A REVIEW ON PHARMACEUTICAL CARE PLAN IN STROKE PATIENTS

Gopu Dharani¹, Dr. D. Akila Devi²

¹Department of Pharmacy Practice, Department of Pharmaceutics², Vels Institute of Science, Technology & Advanced Studies (VISTAS), Chennai, India.

Corresponding author: Gopu Dharani¹, Department of Pharmacy Practice, Vels Institute of Science, Technology & Advanced Studies (VISTAS), Chennai, India.

ABSTRACT:

Pharmaceutical care plan is a dynamic process and a clinical pharmacist requires more knowledge on drugs and their therapeutic activity. Pharmaceutical care plan benefits the patient healthcare outcome and also benefits health care professionals in disease prevention. In stroke patients pharmaceutical care plan plays very key role to identify the Drug Related Problems (DRPs), investigating the nature and frequency of DRPs including untreated indications, sub-therapeutic dosage, excessive dosage, drug use without a clear indication, failure to receive drugs, improper drug selection, drug interactions, and adverse drug reactions. The evidence supporting clinical pharmacist role in care of stroke patients is well-established and literature has been published evaluating pharmacist interventions in stroke patients. The purpose of this review is to explain the detailed evidence of pharmacist interventions on stroke patient.

METHODS: The current literature was reviewed using MEDLINE (PubMed), ELSEVIER up to JULY 2019, and International Pharmaceutical Abstracts. The search was based on the following medical subject heading Disease-related search and MeSH terms included: Drug related problems in ischemic attack, stroke, cerebrovascular accident, cerebral hemorrhage, hemorrhagic stroke, and intracranial hemorrhage were identified and a qualitative analysis is performed.

RESULTS: A total 25 abstracts and full-text studies were identified, studied and included. These studies provided evidence supporting pharmacist interventions in stroke patients and clinical pharmacist can provide his contribution in the detecting and resolving DRP's and benefits safe therapeutic administration in patients. By identifying adverse drug reactions, risk factor reduction can be possible. The evidence strongly suggests that pharmacist interventions plays key role in stroke patient outcomes.

Keywords: Pharmaceutical care plan, Drug related problems, Stroke.

Introduction

Cerebrovascular diseases are the most common devastating disorders. According to the World Health Organization report, worldwide 15 million people suffer from a stroke every year and 5 million people dies. Majorly stroke causes disability and placing a burden on their family and community (World Health Organization 2018). The cerebrovascular diseases include ischemic stroke, hemorrhagic stroke, and cerebrovascular anomalies such as intracranial aneurysms and arteriovenous malformations (AVMs). Drugrelated problems (DRP) are common in hospitalized patients and results in patient morbidity and mortality. DRP is defined as an event that interferes with desired health outcomes, can lead to ineffective pharmacotherapy, and may cause drug-related morbidity and mortality. Pharmaceutical care plan plays major role in identifying, preventing, and resolving DRPs. Clinical pharmacist optimizes the drug therapy, and improves positive clinical outcomes in reasonable economic expenditures. Pharmaceutical care plan also improves patient's health related quality of life. Defining and classifying DRPs is done by A Norwegian system and consists of six main categories: 1) Drug choice 2) Dosing 3) Adverse Drug Reaction 4) Interaction 5) Drug Use and 6) Alternative. This system involves in reduction of adverse drug events and medication errors. A

literature has been published evaluating the role of the pharmaceutical care plan in stroke patients; the main objective of this systematic review is to summarize the role of clinical pharmacist in stroke patient's outcome.

METHODOLOGY

Literature Search: The current literature was reviewed using MEDLINE (PubMed), ELSEVIER up to JULY 2019, and International Pharmaceutical Abstracts. The search was based on the following medical subject heading, Disease-related search and MeSH terms included: Drug related problems in ischemic attack, stroke, cerebrovascular accident, cerebral hemorrhage, hemorrhagic stroke, and intracranial hemorrhage were identified and a qualitative analysis is performed.

Study Selection: Abstracts and full text articles of interest were identified and reviewed by one author. All articles independently screened for identifying relevant articles and discrepancies resolved by discussion. The abstracts/articles had to contain drug related problems in stroke patients and clinical pharmacist interventions to identify and improve the DRP's. The study population had to comprise patients with Ischemic stroke, transient ischemia attack, or a history of stroke or TIA. Studies included if DRP's in stroke population reported clearly and studies support pharmaceutical care plan is beneficial to improve quality of life in stroke patients. Only studies written in English were included.

RESULTS

A total of 930 articles were identified, 25 of which were included in this review. All systemic articles reviewed published as full text articles. Four cross sectional observational study, six prospective study, three randomization clinical trials, two retrospective descriptive studies, two matched case-control study, two prospective interventional studies are included in this systemic review and remaining is included from patient outcomes such as patient satisfaction, health-related quality of life (HRQoL), drug therapy problems. Studies have shown that pharmaceutical care performed in the hospital settings was feasible, a clear benefit and positive impact on patient's Health related quality of life (HRQoL). Mostly 80% of drug-related problems can be prevented. A prospective observational study by Kavithasamy p et al., suggests that occurrence of stroke was more in men than women because of social habits [1]. Dilip Chandrasekhar et al also explained same that occurrence of stroke was more in men than women [2]. The age group of 41 to 70 are mostly has high risk for stroke. Ischemic stroke cases are registered more in number than other strokes. In this study Drug-Drug Interactions are classified according to the severity, and most number of drug interactions occurred is severe. The observed Drug-Drug interactions in this study increases risk of hospitalization and higher health-care cost. Drug-Drug Interactions mainly occurred between antihypertensive, anticoagulants, and antiplatelet. Antihypertensive contributed much to the occurrence of ADRs [1]. A prospective interventional study by Dilip Chandrasekhar et al., divided study population into intervention group and control group. This study suggests population in rural areas has more risk to stroke than in urban. Ischemic stroke cases are registered more in number than any other strokes. Clinical characteristics like right hemisphere brain lesions is more than left hemisphere lesion. Medium Adherence to anticoagulant therapy in interventional group than in control group is observed. The intensified pharmaceutical care given to intervention group resulted in a greater improvement in the HRQoL subscales and summary scores and this study mainly suggest that pharmaceutical care performed in the hospital settings was feasible and had a clear benefit and positive impact on patient's HRQoL [2].A crosssectional descriptive study by Iman ABasheti et al., suggests drug related problems were categorized into efficacy, inappropriate drug adherence and inappropriate knowledge. The identified drug related problems more than 70.0% were of major severity and 80% DRP's under the "efficacy" category were major in severity, 99.3% of TRPs were major in severity under the "inappropriate drug adherence" category. This study also suggests that patients who had hypertension and diabetes mellitus were found to be none-adherent to their medications and it is associated with increased comorbidity [3]. A prospective study was conducted by Carina Hohmann et al... demonstrates that drug related problems mostly occurs at the time prescribing process. This study suggests DRPs occurs due to inappropriate drug indication and drug dosage. Other factors include age, aphasia, high prevalence of comorbidity, polypharmacy, IV route of administration because of impaired oral intake and drugs which requires frequent monitoring and adjusting dose has high risk of medication errors. Patient receiving

heparin, warfarin, antihypertensive medication, and combination of antiplatelet drugshas high risk for drug interactions and medication errors. This study also suggests that drug related problems occur at transcription error or unintended discontinuation of drug therapy. The author mentioned clinical pharmacist involvement in the discharge process, computerized order entry of drugs, monitoring drug-drug interactions prior to administration, and patient counselling will improve therapeutic outcome [4].A retrospective descriptive study by Kannikar Semcharoen et al.., mentioned most common comorbidities were hypertension and other comorbidities were dyslipidemia, diabetes and atrial fibrillation. The possible drug interactions include Warfarin–phenytoin, Warfarin–carbamazepine, Warfarin–prednisolone, Warfarin–co-trimoxazole, Warfarin–amiodarone, Meropenem–sodium valproate, and Clopidogrel–omeprazole. The author mentioned adverse drug reactions such as

Adverse Drug Reactions to Drugs	
Drug	Adverse drug reaction
Atorvastatin	Hepatotoxic, myopathy
Aspirin plus dipyridamole,	
cilostazol	Headache, palpitation, tachycardia
Nicardipine	Phlebitis
Warfarin	Upper gastrointestional bleeding
Hydralazine, carvedilol	Hypotension
Hydralazine, carvedilol	Hypotension

This study shows evidence of pharmaceutical interventions has benefit in patient outcome. Some of the pharmaceutical interventions provided include, the patients receiving dual antiplatelets and has high risk of bleeding, addition of omeprazole reduces risk, reducing dose of piperacillin/tazobactamdose 4.5 g IV q 6 h to 2.25 g IV q 6 h in patients with creatinine clearance of < 20 ml/min, Increasing atorvastatin dose 20 mg to 40 mg for better reduction of low-density lipoprotein cholesterol, changing omeprazole from intravenous to oral route in patients with normal gastrointestinal tract function, discontinuation of antibiotics if there any persistent infection, stop enoxaparin prior to undergoing cardiac catherization, monitoring vancomycin level while using, measurement of international normalized ratio (INR) in patients receiving warfarin, and replacing warfarin by a new oral anticoagulant in patients with history of labile INR despite long history of warfarin usage. 84% of above clinical pharmacist recommendations has been accepted by physician [5]. A study conducted by Carina Hohmann on intervention group and control group that indicates pharmaceutical care has a clear benefit and positive impact on patient's health related quality of life (HRQoL). In this study 12 months comparison done to patients receiving intensified pharmaceutical care and patients with standard care of HRQoL in patients and this study reports patients who do not receive intensified pharmaceutical care have a higher chance of deteriorating HROoL and patient with pharmaceutical care was stable in several subscales and HROoL is maintained [6]. A matched case-control study by Renata Daud-Gallottia explains adverse events are principal in deceased patients due to improper diagnostic or therapeutic procedures and inexperienced nursing activities. Author mentioned severe adverse events leads increase in mortality [7]. A prospective study and conducted by Ramesh A, explains major drug related problems were observed in patients with age group of 60-69 years, among drugdrug interactions were common. The Author explained clearly drugs involved in drug related problems.

TYPES OF DRUG RELATED PROBLEMS	DRUGS INVOLVED
Drug use without indication	Domperidone
	Piperacillin+tazobactam
	Ondansetron
	Paracetamol
	Pantoprazole
	Tramadol
Drug Duplication	Clopidogrel
	Pantoprazole
Subtherapeutic Dose	Amlodipine
	Telmisartan
<u> </u>	Metoprolol
	Nimodipine
Overdose	Ranitidine
	Rabeprazole
Adverse Drug Reactions	amlodipine induced constipation
	atorvastatin induced myopathy
	insulin induced hypoglycemia
Improper drug selection	Rabeprazole
	Ramipril
	Ondansetron
Failure to Receive drugs	LMWH
	Multivitamins
Medication errors	Clopidogrel
	Aspirin
	Atorvastatin
	Mannitol
	Pantoprazole
Untreated indication	Cough with expectoration
	Anemia
	Vomiting

Types of Drug Related Problems and Drugs involved

The author explained most of the ADRs belonged to 'probable' category on WHO scale. Majority was 'Mild' in severity and identified ADRs were 'Not preventable' and none of them were probably preventable and definitely preventable. Also author mentioned involvement of pharmacist in assessment of drug related problems improves patient quality of life mostly in elderly patients with poly-pharmacy. In this study the pharmacist focused in identification of potential DRPs. Most of the cases, there was a change in drug therapy and this recommendations accepted in high. In this study Poly-pharmacy was found to be a potential risk factor leads to drug related problems. By involving pharmaceutical care for early detection of drug related problems improve the therapeutic outcomes [8].

CONCLUSION:

Studies have shown that pharmaceutical care performed in the hospital settings was feasible, a clear benefit and positive impact on patient's Health related quality of life (HRQoL). Studies also suggest occurrence of stroke was more in men than women because of social habits and age group of 41 to 70 are mostly has high risk for stroke. Ischemic stroke cases are registered more in number than other strokes. The most number of drug interactions occurred is severe and observed Drug-Drug interactions in this study increases risk of

hospitalization and higher health-care cost. Drug-Drug Interactions mainly occurred between antihypertensive, anticoagulants, and antiplatelet. Antihypertensive contributed much to the occurrence of ADRs. The drug related problems mostly occurs at the time prescribing process and DRPs occurs due to inappropriate drug indication and drug dosage. Other factors include age, aphasia, high prevalence of comorbidity, polypharmacy, IV route of administration because of impaired oral intake and drugs which requires frequent monitoring and adjusting dose has high risk of medication errors. Patient receiving heparin, warfarin, antihypertensive medication, and combination of antiplatelet drugs has high risk for drug interactions and medication errors. Clinical pharmacist involvement in the discharge process, computerized order entry of drugs, monitoring drug-drug interactions prior to administration, and patient counseling will improve therapeutic outcome. The pharmaceutical care has a clear benefit and positive impact on patient's health related quality of life (HRQoL) and the patients who do not receive intensified pharmaceutical care have a higher chance of deteriorating HRQoL.

REFERENCES

- 1. Athirajith, chaithanya t kumar, joffeysara joy, kavithasamy p*, sambathkumar r department of pharmacy practice, j.k.knattraja college of pharmacy, kumarapalayam, namakkal, tamilnadu 638 183, india; a prospective study of drug–drug interactions and adverse drug reactions among stroke patients in a tertiary care hospital 2016; https://doi.org/10.1016/j.cegh.2018.05.001
- 2. Dilip Chandrasekhar*, Anuja Pradeep, Asha Susan Geoji, AthiraElezebath George, Athira V., Geethu k. Thomas; Impact of intensified pharmaceutical care on health related quality of life in patients with stroke in a tertiary care hospital 2018; https://doi.org/10.22159/ajpcr.2016
- 3. Iman A Basheti, Shahnaz Mohammed Ayasrah, Muayyad Ahmad on Identifying treatment related problems and associated factors among hospitalized post-stroke patients through medication management review 2018; https://doi.org/10.1016/j.jsps.2018.10.005.
- 4. Carina Hohmann Tobias Neumann-Haefelin , Ju^{*}rgen M. Klotz Annette Freidank, Roland RadziwillInt J; Drug-related problems in patients with ischemic stroke in hospital Clin Pharm (2012) 34:828–831, DOI 10.1007/s11096-012-9690-7
- 5. KannikarSemcharoen, SajjaSupornpun, SurakitNathisuwan, JunpornKongwatcharapong et al., Characteristic of drug related problems and pharmacists interventions in a stroke unit in Thailand, International Journal of Clinical Pharmacy (2019),https://doi.org/10.1007/s11096-019-00832-4.
- 6. Carina Hohmann*1,2, Roland Radziwill2, Juergen M Klotz1, Andreas H Jacobs3 Hohmann*et al.*, Health-Related Quality of Life after Ischemic Stroke: The Impact of Pharmaceutical Interventions on Drug Therapy (Pharmaceutical Care Concept) (2010), http://www.hqlo.com/content/8/1/59.
- Renata Daud-Gallottia, Hillegonda Maria DutilhNovaesc, Maria Ceci'liaLorenzib, Jose' Eluf-Netoc, Mirna NamieOkamurac and IrineuTadeuVelascoa al.., Adverse events and death in stroke patients admitted to the emergency department of a tertiary university hospital (2015). European Journal of Emergency Medicine 12:63–71 _c 2005 Lippincott Williams & Wilkins.
- Celin A.T, Seuma J, Ramesh A* Assessment of Drug Related Problems in Stroke Patients Admitted to a South Indian Tertiary Care Teaching Hospital (2012); Indian Journal of Pharmacy Practice Volume 5 Issue 4 Oct - Dec, 2012
- 9. Kirsten K. Viktil1,3 and Hege Salvesen Blix2,3; The Impact of Clinical Pharmacists on Drug-Related Problems and Clinical Outcomes; 2007, Diakonhjemmet Hospital Pharmacy; LovisenbergDiakonale Hospital; Department of Pharmacotherapeutics, Faculty of Medicine, University of Oslo, Oslo, Norway (Received September 13, 2007; Accepted October 22, 2007)
- 10. Bellolio MF, Finley JL, Flemming KD, et al. Implementation of central page activation for acute ischemic strokes. Stroke. 2009; 40(4):E209.
- 11. Pandya KA, Ruf KM, Winstead PS, Cook AM, Weant KA. Pharmacy resident participation with a 24-hour multidisciplinary stroke response team. Am J Health Syst Pharm. 2010;67(22): 1901-7
- 12. Garland J. Achieving target: stroke honor roll status through the utilization of 9 best practice strategies [abstract]. In: Abstracts from the American Heart Association/American Stroke Association 2013

International Stroke Conference and Nursing Symposium;2013 Feb 5-8; Honolulu, USA. Stroke. 2013;44:AWP359.

- Sreenivasan B. Inpatient stroke education in a rehabilitation hospital: patient evaluation [abstract]. In: 7th World Congress for Neuro rehabilitation; 2012 May 16-19; Melbourne, Australia. Neurorehabil Neural Repair 2012;26(6):A321.
- 14. Scott JS, Sampson EF, D'Amato T. Self-medication in hospitalized stroke patients: effectiveness of medication counseling for right and left hemiplegics [abstract]. Presented at the Annual Meeting of the American Society of Health-System Pharmacists; 1989: P-32E.
- 15. Lindblad AJ, Howorko J. Integration of a pharmacist into a stroke prevention clinic team. Can J Hosp Pharm. 2008;61(6):431-6.
- 16. McAlister FA, Majumdar SR, Padwal RS, et al. Case management for blood pressure and lipid level control after minor stroke: PREVENTION randomized controlled trial. CMAJ. 2014; 186(8):577-84.
- 17. Chiu CC, Wu SS, Lee PY, Huang YC, Tan TY, Chang KC. Control of modifiable risk factors in ischemic stroke outpatients by pharmacist intervention: an equal allocation stratified randomized study. J Clin Pharm Ther. 2008;33(5):529-35.
- 18. Nguyen VV, Poon J, Tokuda L, Sayers J, Wallis R, DergalustS.Pharmacist telephone interventions improve adherence to stroke preventive medications and reduce stroke risk factors: a randomized controlled trial [abstract]. In: International Stroke Conference and Nursing Symposium Poster Presentations presented at the International Stroke Conference; 2011 Feb 9-11; Los Angeles, USA. Stroke. 2011; 42:AWP302.
- 19. Hooker AF, Evans H. Does stroke bridge clinic reduce hospital readmission rates in patients who have suffered a stroke [abstract]. In: International Stroke Conference and Nursing Symposium Poster Abstracts; 2012 Feb 1-3; New Orleans, USA. Stroke.2012;43:A2480.
- 20. Bruner L. Stroke bridge clinic: aiming at reduced readmissions for newly diagnosed stroke and complicated TIA patients [abstract].Presented at the American Society of Health-System Pharmacists Summer Meeting; 2012 Jun 9-13; Baltimore, USA.
- 21. Lee J, Graham C, Silliman S, Merino-Juarez J. Evaluation of stroke patients' knowledge of their medications [abstract]. Presented atthe ASHP Midyear Clinical Meeting; 2004 Dec 4-8; Orlando,USA: A121E.
- 22. Hohmann C, Klotz JM, Radziwill R, Jacobs AH, Kissel T. Pharmaceutical care for patients with ischemic stroke: improving thepatients quality of life. Pharm World Sci. 2009;31(5):550-8.
- 23. Fincham JE, Wallace IA, Ivory A. Ticlopidine compliance enhancement: a community pharmacy based study. J Pharmacoepidemiol.2000;8(1):51-66.
- 24. Horning KK, Hoehns JD, Doucette WR. Adherence to clinical practice guidelines for 7 chronic conditions in long-term-care patients who received pharmacist disease management services versus traditional drug regimen review. J Manag Care Pharm.2007;13(1):28-36
- 25. Wardlaw JM, Murray V, Berge E, delZoppo GJ. Thrombolysis for acute ischaemic stroke. Cochrane Database Syst Rev. 2009:(7):CD000213.
- 26. Jauch EC, Saver JL, Adams HP, et al. Guidelines for the early management of patients with acute ischemic stroke: a guideline for healthcare professionals from the American Heart Association/American Stroke Association. Stroke. 2013;44(3):870-947.
- 27. Lindsay MP, Gubitz G, Bayley M, et al. Canadian Best Practice Recommendations for Stroke Care, Updated 2010. On behalf of he Canadian Stroke Strategy Best Practices and Standards Writing Group. Ottawa: Canadian Stroke Network; 2010.Available at: <u>http://www.strokebestpractices.ca/index.php/news/the-5th-canadian-stroke-congress/.</u>
- 28. Marconi GP, Claudius I. Impact of an emergency department pharmacy on medication omission and delay. PediatrEmerg Care.2012;28(1):30-3.
- 29. Patanwala AE, Sanders AB, Thomas MC, et al. A prospective, multicenter study of pharmacy activities resulting in medication error interception in the emergency department. Ann Emerg Med.2012;59(5):369-73.
- 30. Abu-RamailehAM, Shane R, Churchill W, Steffenhagen A, PatkaJ,Rothschild JM. Evaluating and classifying pharmacists' qualityinterventions in the emergency department. Am J Health Syst Pharm. 2011;68(23):2271-5

- 31. Cesarz JL, Steffenhagen AL, Svenson J, Hamedani AG. Emergency department discharge prescription interventions by emergency medicine pharmacists. Ann Emerg Med. 2013;61(2):209-214.
- 32. Stroke Unit Trialists' Collaboration. Organised inpatient (stroke unit)care for stroke. Cochrane Database Syst Rev. 2007:(4):CD000197.
- 33. Kaboli PJ, Hoth Ab, McClimon BJ, Schnipper JL. Clinical pharmacists and inpatient medical care. Arch Intern Med. 2006;166(9):955-64.
- 34. Stading J, Herrmann J, Walters R, Destache C, Chock A. Impact of pharmacist intervention on diabetes patients in an ambulatory setting. Diabetes Spectr. 2009;22(4):241-6; Available at: http://spectrum.diabetesjournals.org/content/diaspect/22/4/241.full.pdf.
- 35. Sease JM, Franklin MA, Gerrald KR. Pharmacist management of patients with diabetes mellitus enrolled in a rural free clinic. Am Health Syst Pharm. 2013;70(1):43-7.
- 36. Cording MA, Engelbrecht-Zadvomy EB, Pettit BJ, Eastham JH,Sandoval R. Development of a pharmacist-managed lipid clinic.AnnPharmacother. 2002;36(5):892-904.
- 37. Casaubon LK, Suddes M, on behalf of the Acute Stroke Care Writing Group. Acute inpatient stroke care. In: Lindsay MP, GubitzG,Bayley M, Phillips S editors Canadian Best Practice Recommendations for Stroke Care. Ottawa: Heart and Stroke Foundation of Canada, and the Canadian Stroke Network; 2013,
- 38. Quality of Life Among Stroke Survivors Evaluated 1 Year After Stroke Experience of a Stroke Unit; Javier Carod-Artal, MD, PhD; Jose´ Antonio Egido, MD; Jose´ Luis Gonza´lez, MD; E. Varela de Seijas, MD, PhD.

