   |
|        |       | Treatment | 53     | 54.6%   |
| Total  |       | 97        | 100.0% |         |
| Visit  | 5     | 17        | 17.5%  |         |
|        | 4     | 20        | 20.6%  |         |
|        | 3     | 20        | 20.6%  |         |
|        | 2     | 20        | 20.6%  |         |
|        | 1     | 20        | 20.6%  |         |
|        | Total | 97        | 100.0% |         |

## Goodness of Fit<sup>a</sup>

|                                                                                   | Value    |
|-----------------------------------------------------------------------------------|----------|
| Quasi Likelihood under Independence Model Criterion (QIC) <sup>b</sup>            | 7393.042 |
| Corrected Quasi Likelihood under Independence Model Criterion (QICC) <sup>b</sup> | 7389.820 |

Dependent Variable: 憂鬱分數  
Model: (Intercept), Treatment, Visit, Gender, Age, Treatment \* Visit

a. Information criteria are in smaller-is-better form.

b. Computed using the full log quasi-likelihood function.

QIC/QICC數值越小越好

## Parameter Estimates

| Parameter                 | B              | Std. Error | 95% Wald Confidence Interval |        | Hypothesis Test |    |      |
|---------------------------|----------------|------------|------------------------------|--------|-----------------|----|------|
|                           |                |            | Lower                        | Upper  | Wald Chi-Square | df | Sig. |
| (Intercept)               | 23.711         | 17.2393    | -10.078                      | 57.499 | 1.892           | 1  | .169 |
| [Treatment=2]             | 3.616          | 3.3682     | -2.986                       | 10.218 | 1.153           | 1  | .283 |
| [Treatment=1]             | 0 <sup>a</sup> | .          | .                            | .      | .               | .  | .    |
| [Visit=5]                 | -3.502         | 3.4075     | -10.180                      | 3.177  | 1.056           | 1  | .304 |
| [Visit=4]                 | -3.364         | 3.7090     | -10.633                      | 3.906  | .822            | 1  | .364 |
| [Visit=3]                 | -5.636         | 2.7732     | -11.072                      | -.201  | 4.131           | 1  | .042 |
| [Visit=2]                 | .091           | 4.6859     | -9.093                       | 9.275  | .000            | 1  | .985 |
| [Visit=1]                 | 0 <sup>a</sup> | .          | .                            | .      | .               | .  | .    |
| Gender                    | -2.306         | 2.2536     | -6.723                       | 2.111  | 1.047           | 1  | .306 |
| Age                       | -.166          | .4758      | -1.099                       | .767   | .122            | 1  | .727 |
| [Treatment=2] * [Visit=5] | -.113          | 4.8978     | -9.712                       | 9.487  | .001            | 1  | .982 |
| [Treatment=2] * [Visit=4] | -.859          | 5.5842     | -11.803                      | 10.086 | .024            | 1  | .878 |
| [Treatment=2] * [Visit=3] | 6.414          | 5.4913     | -4.349                       | 17.177 | 1.364           | 1  | .243 |
| [Treatment=2] * [Visit=2] | 5.242          | 6.0191     | -6.555                       | 17.040 | .759            | 1  | .384 |
| [Treatment=2] * [Visit=1] | 0 <sup>a</sup> | .          | .                            | .      | .               | .  | .    |
| [Treatment=1] * [Visit=5] | 0 <sup>a</sup> | .          | .                            | .      | .               | .  | .    |
| [Treatment=1] * [Visit=4] | 0 <sup>a</sup> | .          | .                            | .      | .               | .  | .    |
| [Treatment=1] * [Visit=3] | 0 <sup>a</sup> | .          | .                            | .      | .               | .  | .    |
| [Treatment=1] * [Visit=2] | 0 <sup>a</sup> | .          | .                            | .      | .               | .  | .    |
| [Treatment=1] * [Visit=1] | 0 <sup>a</sup> | .          | .                            | .      | .               | .  | .    |
| (Scale)                   | 86.657         |            |                              |        |                 |    |      |

Dependent Variable: 憂鬱分數

Model: (Intercept), Treatment, Visit, Gender, Age, Treatment \* Visit

a. Set to zero because this parameter is redundant.

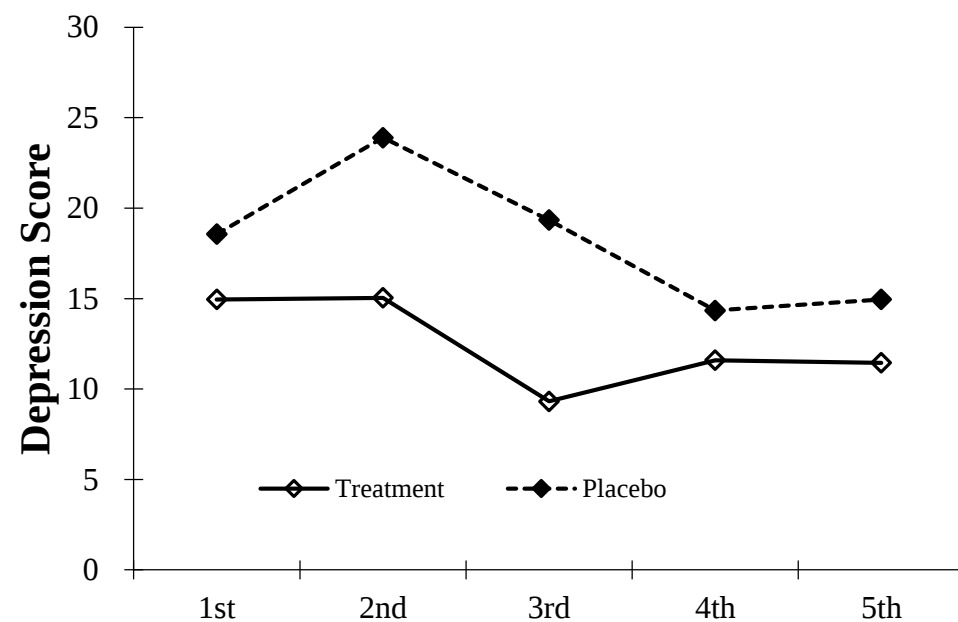
# GEE-Output

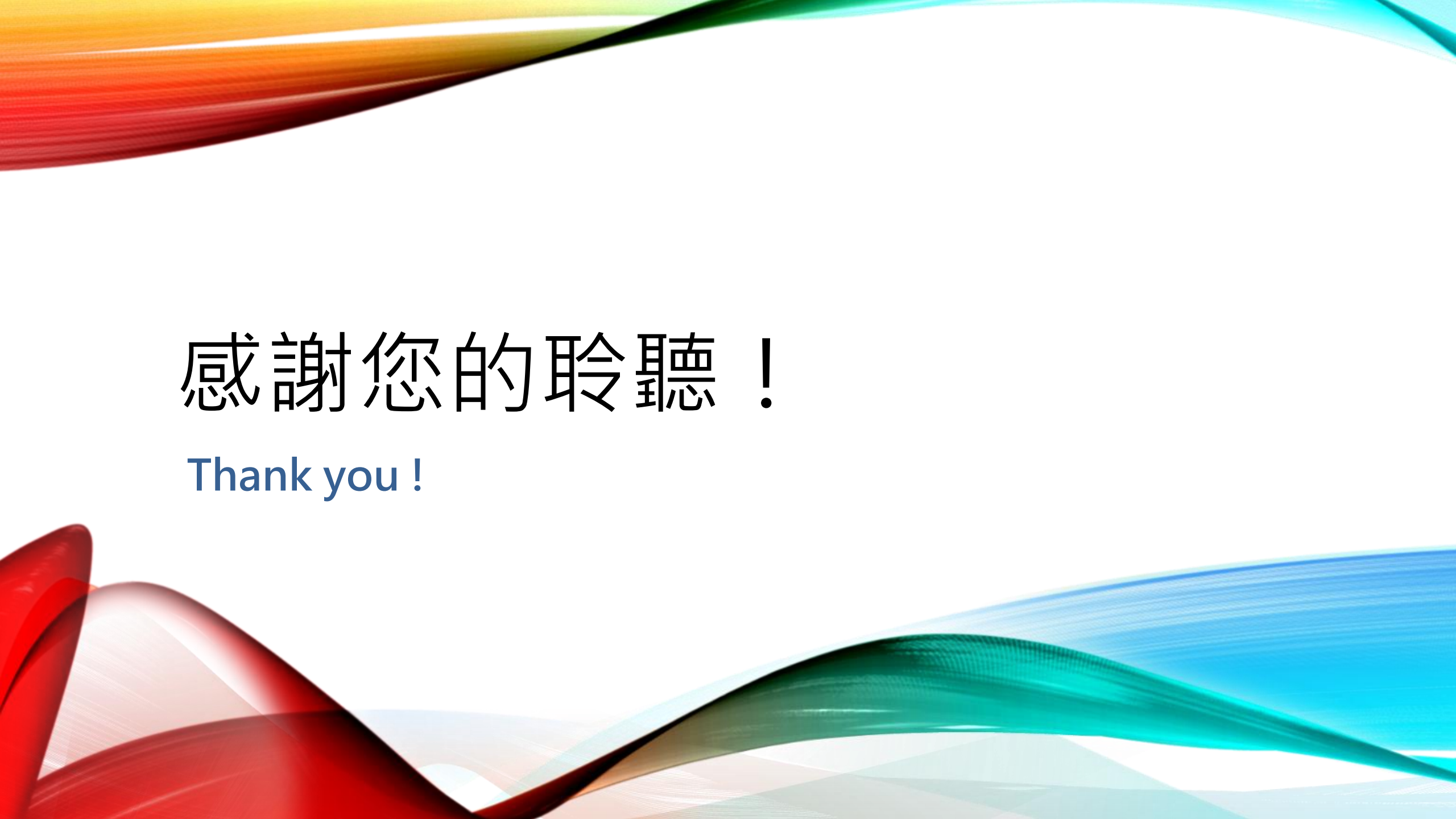
Estimates

| 治療方式      | Visit | Mean   | Std. Error | 95% Wald Confidence Interval |         |
|-----------|-------|--------|------------|------------------------------|---------|
|           |       |        |            | Lower                        | Upper   |
| Placebo   | 5     | 14.953 | 45.0687    | -73.380                      | 103.286 |
|           | 4     | 14.345 | 44.2540    | -72.391                      | 101.081 |
|           | 3     | 19.345 | 44.3947    | -67.667                      | 106.357 |
|           | 2     | 23.901 | 44.2926    | -62.911                      | 110.712 |
|           | 1     | 18.567 | 44.0893    | -67.846                      | 104.981 |
| Treatment | 5     | 11.449 | 42.0461    | -70.959                      | 93.858  |
|           | 4     | 11.587 | 43.0089    | -72.708                      | 95.883  |
|           | 3     | 9.315  | 43.0638    | -75.089                      | 93.718  |
|           | 2     | 15.042 | 43.0606    | -69.355                      | 99.439  |
|           | 1     | 14.951 | 42.8486    | -69.031                      | 98.933  |

Covariates appearing in the model are fixed at the following values:

Gender=1.54; Age=31.42





感謝您的聆聽！

Thank you !