A Prospective Study Of Prescription Pattern and Assessing the Psychiatric Co-Morbidities In Dementia Patients In A Tertiary Care Hospital In South Kerala

Neethu J¹*, Aranya K Sreekumar¹, Jibin John¹, Kavya Shyam¹, Reshma S Reghu², Prasobh GR³

¹.Associate professor, Department of pharmacy practice, Sreekrishna college of pharmacy and research centre, Thiruvananthapuram, Kerala, India
². Fifth year Pharm D students, Sreekrishna college of pharmacy and research centre, Thiruvananthapuram,Kerala, India
³. Principal ,Head Of Department of pharmacy practice, Sreekrishna college of pharmacy and research centre, Thiruvananthapuram, Kerala , India

ABSTRACT

Dementia is one of the most common disorders among elderly patients. Worldwide 7.7 million new cases of dementia were anticipated each year, implying one new case every 4.1 seconds. The aim of the study is to assess the psychiatric co morbidity and prescribing pattern in a tertiary care hospital in south India. This was an Prospective Observational Study conducted at tertiary care hospital. Prospective data was collected from outpatient and inpatient department. Out of 16 patients the common type of dementia reported was Alzheimer’s disease found in 56.3% of the cases followed by vascular dementia and the prescribed drugs were Memantine (37.5%) is the prescribed drug for most of the dementia cases followed by donepezil (25%), Memantine + donepezil combination (18.75%), Rivastigmine (12.5%), Memantine + Rivastigmine combination (6.2%). The common type of dementia reported was Alzheimers and Memantine was the most commonly prescribed drug in mono-therapy and among combination therapy Memantine–donepezil combination was mostly prescribed.

Keywords: dementia, prescribing pattern, psychiatric co morbidity
INTRODUCTION

Dementia is defined as an acquired deterioration in cognitive abilities that impairs the successful performance of activities of daily living. Memory is the most common cognitive ability lost with dementia. In addition to memory, other mental faculties were affected; these include language, visuospatial ability, calculation, judgment and problem solving. Dementia syndromes result from disruption of specific large scale neuronal networks, the location and severity of synaptic and neuronal loss combine to produce a clinical features. Behaviour and mood are modulated by noradrenergic, serotonergic and dopaminergic pathways whereas the cholinergic signalling is critical for attention and memory functions.\(^1\)

The number of people living with dementia worldwide was estimated at 47.47 million as of 2015, reaching 75.63 million in 2030 and 135.46 million in 2050.\(^2\) From 2015 to 2050, numbers of people living with dementia will have increased slightly less than twofold in Europe, somewhat more than twofold in north America, threefold in Asia, and fourfold in Latin America and Africa. While 37% of the people living with dementia live in high-income countries, 63% live in low and middle-income countries. The incidence of dementia increased exponentially with increasing age, based on the available estimates for the global incidence of dementia dating from 2010.\(^3\) 7.7 million new cases of dementia are reported each year worldwide, implying one new case every 4.1 seconds. This means that there were 3.6 million (46%) new cases per year in Asia, 2.3 million (31%) in Europe, 1.2 million (16%) in the Americas, and 0.5 million (7%) in Africa. It is expected that the burden of dementia will be increasing in developing countries due to increase in longevity and increasing prevalence of risk factors such as hypertension and stroke. In India 1.8 million people are affected with dementia.\(^3\)

Alzheimer's dementia begins in the transentorrhinal region spreads to the hippocampus and then moves to lateral, posterior, temporal and parietal neocortex, eventually causing more widespread degeneration. Vascular dementia is associated with focal damage in random network of cortical and subcortical regions or white matter tract that disconnect nodes within disturbed networks. patients with dementia that begins in frontal or subcortical regions such as fronto-temporal dementia or Huntington’s disease are less likely to begin with memory problems and more likely to have difficulties with mood and behaviour.\(^1\)
The treatment of dementia is basically directed towards management of cognitive and Behavioural symptoms of dementia. Particularly effective treatments are available for most common symptoms of dementia. Acetylcholinesterase inhibitors such as Donepezil, Rivastigmine, and Galantamine have been recommended to be used in mild to moderate cases of dementia while NMDA receptor antagonist Memantine is the only drug recommended for severe cases of dementia. The pathologic action of Donepezil, Rivastigmine and Galantamine is inhibition of the cholinesterase, primarily acetyl choline-esterase, with the resulting increase in cerebral acetyl choline level. Memantine appears to acts by blocking over-excited N-Methyl D-Aspartate (NMDA) glutamate receptor1.

With increasing life expectancy, the prevalence of dementia will increase dramatically in the next few decades, and will have a phenomenal emotional and economic impact. Thus cognitive impairment and dementia are one among the major public health concerns of the 21st century. In this context there seems to be scarcity of data regarding dementia. This study highlights the present scenario of dementia in relation to epidemiology, psychiatric co morbidity and drug utilization pattern in a tertiary care hospital in south kerala 2-4.

Neuropsychiatric and social deficits also seen in many subtypes of dementia resulting in depression, apathy, hallucinations, delusions, agitation, insomnia and disinhibition. The most common forms of dementia are progressive but some are static and unchanging or fluctuate from day to day or even minute4.

MATERIALS AND METHOD

All relevant information regarding the study will be collected from case records and direct interview with the caregivers in the presence of the physician. Data from patients or caregiver will be collected by using a specially designed proforma.

Pharmacotherapy data (indication, dose, posology, scheduling, and reports on the access to medicines) will be collected from the medical prescription, patient diary and all medicines (over-the-counter and prescription medicines) brought by the patient to the appointment with the physician.

The patients or caretaker will then be educated about the disease and medication. To assess the patient condition DSRS (Dementia severity Rating scale) questionnaire and SLUMS Scale will be used on all the patients during each visit.

To assess psychiatric co morbidities in dementia patients using HADS (Hospital anxiety and depression scale) will be used.

To improve the quality of life in patient by giving effective counselling to the caregiver

RESULTS AND DISCUSSION
Average age of the study population was 79.9 ± 6.1 and age ranges from 69-89 years. 12.5% of the patients were in the age group ≤70 years, 6.3% were 71-75 years,31.3% were in the 76-80 years and 81-85 years and 18.8% >86 years. Dementia is mostly observed in the age groups from 76 – 85 years and least observed in the age group from 71-75.

Table 1: Dementia Patients Age Distribution

| Age in years | Frequency | Percentage |
|--------------|-----------|------------|
| ≤70          | 2         | 12.5       |
| 71 - 75      | 1         | 6.3        |
| 76 - 80      | 5         | 31.3       |
| 81 - 85      | 5         | 31.3       |
| >86          | 3         | 18.8       |
| Total        | 16        | 100.0      |

In the gender distribution males dominated in the study population which is 56.3% while in females it is 43.8% as depicted in Table 2.
Figure 2: Prevalence of Dementia according to Gender.

Table 3: Education status of Dementia Patients.

| Qualification        | Frequency | Percentage |
|----------------------|-----------|------------|
| Illiterate           | 2         | 12.5       |
| High school          | 1         | 6.3        |
| Higher secondary     | 2         | 12.5       |
| Graduate             | 9         | 56.3       |
| Postgraduate and above | 2      | 12.5       |
| Total                | 16        | 100.0      |

Regarding education of patients in the sample, it was noted that 56.2% of the patients were college graduates, 12.5% of the patients were high school graduates, 12.5% were higher secondary graduates and 12.5% were post graduates and 12.5% of the patients were illiterate.

Figure 3: Education status of Dementia patients.

Table 4: Occupation of Dementia Patients

| Occupation      | Frequency | Percentage |
|-----------------|-----------|------------|
| Employed        | 7         | 43.8       |
| Business        | 4         | 25.0       |
| Manual labour   | 1         | 6.3        |
| Unemployed      | 4         | 25.0       |
Regarding the occupational status of dementia patients, 43.8% of the patients were employed, 25% of the patients were businessmen, 25% of them were unemployed and 6.3% of the patients were manual labour.

![Occupational Status of Dementia Patients](image1)

**Figure 4: Occupational Status of Dementia Patients**

| Marital status | Frequency | Percentage |
|----------------|-----------|------------|
| Married        | 13        | 81.3       |
| Unmarried      | 1         | 6.3        |
| Separated      | 1         | 6.3        |
| Widow/Widower  | 1         | 6.3        |
| Total          | 16        | 100.0      |

Regarding the marital status of the patients, 81.3% of the patients were married and 6.3% of the patients were unmarried. 6.3% of them were separated and 6.3% of the patients were widowed.

![Marital status of Dementia Patients](image2)
Figure 5: Marital status of Dementia Patients.

Table 6: Socio-Economic status of Dementia Patients.

| SES        | Frequency | Percentage |
|------------|-----------|------------|
| Upper      | 2         | 12.5       |
| Upper middle | 7         | 43.8       |
| Lower middle | 4         | 25.0       |
| Upper lower | 3         | 18.8       |
| Total      | 16        | 100.0      |

Regarding the socio-economic status of dementia patients, 43.8% of the patients were of upper class, 43.8% were of upper middle class, 25% of the patients were of lower middle class and 18.8% of the patients were of upper lower class.

Figure 6: Socio-economic status of dementia patients.

Table 7: Depiction of Subtypes Of Dementia

| Type of dementia      | Frequency | Percent |
|-----------------------|-----------|---------|
| Alzheimer’s disease   | 9         | 56.3    |
| Vascular dementia     | 3         | 18.8    |
| Lewy body dementia    | 1         | 6.3     |
| Fronto-temporal dementia | 1       | 6.3     |
| Beclouded dementia    | 1         | 6.3     |
| Subcortical dementia  | 1         | 6.3     |
| Total                 | 16        | 100.0   |

Out of 16 patients the common type of dementia reported was alzheimers disease found in 56.3% of the cases followed by vascular dementia which is found in 18.8% of the cases followed by lewy body dementia, fronto-temporal dementia and beclouded dementia and sub cortical dementia, which were found to be 6.3% each.
According to our study we have found out that the most presenting symptom is forgetting recent events with a percent of 100, 62.5% of unaware of time & place, 68.8% of difficulty in recognizing relatives, 31.3% of difficulty in performing familiar task, 43.8% of lost in familiar places, 25% of poor concentration, 18.8% of difficulty in communication, 25% of difficulty in walking, 12.5% disturbed sleep, 6.3% of loss of interest, 18.8% of behavioural changes, 50% of mood swings & also there were no patients presenting with poor judgment.
Figure 8: Symptoms of Dementia

Table 9: Representation of comorbidity distribution of dementia patients

| Comorbidities          | Frequency | Percentage |
|------------------------|-----------|------------|
| HTN                    | 12        | 75.0       |
| DM                     | 12        | 75.0       |
| CAD                    | 4         | 25.0       |
| Dyslipidemia           | 3         | 18.8       |
| Thyroid                | 0         | 0.0        |
| Surgery                | 5         | 31.3       |
| Liver disease          | 0         | 0.0        |
| Kidney disease         | 1         | 6.3        |
| Seizure                | 0         | 0.0        |
| Parkinsonism           | 2         | 12.5       |

Distribution of different comorbidities associated with dementia patients is represented in figure 9. Regarding the comorbidity pattern in dementia patients, it was observed that hypertension and diabetes were found to be the most common type of comorbidity. This is about 75% each, which is followed by surgery and CAD 4.3% and 25% respectively. Followed by dyslipidemia 18.8%, parkinsonism 12.5% and kidney disease 6.3%. None of the patients were having liver disease, thyroid disease or seizure.
Figure 9: Representation of Comorbidity Distribution of Dementia

Table 10: Representation of Psychiatric Comorbidity Distribution of Dementia

| Psychiatric Co morbidity | Frequency | Percentage |
|--------------------------|-----------|------------|
| Depression               | 6         | 37.5       |
| Anxiety                  | 9         | 56.25      |
| NIL                      | 1         | 6.25       |

In total of 16 cases it was observed that 37.5% of patients was having depression and 56.25% patients was found to have anxiety, 6.25% of patients were having neither depression or nor anxiety.

Figure 10: Representation of Psychiatric Comorbidity Distribution Of Dementia
Table 11: Depiction Of Dose Distribution Of Different Drugs Used In Dementia

| Drug          | No Of Patients | %  | 5mg No. Of Patients | %  | 10 mg No. Of Patients | %  | 1.5 mg No. Of Patients | %  |
|---------------|----------------|----|---------------------|----|-----------------------|----|-----------------------|----|
| Memantine     | 10             | 62.5 | 3                   | 18.8 | 7                     | 43.8 |
| Donepezil     | 7              | 43.75 | 3                   | 18.8 | 4                     | 25.0 |
| Rivastigmine  | 3              | 18.75 |                    |     |                       |     |

Prescription analysis of dementia patients depicts that most popular drug of choice for the given condition was Memantine in a dose range of 5 to 10 mg. The doses of drugs used were 5mg dose in 18.8% of cases and 10mg dose in 43.8% of cases in Memantine, which is followed by Donepezil in a dose range of 5 to 10 mg. The doses of drugs used were 5mg dose in 18.8% of cases and 10mg dose in 25.0% of cases in Donepezil, while for Rivastigmine it was 1.5mg dose in 18.8%.

Table 12: Prescribing Patterns In Dementia Patients

| Drugs               | Frequency | Percentage |
|---------------------|-----------|------------|
| Memantine           | 6         | 37.5       |
| Donepezil           | 4         | 25         |
| Memantine+ Donepezil| 3         | 18.75      |
| Rivastigmine        | 2         | 12.5       |
| Memantine+ Rivastigmine | 1   | 6.2        |
| Total               | 16        | 100        |

In this study NMDA receptor antagonist, memantine (37.5%) is the prescribed drug for most of the dementia cases followed by donepezil (25%), memantine+ donepezil combination (18.75%), rivastigmine (12.5%), memantine+ rivastigmine combination (6.2%).
Table 12: SLUMS Score of Dementia Patients

| SLUMS score | Frequency | Percentage |
|-------------|-----------|------------|
| MCI         | 1         | 6.3        |
| Dementia    | 15        | 93.8       |
| Total       | 16        | 100.0      |

SLUMS scale was conducted to determine whether the patient having dementia or mild cognitive impairment. According to the criteria, about 93.8% patients was found to have dementia and 6.3% having mild cognitive impairment.

![SLUMS Score of Dementia Patients](image)

Figure 12: SLUMS Score of Dementia Patients

Table 13: DSRS score of dementia patients.

| DSRS     | Frequency | Percentage |
|----------|-----------|------------|
| Mild     | 5         | 31.3       |
| Moderate | 8         | 50.0       |
| Severe   | 3         | 18.8       |
| Total    | 16        | 100.0      |

Dementia severity rating scale was conducted to determine the severity of dementia. 50% of patients was observed to have moderate dementia followed by 31.3% patients with mild dementia and 18.8% patients with severe dementia.

![DSRS Score of Dementia Patients](image)

Figure 13: DSRS score of dementia patients
Table 14: Hads Score In Dementia Patients

| Hads  | Frequency | Percentage |
|-------|-----------|------------|
| Depression | 6   | 36.5       |
| Anxiety   | 9    | 56.25      |
| Nil        | 1    | 6.25       |

Was conducted in patients to assess the psychiatric comorbidity. In total of 16 cases, it was observed that 37.5% of patients was having depression and 56.25% patients was found to have anxiety, 6.25% of patients were having neither depression nor anxiety.

DISCUSSlON

Dementia is observed in elderly throughout the world but they are poorly recognized and treated in developing countries including India. The disease is important since there are an estimated 35.6 million people with dementia worldwide and more than half of these individuals have Alzheimer’s disease (AD) that result in a progressive decline in cognition, function communication and frequent occurrence of neuropsychiatric symptoms. The prescription pattern of drugs used in dementia patients has not been studied much in developing countries where cost may be the most important factor which determine their choice. In this study, we highlight the present Indian scenario of dementia in relation to its severity, comorbidities, and drug utilization in this particular region.

In our study average age of the study population was 79.9 ± 6.1 and age ranges from 69-89 years. Dementia is mostly observed in the age groups from 76 – 85 years and least observed in the age group from 71- 75 and males dominated the study population which is 56.3%. In one of the similar study conducted at multiple centers by Gil-Neciga and Gobartt, it was found that the mean age of patients was 77 ± 6.6 years and that female dominated the study population. On contrary, in the study done by Jeschke et al. during the 5-year study period (2004–2008), in 577 patients with
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Dementia, 81 years was found to be the median age of the patient and females dominated the study population which is, 69% 10. The comparatively lower population of dementia patients in our study might be due to the less preference given to females in the Indian community and therefore neglected and not brought to the hospitals when treatment is essential.

In the study it was noted that 56.2% of the patients were college graduates, 12.5% of the patients were high school graduates, 12.5% were higher secondary graduates and 12.5% were post graduates and 12.5% of the patients were illiterate. In a similar study conducted by Mansi et al, it was noted that almost 8.0% of the patients were illiterate. A total of 56.0% of the patients were high school graduates. The percentage of population who were graduates was 24.0% while postgraduates were very less 12.0%.

However regarding the Socioeconomic status most of the patients belongs to lower socioeconomic status which may be one of the important risk factors responsible for dementia. In this study it was noted that 12.5% of the patients were of upper class and 43.8% of the patients were of upper lower class.

In a study conducted by Qian et al. found that individuals with lower socioeconomic status come into memory clinic after the disease has progressed to dementia, while higher socioeconomic status individuals present earlier when the disease is still in its mild cognitive impairment stage. They concluded that higher socioeconomic status is associated with better cognitive functioning and increased use of cognitive enhancers 11.

In the present study we found that amongst the types of Dementia the common type of dementia reported was Alzheimers disease found in 56.3% of the cases followed by vascular dementia which is found in 18.8% of the cases followed by lewy body dementia, fronto-temporal dementia and beclouded dementia and sub cortical dementia, which were found to be 6.3% each. In a clinic-based study from South India also depicts similar results, in which Alzheimer’s disease was found to be the commonest in 38.3% of the cases, vascular dementia in 25.4% of the cases, followed by fronto temporal dementia (FTD) in 18.7%, Lewy body disease (DLB) in 8.9%, and mixed dementia in 8.6% of the patients, respectively 12.

Regarding the comorbidity pattern in dementia patients, it was observed that hypertension and diabetes were found to be the most common type of comorbidity. This is about 75% each, which is followed by surgery, CAD, dyslipidaemia, parkinsonism and kidney disease respectively. A study reported by Poblador-Plou et al. also found similar results, where the two most frequent comorbidities both for men and women with dementia were hypertension and diabetes. Other comorbidities significantly associated with dementia were Parkinson’s disease, congestive heart
failure, cerebrovascular disease, anemia, cardiac arrhythmia, chronic skin ulcers, osteoporosis, thyroid disease, retinal disorders, prostatic hypertrophy, insomnia and anxiety, and neurosis. In the present study it was observed that 37.5% of patients was having depression and 56.25% patients was found to have anxiety, 6.25% of patients were having neither depression nor anxiety.

CONCLUSION

This study shows the drug utilization pattern of dementia patients and also assess the psychiatric comorbidities associated with dementia and quality of life of most of the dementia patients are compromised, it is mainly due to the lack of knowledge about the disease and poor medication taking behaviour of the patients.

During our study we provided brief information about the disease to the patients, dietary advice, lifestyle modification measures were given to the patient. Information about each prescribed drug and the importance of taking medication properly were advised to the patient. Data collections are performed through specially designed pro-forma, DSRS questionnaire, HADS questionnaire, SLUMS questionnaire. DSRS and SLUMS scale was used to assess the severity of dementia and HADS scale was used to assess the psychiatric comorbidities. The ratings of these scales were made during each time of patient visit to the doctor.

Pharmacists are in ideal position to provide patient education and optimize the therapeutic outcome. Patient counselling provide a great opportunity to improve the medication adherence of the patient and symptomatic improvement was also observed, further more we also assess the prescribing pattern of dementia in a tertiary care sector from which we came to observe that Memantine is the most commonly prescribed drug in mono-therapy and among combination therapy Memantine–donepezil combination was mostly prescribed.

Our study evaluates the possibility of occurrence of any sort of psychiatric comorbidities with dementia and those patients who were found vulnerable could seek medical care for their condition. The counselling provided was effective in improving the knowledge, attitude and practice of patients with dementia. Pharmacists are in an ideal position to provide patient education and optimize patient care. Greater understanding about the illness and a change of attitude and practice would in turn results in a better therapeutic outcome. For future research the study should be done in a large sample and for long duration of period as a multi-center study.

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