# 行政院國家科學委員會專題研究計畫 成果報告

# 巴金森病人之情緒功能:長期追蹤研究(111)

<u>計畫類別</u>: 個別型計畫 <u>計畫編號</u>: NSC91-2413-H-002-008-<u>執行期間</u>: 91 年 08 月 01 日至 92 年 07 月 31 日 執行單位: 國立臺灣大學心理學系暨研究所

## <u>計畫主持人:</u>花茂棽

共同主持人: 吳瑞美, 吳逸如, 陳獻宗

## 報告類型:精簡報告

<u>處理方式:</u>本計畫可公開查詢

## 中 華 民 國 92 年 11 月 7 日

巴金森病是一種常見中樞神經系統的退化疾病。運動障礙、認知功能缺損、 以及情緒功能失調,特別憂鬱問題是主要的症狀。但是文獻上所呈現這類患者罹 患憂鬱與焦慮問題的盛行率,卻是不大一致。本研究計劃以三年的時間,利用追 蹤方式來探討這個爭議未決的問題。

完成第三年研究計劃,總共收集到 192 位成年受試參與本研究,其中包括 127 位巴金森患者和 65 位正常受試。依照侯-葉氏(Hoehn-Yahr,1967)運動障礙評量 表 127 患者中,58 位屬於第一級輕度運動障礙、52 位第二級輕中度運動障礙、 以及 18 位第三級中度運動障礙.。每一位受試都分別接受一組認知功能之神經心 理測驗,以及一組情緒功能之測量。受試之主要照顧者同時也接受一組情緒功能 之測量。完成第一年追蹤研究之受試者總共53人;其中21 人屬於第一級輕度運 動障礙、17 位第二級輕中度運動障礙、9 位第三級中度運動障礙,以及正常受試 者 6 位。完成第二年追蹤研究之受試者總共22 人;其中4 人屬於第一級輕度運 動障礙、3 位第二級輕中度運動障礙、5 位第三級中度運動障礙,以及正常受試 者 10 位。

初步資料分析之結果顯示巴金森病患呈現情緒功能失調的問題 主要包括了 憂鬱 焦慮 身心不適 強迫性行為等症狀。大約 40%的巴金森患有憂鬱的問題: 第一級輕度運動障礙患者有 43%患有這種情緒症狀、第二級輕中度運動障礙患 者有 30%、第三級中度運動障礙患者有 50%。這三組巴金森病患中約有 55%罹 患失智症狀群,不過只有40%的這些失智患者同時出現憂鬱症狀。完成第一年追 蹤研究之患者中,約有32%呈現憂鬱問題。第一級輕度運動障礙患者有24%、第 二級輕中度運動障礙患者有 35%、第三級中度運動障礙患者有 44%這種情緒症 狀。憂鬱症狀程度達到重憂鬱症(major depressive disorder)約有 15%;其中第一級 輕度運動障礙患者約 15%、第二級輕中度運動障礙患者 18%、第三級中度運動障 磁患者 11%出現該疾患。完成第一年追蹤研究之患者中約有 64%罹患失智症狀 群。完成第二年追蹤研究之患者中,約有50%呈現憂鬱問題。第一級輕度運動障 礙患者有 50%、第二級輕中度運動障礙患者有 100%、第三級中度運動障礙患者 有 33%這種情緒症狀。憂鬱症狀程度達到重憂鬱症(major depressive disorder)約 有 33%;其中第一級輕度運動運動障礙患者約 50%、第二級輕中度運動障礙患者 0%、第三級中度運動障礙患者 33%出現該疾患。完成第二年追蹤研究之患者中 約有 40%罹患失智症狀群,不過只有 50%的這些失智症患者同時出現憂鬱症狀。

關鍵詞:情緒狀態、巴金森病、單氨類化學物質失調

#### ABSTRACT

Parkinson's disease (PD) is a prevailing degenerative disease of central nervous system. Motor symptoms, cognitive impairments including dementia, and emotional disturbances, especially depression are the cardinal features of the disease. The prevalence rate of depression and anxiety in the parkinsonian patients has a great variation. The present 3-year follow-up study, thus, is designed to examine emotional function in these patients.

In this third-year study, we included a total of 192 adult participants, 127 patients with idiopathic PD and 65 normal controls. On the basis of the motor staging of Hoehn and Yahr (1967), Group 1 consisted of 58 patients with the staging I, Group 2 included 52 patients with the staging II, and Group 3 were composed of 18 patients with the staging III. Each subject received a series of non-emotional neuropsychological test battery, and emotional status measures. The significant informant of each subject also received a series of emotional function measures. 53 patients completed the first-year follow-up. Among these, 21 of them were rated as the staging I in terms of their motor disabilities; 17 the staging II; and 9 the staging III, as well as 6 normal counterparts. 22 of these subjects have received the second-year follow-up. Among them, 4 patients were rated as the staging I of Hoehn-Yahr Scale, 3 the staging II, and 5 the staging III, as well as 10 of normal control subjects.

Because of a high dropout rate, only 6 out of 12 patients completed the 2<sup>nd</sup>-year follow-up of emotional status examination. The preliminary results revealed that about 50% of our patients manifested depression symptom. Among them, 50% of patients with staging I, 100% with staging II, and 33% with staging III were evident of this symptom. About 33% suffered from Major Depressive Disorder in which 50%, 0%, and 33% with the motor stagings I-III respectively had this disorder. About 40% of our patients suffered from the syndrome of dementia based on the demented diagnostic criteria of the DSM-IV. However, about 50% of these demented patients were also evident of depression problem.

Keywords: Emotional Status, Parkinson's Disease, Monoaminergic Imbalance

#### INTRODUCTION

Review of the literature indicates that the prevalence rate of depression in the parkinsonian patients has a great variation, ranging from 4% to 70% with an average of around 40% (Cummings, 1992). This considerable variation is mainly attributable to methodological discrepancy, such as the divergent source patient populations, and various ways of depression diagnosis (e.g., based on the DSM system, semi-structured interview, or psychometric measures) (Sano, Marder & Dooneief, 1996). The investigators (Hoen & Yahr, 1967; Sano et al., 1989) reported about 51% of depression in PD based on clinic/hospital-based studies, and about 32% based on population-based investigations. Taken these data together, it implicates that at least about 50% of the patients with PD do not suffer from depression. Depression has been thought to be associated with a widespread monoaminergic dysfunction, and degeneration of serotonic and noradrenergic projections occurs alongside degeneration of dopaminergic neurons in the patients with PD (Beatty, 1995; Conn, 1995; Fibigier, 1984; Mayeux, 1990). Accordingly, it is expected that the patients with PD but without depression will be evident of depression gradually along with the neural degenerative processes. This issue, however, has been lacking in systematic investigation.

Otherwise emotional disturbances, such as anxiety, mania, hypomania, and psychosis characterized by paranoid delusions and hallucinations have also reported in the patients with PD (Iruela, Ibanez-Rojo, Palanca, & Caballero, 1992; Stein, Heuser, Juncos & Uhde, 1990). These problems, however, are rare and most of them were associated with the patient's taking antiparkinosnian medications (Celesia & Barr, 1970; Factor & Brown, 1992; Goodwin, 1971; Jouvent et al., 1983; Lang et al., 1982).

A prospective longitudinal study of emotional function changes, particularly for depression, in the patients with PD is limited. Using such a research design with one-year follow-up, Starkstein and his colleagues (1990, 1992) have investigated depression in the patients with PD. The results revealed that both depressed and non-depressed patients had a significant deterioration of cognitive function (the former was remarkably more severe than the latter), and about 18 % (10 out of the 55) of the non-depressed patients with PD were progressively evident of depression a year later. The findings seem to partially support the hypothesis of depression associated with a widespread monoaminergic imbalance in the patients with PD (Fibigier, 1984; Mayeux, 1990). Because these results were only based on a one-year follow-up, the hypothesis, however, deserves further investigation.

The report of emotional disturbances, particularly depression, in the patients with

PD in Taiwan is meager. In order to explore the aforementioned issues (i.e., the prevalence of depression and otherwise emotional disturbances, and the hypothesis of monoaminergic imbalance and its relation to depression and anxiety), and to document the literature in Taiwan, we design this three-year longitudinal study. In our study, 2 groups of participants, 1 cohort of parkinsonian patients and 1 group of normal controls will be included. The goal of the first-year study is to complete the initial and part of 1-year follow-up evaluation of the emotional status and cognitive function. The objective of the second-year study is to complete 1-year follow-up and part of 2-year follow-up evaluation of and cognitive function; and the third-year is to complete 2-year and part of 3-year follow-up evaluation of emotional status and cognitive function.

The specific aims of the study, thus, are to examine the following questions: 1) Is there an impairment of emotional status in the patients with PD? 2) If so, does the deficit only involve depression or also include otherwise emotional function evaluated? 3) If depression symptoms do occur in the patients, are they persistent in nature and can thus these patients be diagnosed as Major Depressive or Dysthymic Disorder? 4) If anxiety symptoms do occur in the patients, are they persistent in nature and then can these patients be classified as Generalized Anxiety Disorder? 5) If depression does occur, is it significantly associated with dementia? 6) If depression or anxiety does not occur in the patients with PD at first, will it be evident in company with the progression of the disease?

#### METHOD

**Participants.** A total of 192 adult participants, including three groups of patients with idiopathic PD with varying degree of motor disabilities and one normal control group, participated in the  $1^{st}$ - and  $2^{nd}$ -year study. Groups (Gps) were matched for age and educational level. Although 47 patients completed the 1st-year follow-up, only 12 patients completed the  $2^{nd}$ -year follow-up. The VIQ of Gps 4 and 1 were significantly higher than that of Gps 2 and 3 (Table 1). The diagnosis of PD was based on the Parkinson's Disease Society Brain Bank in London (PDSBB) guidelines (Fahn & Elton, 1987), and the patients with severe motor symptoms (staging 5) according to the criteria derived from Hoehn and Yahr (1967) were excluded in the study. All of the patients were also free of any other CNS, and psychiatric history.

All participants were right-handed in which hand dominance was ascertained by the history that the participant has always used his/her right hand preferentially for doing skillful activities, such as writing and holding chopsticks.

**Tests and Procedure**. After giving informed consent, each participant received a series of neuropsychological tests. These tests included following cognitive tests: the Temporal Orientation Test (Benton, Hamsher, Varney & Spreen, 1983), the

Orientation to Personal Information and Place (Hamsher, 1983), the Object Naming Test (Spreen & Benton, 1969), the Semantic Association of Verbal Fluency (Hua, 1987), the Token Test (Benton & Hamsher, 1978), the verbal subtests of the WAIS-R (Wechsler, 1981), the Judgment of Line Orientation (Benton et al., 1983), the Facial Recognition Test (Benton et al., 1983), the Word Sequence Learning (Hua, 1987), the Benton Visual Retention Test (Benton, 1974), the Wisconsin Card Sorting (Nelson, 1974), and the Trails Making A and B (Reitan & Wolfson, 1993). The battery also included the following emotional status measures: Symptom Checklist-90-R (Derogatis, 1977), Beck Depression Inventory (Beck, 1987), and a semi-structured Standard Neurobehavioral Interview Inventory (Hamsher, 1983). Evaluations were administered in an examining room between 11AM and 2 PM to minimize any possible effects of diurnal mood variation on interview response, as suggested by Starkstein and his colleagues (1992). All these measures are Chinese versions.

In order to obtain reliable and valid ratings of the patient's emotional status, we asked the significant informants, particularly family caregivers of the patient, to rate the patient's emotional function. All of the significant informants of the patients were free of emotional disturbances, dementia, and psychiatric history. Each participant received the above evaluation at around 1 week after the diagnosis of PD, and at 12, 24, and 36 (if possible) months following the first evaluation.

### RESULTS

Since there was a high dropout rate in the 2<sup>nd</sup>-year follow-up and only a small sample of subjects participated in the study, the non-parametric statistical procedures, Kruskall-Wallis one-way ANOVA, were used to analyze the test score or scale rating differences between the patients and normal controls. The post-hoc pairwise comparison procedure, Nemenyi's contrasts, was subsequently employed if H values reached a statistically significant level. On the non-emotional status measures, performance of normal control subjects on these tests overpowered that of patients in which performance of patients with motor staging I was better than that of patients with motor stagings II and III. that of their normal counterparts (Tables2-5). However, only the correct score differences of the Word Sequence Learning Test between normal control subjects and patients with motor staging III on the Word Sequence Learning reached a statistically significant level.

On the emotional status measures (Table 6), the patients' score on the Beck Depression Inventory was higher than that of their normal counterparts. The patients' rating scores on Somatic, Obsessive-Compulsive, Depressed Mood, and Anxiety Symptoms subscales on the SCL-90-R were higher than their normal counterparts. However, score differences of these tests were not statistically significant.

Concerning the ratings of the patients' and normal controls' significant

informants on the subjects' emotional status with the Beck Depression Inventory and the 4 subscales of SCL-90-R, the scores of the patients' significant informants were also higher than those of the normal controls (Table 7). However, these rating differences did not reach a statistically significant level. The ratings between the patients and their significant informants on both of the emotional scales were consistent.

The examiner's ratings of the four group subjects' emotional status based on the items 9, 13, 15, 17, and 21 (which are associated with depression), and items 4 and 25 (which are related to anxiety) of the Neurobehavioral Rating Scale Interview are indicated in Table 8. The results revealed that on the depressed mood and anxiety ratings, the mean score of Gps 1, 2 and 3 was higher than that of their normal counterparts. However, these differences did not reach a statistically significant level.

We used a preliminary cut-off point score of 12 or more; that is, 1.5 standard deviation above the mean score of the normal controls on the Beck Depression Inventory, to determine whether the patient had depressed mood. On the basis of this cut-off point score, we classified about 50% (6 out of 12) of our patients evident of depression symptom. Among these depressed patients, about 50% (3 out of 6) patients were rated as the motor staging I, about 100% (2 out of 2) the staging II, and 33% (1 out of 3) the staging III. There were about 33% evident of Major Depressive Disorder(MDD). Among these patients, 50% with motor staging I, 0% with motor staging II.

On the basis of the dementia criteria of the DSM-IV, 40% (4 out of 10) of our patients were included in this diagnostic category. Among these demented patients, about 50% (2 out of 4) manifested depression.

#### DISCUSSION

Is there an impairment of emotional status in the patients with PD? On the basis of our preliminary results, our patients did have emotional problems. We found that about 50% of our patients with PD were evident of depressed mood. In the literature, depression has often been observed in parkinsonian patients (Mayeux et al., 1986; Ring et al., 1994). The prevalence rate of depression in the patient population was around 40% in average ranging from 4% to 70% (Cummings, 1992). Our results indicating the evidence of depression problem in the parkinsonian patients did corroborate the prior findings. Furthermore, about 50% of our parkinsonian patients having this emotional function seemed to be not only compatible with the prevalence rate reported in western literature (Cummings, 1992), but also with the recent observations (Liu et al., 1997) based on a sample of Taiwanese patients with idiopathic PD. However, our results of about 33% our patients having major depressive disorder based on the current available 2nd-year follow-up seemed to be

inconsistent with the prior findings (e.g., Kostic et al., 1994; Liu et al., 1997). This might be partially due to our small sample of patients (About 15% patients with major depression in our 1<sup>st</sup>-year follow-up was consistent with prior researchers' findings).

Is depressed mood associated with degree of motor disabilities in our patients with PD? The answer seemed to be negative. Liu and his colleagues (1997) noted that most of their depressed patients had nothing to do with their motor disabilities in terms of motor symptom staging (mainly, I and II) of Hoehn and Yahr (1967). Likewise, our patients included the motor stagings I-III, and we did find that the depression proportion of patients with motor staging I was lower than that of patients with staging II. However, the depression proportion of our patients who had moderate motor severity (stage III) manifested depression symptom was lower than that of patients with staging I. On the basis of the findings of both studies, we would suggest that depression problem evident in patients with idiopathic PD might not have a remarkable association with their motor disabilities. In light of scanty literature on this issue, our tentative claim awaits further investigation.

If so, does the deficit only involve depression or also include otherwise emotional function evaluated? The answer appeared to be partially positive. In addition to depression evident in our patients with PD, otherwise emotional disturbances, including somatic, obsessive-compulsive, and anxiety symptoms were slightly to mildly also manifest, particularly for patients with staging III. Among these symptoms, anxiety seemed to be slightly remarkable compared with other problems.

If depression does occur, is it significantly associated with dementia? The answer might be partially true. In the literature, the prevalence rate of dementia in patients with PD has been controversial, ranging from 2% to 93%, and the variation probably attributable to different definitions of dementia and population investigated (e.g., Dubois, Boller, Pillon, & Agid, 1991). In the recent reports, Chui (1989), Mayeux and his co-workers (1988), and Rajput (1992) found that the prevalence rate in the demented patients with PD was in the range from 10% to 40%, while Dubois and colleagues (1991) reported about 50% patients suffering from dementia. It appeared that the frequency of our demented patients was consistent with researchers' observations Mayeux and his co-workers' (1988) while it was higher than that of others' (e.g., Dubois et al., 1991). These inconsistent results await further investigation. In addition, the present results revealed that dementia did not seem to be positively associated with the degree of motor disabilities in our patients with PD. Our results further corroborated these previous findings (e.g., Sagar, 1999).

The investigators (Mayeux et al., 1981) claimed that there was a remarkable relationship between depression and dementia in the patients with PD. We found that 50% of our demented patients with PD were also evident of depression symptom and

these results seemed to partially substantiate Mayeux and his colleagues' observations..

Self-awareness is one of the frontal lobe functions (Stuss & Alexander, 2000), and the neuropathological involvement of frontal-striatal loop is generally evident in patients with PD (e.g., Lichter & Cummings, 2001). Accordingly, these patients' subjective complaints of emotional disturbances, such as depressed mood on the conventional inventories (e.g., Beck Depression Inventory) or interview scales (Hamilton Depression Scale) might not be valid and reliable because of their poor self-awareness functioning. In the present study, we determined whether the patient manifested depressed mood based on multiple data sources, including the patient's self-report, his/her significant informant's description, and the examiner's clinical observations, rather than the patient's self-report only used in the prior studies. Thus, it appeared that our results might be more justifiable than those previous ones, and the contradictory findings might also be partially attributable to this methodological variation.

#### REFERNCES

- Adams, R.D., Victor, M., & Ropper, A.H. (1997). Principles of neurology (6<sup>th</sup> ed.). New York: McGraw-Hill.
- 2. Bayles, K.A. (1990). Language and Parkinson disease. Alzheimer Disease & Associated Disorders, 4, 171-180.
- 3. Beatty, J. (1955). Principles of behaivoral neuroscience. Chicago, ILL: Brown & Benchmark Publishers.
- 4. Beck, A.T. (1987). Beck Depression Inventory. San Antonio, TX: The Psychological Corporation.
- Benton, A.L. (1974). Revised Visual Retention Test (4<sup>th</sup> ed.). New York: The Psychological Corporation.
- 6. Benton, A.L., & Hamsher, K.desS. (1978). Multilingual Aphasia Examination. Iowa City: University Hospital of Iowa, Department of Neurology.
- Benton, A.L., Hamsher, K.deS., Varney, N., & Spreen, O. (1983). Contributions to neuropsychological assessment: a clinical manual. New York: Oxford University Press.
- Boller, F., Passafiume, D., Keefe, N.C., Rogers, K., Morrow, L., & Kim, Y. (1984). Visuospatial impairment in Parkinson's disease: role of perception and motor factors. Archives of Neurology, 41, 485-490.
- 9. Brown, R., & Jahanashi, M. (1995). Depression in Parkinson's disease: a psychosocial viewpoint. Advanced Neurology, 65, 61-84.
- 10. Celesia, G.G., & Barr, A.N. (1970). Psychosis and other psychiartic manifestations of levodopa therapy. Archives of Neurology, 23, 193-200.

- 11. Conn, P.M. (1995). Neuroscience in Medicine. Philadelphia, PA: J.B. Lippincott Company.
- 12. Cummings, J.L. (1992). Depression and Parkinson's disease: a review. American Journal of Psychiatry, 149, 443-454.
- Davison, G.C., & Neale, J.M. (1998). Abnormal Psychology (7<sup>th</sup> ed.). New York: John Wiley & Sons, Inc.
- Derogatis, L.R. (1977). SCL-90-R version manual-I. Baltimore: Clinical Psychometrics Research Unit, Johns Hopkins University School of Medicine.
- 15. Factor, S.A. & Brown, D. (1992). Clozapine prevents recurrence of psychosis in Parkinson's disease. Movement Disorder, 7, 125-131.
- Fahn, S., & Elton, E. (1987). For the UPDRS Development Committee. Unified Parkinson's Disease Rating Scale. In Fahn, S., Marsden, C.D., Goldstein, M., Calne, C.D. (eds.). Recent Development in Parkinson's Disease. New York: Macmillan Publishing Co Inc.
- 17. Fibigier, H.C. (1984). The neurobiological substrates of depression in Parkinson's disease: a hypothesis. Canadian Journal of Neurological Sciences, 11, 105-107.
- Goodwin, F.K. (1971). Psychiatric side effects of levodopa in man. JAMA, 218, 1915-1920.
- Hamsher, K. deS. (1983). Orientation to Personal Information and Place.
  Milwaukee: University of Wisconsin Medical School, Department of Neurology.
- Hamsher, K.deS. (1983). Standard Neurobehavioral Interview Inventory: Milwaukee: University of Wisconsin Medical School, Department of Neurology.
- 21. Hoehn, M.M., & Yahr, M.D. (1967). Parkinsonism: onset, progression, and mortality. Neurology, 17, 427-442.
- 22. Hua, M.-S. (1986). Word Sequence Learning-Revised. Unpublished manuscript. Chungli, Taiwan: Chung Yuan University, Department of Psychology.
- Ingram, R.E., Kendall, R.C., Smith, T.W., Donnell, C., & Ronan, K. (1987). Cognitive specificity in emotional distress. Journal of Personality and Social Psychology, 53, 734-742.
- Iruela, I.M., Ibanez-Rojo, V., Palanca, I., & Caballero, L. (1992). Anxiety disorders and Parkinson's disease (Letter). American Journal of Psychiatry, 149, 719-720.
- 25. Jouvent, R., Abensour, P., Bonnet, A.M., et al. (1983). Antiparkinsonian and antidepressant effects of high doess of bromocriptine. Biological Psychiatry, 5, 141-145.
- Kuzis, G., Sabe, L., Tiberti, C., Leiguarda, R., & Starkstein, S.E. (1997). Cognitive functions in major depression and Parkinson's disease. Archives of Neurology, 54, 982-986.

- 27. Lang, A.E., Quinn, N., Brincat, S., et al. (1982). Pergolide in later-stage Parkinson's disease. Annals of Neurology, 12, 243-247.
- 28. Levin, B.E., Llabre, M.M., & Weiner, W.J. (1989). Cognitive impairments associated with early Parkinson's disease. Neurology, 39, 557-561.
- 29. Liu, C.-Y., Wang, S.-J., Fuh, J.-L., Lin, C.-H., et al. (1997). The correlation of depression with functional activity in Parkinson's disease. Journal of Neurology, 244, 493-498.
- Mayberg, H.S., Mahurin, R.K., & Brannon, S.K. (1995). Parkinson's depression: discrimination of mood-sensitive and mood-insensitive cognitive deficits using fluoxetine and FDG PET. Neurology, 45, A166.
- Mayberg, H.S., Starkstein, S.E., Sadzot, B., Preziosi, T., Andrezejewski, P.L., Dannals, R.F., Wagner, H.N., & Robinson, R.G. (1990). Selective hypometabolism in the inferior frontal lobe in patients with Parkinson's disease. Annals of Neurology, 28, 57-64.
- 32. Mayeux, R. (1990). Dementia in extrapyramidal disorders. Current Opinions of Neurology, 3, 98-102.
- Mayeux, R., Stern, Y., & Williams, J.B.W. (1986). Clinical and biochemical features of depression in Parkinson's disease. American Journal of Psychiatry, 143, 756-759.
- 34. Nelson, H.E. (1976). A modified card sorting test sensitive to frontal lobe defects. Cortex, 12, 313-324.
- 35. Reynolds, C.F. (1992). Treatment of depression in special populations. Journal of Clinical Psychiatry, 53, 45-53.
- Ring, A., Bench, C.J., Trimble, M.R., Brooks, D.J., Frackowiak, R.S.J., & Dolan, R.J. (1994). Depression in Parkinson's disease: a Positron Emission study. British Journal of Psychiatry, 165, 333-339.
- Sano, M., Marder, K., & Dooneief, G. (1996). Basal ganglia diseases. In Fogel, B.S., Schiffer, R.B., & Rao, S.M. (Eds.). Neuropsychiatry. Baltimore: Williams & Wilkins.
- Sano, M., Stern, Y., Williams, J.B.W., Cote, L., Rosenstein, R., & Mayeux, R. (1989). Archives of Neurology, 46, 1284-1286.
- Schiffer, R.B., Kurlan, R., Rubin, A., & Boer, S. (1988). Evidence for atypical depression in Parkinson's disease. American Journal of Psychiatry, 145, 1020-1022.
- 40. Seibert, P.S., & Ellis, H.C. (1991). Irrelevant thought, emotional mood states, and cognitive task performance. Memory & Cognition, 19, 507-513.
- 41. Speedie, L., O'Donnell, W., Rabins, P., Pearlson, G., Poggi, M., & Gonzalez Rothi, L. (1990). Language performance deficits in elderly depressed patients.

Aphasiology, 4, 197-205.

- 42. Spencer, K.A., Tompkins, C.A., & Schulz, R. (1997). Assessment of depression in patients with brain pathology: the case of stroke. Psychological Bulletin, 122, 132-152.
- Spreen, O., & Benton, A.L. (1969). Neurosensory Center Comprehensive Examination for Aphasia. Victoria, BC: Neuropsychological Laboratory, Department of Psychology, University of Victoria.
- 44. Starkstein, S.E., Mayberg, H.S., Leiguarda, R., Preziosi, T., & Robinson, R.G. (1992). A prospective longitudinal study of depression, cognitive decline, and physical impairments in patients with Parkinson's disease. Journal of Neurology, Neurosurgery & Psychiatry, 55, 377-382.
- 45. Starkstein, S.E., Preziosi, T.J., Forrester, A.W., & Robinson, R.G. (1990). Specificity of affective and autonomic symptoms of depression in Parkinson's disease. Journal of Neurology, Neurosurgery & Psychiatry, 53, 869-873.
- 46. Stein, M.B., Heuser, I.J., Juncos, J.L., Uhde, T.E. (1990). Anxiety disorders in patients with Parkinson's disease. American Journal of Psychiatry, 147, 217-220.
- Troster, A.I., Stalp, L.D., Paolo, A.M., Fields, J.A., & Koller, W.C. (1995). Neuropsychological impairment in Parkinson's disease with and without depression. Archives of Neurology, 52, 1164-1169.
- 48. Wechsler, D. (1981). Wechsler Adult Intelligence Scale-Revised manual. San Antonio, TX: Psychological Corp.
- White, K.E., & Cummings, J.L. (1997). Neuropsychiatric aspects of Alzheimer's and other dementing illness. In Yudofsky, S.C., & Hales, R.E (Eds.): The American Psychiatric Press Textbook of Neuropsychiatry (3<sup>rd</sup>. ed.). Washington, DC: American Psychiatric Press, Inc.

#### Table 1. Demographic Data and VIQ Score of Subject Groups (2nd year follow-up)

	Gp1 (n=4)			Gp2			Gp3			Gp4		Н	р	
			(n=3)			(n=5)			(n=10)					
	М	(SD)	Range	М	(SD)	Range	М	(SD)	Range	М	(SD)	Range		
Age(years)	55.75	11.38	44-66	53.67	13.65	38-63	71.20	6.22	64-81	63.00	8.38	51-77	7.45	>.05
Edu.(Yrs)	11.50	2.08	9-14	9.00	3.00	6-12	8.25	4.92	3-16	13.00	3.18	6-16	6.19	>.05
VIQ	97.50	3.11		78.00	10.82		76.80	12.50		112.10	11.40		16.00	<.05
Gender(M/F)		4/0			1/2			2/3			3/7			

WAIS-R: Wechsler Adult Intelligence Scale-Revised; H: Kruskal-Wallis one way ANOVA; a: significant pairwise contrast between Gp1 and otherwise

Gps; b: significant pairwise contrast between Gp2 and otherwise Gps; c: significant pairwise contrast between Gp3 and otherwise Gps; d: significant

pairwise contrast between Gp4 and otherwise Gps

Gp1: Hoehn & Yahr staging 1; Gp2: Hoehn & Yahr staging 2; Gp3: Hoehn & Yahr staging 3; Gp4: Normal controls

Table 2. Learning and Memory Test Performance of Subject Groups (2nd year follow-up)

	Gp1		Gp2		Gp3		Gp4		Н	р
	Mean	SD	Mean	SD	Mean	SD	Mean	SD		
Orientation										
ТО	0.00	0.00	0.00	0.00	1.80	3.49	0.10	0.32	4.19	>.05
OPIP	11.25	0.96	11.67	0.58	10.20	2.05	11.80	0.42	3.78	>.05
Remote Memory Test	46.25	2.06	41.33	8.96	38.20	9.58	47.70	2.31	5.91	>.05
Remote Memory Test (Recognition)	48.50	1.73	47.33	3.79	44.20	5.17	49.10	2.02	6.18	>.05
Recent Memory Test	37.75	2.87	32.00	1.73	28.00	6.28	36.80	3.16	11.91	<.05
Recent Memory Test (Recognition)	40.00	0.00	38.00	1.73	34.60	4.83	39.00	1.76	8.40	<.05
Verbal learning and memory										
WSL										
Correct	50.75	2.63	44.33	11.68	25.50 <sup>d</sup>	14.20	54.80°	4.61	12.62	<.05
Position	33.00	4.55	26.67	22.85	11.75	3.86	46.90	17.26	9.40	<.05
Recall	5.00	1.15	3.00	3.00	0.00	0.00	3.10	2.02	6.45	>.05
Cue	5.75	0.50	5.33	1.15	4.50	0.71	4.00	1.63	5.41	>.05
Recognition	29.50	0.58	28.00	2.65	24.00	2.37.	27.00	5.08	4.20	>.05
BVRT										
Correct	7.25	1.71	6.50	3.54	2.20	1.64	7.10	2.33	8.34	<.05
Error	$3.00^{\circ}$	2.16	$6.00^{\circ}$	7.07	14.80	4.76	3.40	3.20	10.10	<.05

TO: Temporal Orientation; OPIP: Orientation to Personal Information and Place; WSL: Word Sequence Learning-Revised; BVRT: Be-

nton Visual Retention Test

Gp1: Hoehn & Yahr staging 1; Gp2: Hoehn & Yahr staging 2; Gp3: Hoehn & Yahr staging 3; Gp4: Normal controls

Table 3. Visual Perception Test Performance of Subject Groups (2nd year follow-up)

	Gp1		Gp2		Gp3		Gp4	Gp4			
	Mean	SD	Mean	SD	Mean	SD	Mean	SD			
JLO	23.75	2.99	16.67	2.52	18.25	6.18	22.70	3.53		6.43	
3-DBC											
Correct	28.25	0.96	24.67	4.04	26.75	1.71	27.90	2.33		3.91	
FRT	39.75	0.96	38.00	8.19	37.40	2.51	44.70	3.59		10.53	

JLO: Judgement of Line Orientation; 3-DBC: Block Construction-Model; FRT: Facial Recognition Test

Gp1: Hoehn & Yahr staging 1; Gp2: Hoehn & Yahr staging 2; Gp3: Hoehn & Yahr staging 3; Gp4: Normal controls

Table 4. Core Linguistic Test Performance of Subject Groups (2nd year follow-up)

	Gp1	Gp1		Gp2		Gp3			Н	Р
	Mean	SD	Mean	SD	Mean	SD	Mean	SD		
TT	42.50	1.91	40.00	4.00	34.50	7.72	41.30	2.11	4.98	>.05
VN	55.50	1.91	52.67	2.31	48.40	7.54	54.40	3.24	5.65	>.05

TT: Token Test; VN: Visual Naming

Gp1: Hoehn & Yahr staging 1; Gp2: Hoehn & Yahr staging 2; Gp3: Hoehn & Yahr staging 3; Gp4: Normal controls

#### Table 5. Frontal Lobe Function Test Performance of Subject Groups (2nd year follow-up)

	Gp1		Gp2		Gp3		Gp4		Н	р
	Mean	SD	Mean	SD	Mean	SD	Mean	SD		
WCST-M										
No. of complete										
categories	4.75	2.06	3.33	3.21	1.00		6.20	1.14	5.10	>.05
No. of preservative errors	3.50	4.51	4.67	3.51	6.00		3.00	2.31	2.21	>.05
No. of non-preservative errors	7.25	3.40	11.33	8.50	17.00		4.80	4.18	4.82	>.05
% of preservative errors	0.22	0.20	0.29	0.12	0.26		0.38	0.24	2.11	>.05
Trail Making A (sec.)	43.25	18.54	52.00	14.80	160.50	66.97	46.60	20.96	9.07	<.05
Trail Making B (sec.)	110.33	50.21	182.00	175.36	453.00		98.88	45.48	2.77	>.05
VF	24.00	3.46	32.33	2.31	24.80	6.87	42.70	8.67	14.52	<.05
Similarity (WAIS-R)	8.50	1.00	6.33	2.52	5.40	2.61	11.90	2.08	15.31	<.05
Digit Span(forward-backward	3.00	1.15	3.67	0.58	2.00	1.00	3.60	1.17	6.61	>.05

WCST-M: Wisconsin Card Sorting Test-Modified; VF: Semantic Association of Verbal Fluency

Gp1: Hoehn & Yahr staging 1; Gp2: Hoehn & Yahr staging 2; Gp3: Hoehn & Yahr staging 3; Gp4: Normal controls

#### Table 6. Emotional Status Measure Performance of Subject Groups (2nd year follow-up)

	Gp1		Gp2		Gp3		Gp4		Н	Р
	Mean	SD	Mean	SD	Mean	SD	Mean	SD		
BDI	11.50	3.54	24.00		6.00	4.24	7.50	6.20	3.90	>.05
SCL-90-R										
Somatization	0.52	0.49	1.20	0.29	1.13	1.00	0.53	0.37	5.46	>.05
Obsessive-compulsive	0.73	0.38	0.77	0.25	1.40	0.87	0.74	0.58	2.43	>.05
Depression	0.64	0.51	0.62	0.41	1.21	0.98	0.45	0.46	2.75	>.05
Anxiety	0.63	0.35	0.30	0.36	1.30	0.98	0.34	0.38	5.43	>.05

BDI: Beck Depression Inventory; SCL-90R: Symptom Checklist-90-R

Gp1: Hoehn & Yahr staging 1; Gp2: Hoehn & Yahr staging 2; Gp3: Hoehn & Yahr staging 3; Gp4: Normal controls

Table 7. Emotional Status	Measure Performance of	Significant Informant (	Groups (2nd year follow-up)

	Gp1		Gp2		Gp3		Gp4			Н	р
	Mean	SD	Mean	SD	Mean	SD	Mean	SD	_		
BDI_F	4.00		16.50	6.36	7.00	1.73	6.63	7.17		3.14	>.05
SCL-90-R											
Somatization	0.08		1.00		0.86	0.13	0.33	0.47		4.15	>.05
Obsessive-compulsive	0.50		1.10		0.75	0.48	0.40	0.37		2.92	>.05
Depression	0.38		0.62		0.64	0.29	0.35	0.40		1.46	>.05
Anxiety	0.40		0.80		0.63	0.35	0.27	0.39		3.82	>.05

Gp1: Hoehn & Yahr staging 1; Gp2: Hoehn & Yahr staging 2; Gp3: Hoehn & Yahr staging 3; Gp4: Normal controls

Table 8. The Examiner's Rating Scores on the Emotional Status Interview Scales (2nd year follow-up)

	Gp1		Gp2		Gp3		Gp4		Н	р
	Mean	SD	Mean	SD	Mean	SD	Mean	SD	-	
Depression	0.60		0.40		0.93	0.38	0.10	0.14	8.73	<.05
Anxiety	0.50		1.00		1.25	0.35	0.00	0.00	12.90	<.05

Gp1: Hoehn & Yahr staging 1; Gp2: Hoehn & Yahr staging 2; Gp3: Hoehn & Yahr staging 3; Gp4: Normal controls

### 計畫成果自評

本研究 (第二年追蹤研究) 內容與原計劃大致符合,但達成目標與預期有一段距離。但由於病 人的病情嚴重程度、或意願問題無法繼續參與追蹤研究,使得個案數較少並與預期有落差,因此一 些欲探討的問題無法如願達成。第一年追蹤研究個案數較符合預期,未來將以該研究結果來完成論 文,並投遞至期刊發表。

本研究發現巴金森病人有情緒問題,尤其是憂鬱 (50%病人),但該問題與病人的運動障礙嚴重性沒 有緊密關係,40%病人罹患失智症的情形則與國外研究大致符合。雖然本研究中有憂鬱症狀的病人 裡,有40-50%同時罹患失智症目前,但可能不能以憂鬱來解釋巴金森病人罹患失智症的原因。