



Pharmaceutical Care Strategies for Stroke Patients: A Comprehensive Review

**Sainath Chary Gannoji^{a++*}, Soujanya Durishetty^{a++},
Raheemunnissa^{a++}, Vedavathi Sai G.^{b++},
Shivankitha Kattekola^{c#}, Purbai Dheeraj^{d++}
and Parsi Lavanya^{at}**

^a St. Paul's College of Pharmacy, Hyderabad, Telangana, India.

^b East West College of Pharmacy, Bangalore, Karnataka, India.

^c Sri Venkateshwara College of Pharmacy, Hyderabad, Telangana, India.

^d Malla Reddy Pharmacy College, Hyderabad, Telangana, India.

Authors' contributions

This work was carried out in collaboration among all authors. All authors read and approved the final manuscript.

Article Information

Open Peer Review History:

This journal follows the Advanced Open Peer Review policy. Identity of the Reviewers, Editor(s) and additional Reviewers, peer review comments, different versions of the manuscript, comments of the editors, etc are available here: <https://www.sdiarticle5.com/review-history/106669>

Review Article

Received: 24/07/2023

Accepted: 01/10/2023

Published: 11/10/2023

ABSTRACT

Stroke is a clinical syndrome characterized by rapidly developing clinical symptoms of focal (or global, in the case of coma) brain dysfunction lasting more than 24 hours or causing death with no apparent cause other than vascular. Stroke is broadly divided into subarachnoid hemorrhage, ischemic stroke, and hemorrhagic stroke. Stroke is the second leading cause of death and the third leading cause of disability worldwide. Stroke is the most common non-communicable disease

⁺⁺ Pharm D VI Year;

[#] B. Pharm Graduate;

[†] Pharm. D, Assistant Professor;

^{*}Corresponding author: Email: sainathcharygannoji@gmail.com;

(NCD) in India, accounting for 3.5 percent of Disability-Adjusted Life Years (DALY). According to studies, the prevalence of strokes in India is 116-163 strokes per 100,000 people. Recently, the ICMR published a report titled "India: According to the National Health Service, stroke was the fourth leading cause of death and the fifth leading cause of Disability-Adjusted Life Years (DALYs) in 2016". All the literature evaluating pharmacy interventions in stroke patients and the role of the clinical pharmacist in the evidence supporting stroke care are well established. This review aims to provide a comprehensive explanation of pharmacist interventions for stroke patients. Outcomes can be improved by clinical pharmacist involvement in the prescribing process, computerized entry of medication orders, pre-administration monitoring of drug interactions, and patient counseling.

Keywords: Stroke; clinical pharmacists; pharmaceutical care plan; drug related problems.

1. INTRODUCTION

"The World Health Organization defines stroke as a clinical syndrome characterized by rapidly developing clinical signs of focal (or global, in the case of coma) disturbance of cerebral function lasting more than 24 hours or resulting in death with no apparent cause other than a vascular origin"[1]. "Stroke is broadly divided into subarachnoid hemorrhage, ischemic stroke, and hemorrhagic stroke. Ischemic stroke happens because of a blockage of the vein, which restricts the blood supply to the mind, while hemorrhagic stroke happens because of a breakage of a vein driving spillage of blood in the intracranial cavity" [2]. "Worldwide, stroke is the second-leading cause of death and the third-leading cause of disability"[3]. "Globally, 68% of all strokes are ischemic, while 32% are hemorrhagic [3]. The numbers in the United States are slightly different, with 87% being ischemic, 10% being hemorrhagic, and about 3% being subarachnoid hemorrhage" [4,5]. In India, stroke, a major non-communicable disease (NCD), accounts for 3.5 percent of Disability-Adjusted Life Years (DALY).

According to studies, the prevalence of strokes in India is 116-163 strokes per 100,000 people. Recently, the ICMR published a report titled "India: Stroke was the fourth leading cause of death and the fifth leading cause of Disability-Adjusted Life Years (DALYs) in 2016, according to National Health Data". "There is a wealth of literature evaluating pharmacist interventions for stroke patients, and the role of the clinical pharmacist in stroke support is well established. Proven, the purpose of this review is to provide a comprehensive overview of pharmacist interventions with stroke patients. The pharmacist needs a better understanding of drugs and their therapeutic effects because drug regimen development is an ever-evolving process. A treatment plan improves both the patient's health and the ability of healthcare

professionals to prevent disease. The stroke medication plan plays a vital role in differentiating drug problems (DRPs) and investigating the nature and recurrence of DRPs, including untreated symptoms, under-corrected measurements, inappropriate dosing, overt drug use, failure to obtain medications, inappropriate drug prescription, drug interactions and side effects. According to the multicentre Trial of Acute Stroke Treatment (TOAST), there are three kinds of ischemic stroke" [6].

1. Large vessel stroke
2. Small vessel stroke or Lacunar stroke
3. Cardioembolic stroke

Thrombotic or embolic occlusion of the major arteries of the brain, such as the internal carotid artery, middle cerebral artery, anterior cerebral artery, or vertebrobasilar system, may result in large artery strokes. The involvement of smaller or perforating blood vessels that supply the brain's deeper structures is the primary cause of lacunar strokes.

"Time is the most crucial factor in treating an acute ischemic stroke. When an ischemic stroke occurs, the patient loses 190,000 brain cells every minute, approximately 14,000,000 nerve connections every minute, and 12 kilometers (7.5 miles) of nerve fibers every minute. For every hour that the brain is without blood, it ages by 3.6 years"[7]. Acute ischemic stroke can be treated with one of two methods: Mechanical thrombectomy and intravenous thrombolysis.

The next steps must be taken once an acute stroke clinical diagnosis has been made.

- Verify the patient's medical stability.
- Look for neurological symptoms that can be fixed.
- Determine the type of stroke ischemic or hemorrhagic and how to treat it.

- Find out what causes stroke (which goes beyond the scope of this chapter).

Pharmaceutical interventions were defined as medication optimization, where medication management included medication evaluation, medication coordination, medication problem identification, and resolution. Findings are divided into primary and secondary prevention optimization to enable comparison of similar results across different treatment programs.

A person's risk of having a stroke can be assessed using the acronym FAST

Face: invite the person to smile; this helps distinguish dead or missing part of the face.
Hands: Ask the person to raise both hands. Check if one arm has fallen lower than the other.
Speaking: Ask the person to repeat a short sentence.

2. CARE PROVIDED AT DIFFERENT LEVELS OF HEALTHCARE

The Primary Health Center's (PHC) role is limited to risk assessment, prompt recognition of symptoms, stabilization, and referral to higher centers with management facilities due to the complexity of stroke treatment.

- Primary health care: primary prevention, early recognition and referral, rehabilitation.
- Secondary health care: acute stroke management, secondary prevention and follow-up, rehabilitation.
- Tertiary health care: complex and higher-level management of acute cases, follow-up of stroke for enablement and support services and rehabilitation of residual impairment.

3. PHARMACOLOGICAL TREATMENT OF STROKE

"Whether the stroke is ischemic or hemorrhagic, stroke treatment can be divided into two categories: stroke-specific treatment and prevention of stroke".[8] Tissue plasminogen activator (tPA) and antiplatelet agents are pharmacotherapeutic options for primary ischemic stroke under certain conditions. The goal of pharmacotherapeutic treatment of hemorrhagic stroke is to control the patient's blood pressure and intracranial pressure. Acute

ischemic stroke (AIS) is usually treated with alteplasin intravenous thrombolysis. Antiplatelet therapy with aspirin (acetylsalicylic acid) has been shown to reduce the risk of early recurrent stroke when started within 48 hours of an ischemic stroke, but it does not treat a stroke that has already occurred. Aspirin and newer antiplatelet drugs have shown promising results in preventing early recurrence, and clinical trials are ongoing. Patients receiving acute therapeutic anticoagulation with unfractionated heparin (UFH) or low molecular weight heparin (LMWH) have shown no clinical benefit compared with antiplatelet agents in acute ischemic stroke. The role of acute anticoagulation in certain situations with a high risk of early stroke requires further investigation.

"Tissue plasminogen activator (tPA): Alteplase is the only IV Tissue plasminogen activator (tPA) agent approved by the FDA for the treatment of ischemic stroke. This indication was approved once the drug's effectiveness in the NINDS rt-PA Stroke Study was established. The International Stroke Registry's Safe Implementation of Thrombolysis in Stroke (SITS-ISTR) A 3-to-4.5-hour research found no differences in mortality, a modified Rankin Scale score of 0 to 2, or symptomatic intracerebral hemorrhage between individuals treated within 3 hours and those treated within 3-to-4.5 hours"[9,10]. "Despite the fact that stroke patients who qualify for IV Tissue Plasminogen Activator (tPA) should be treated within 3 hours, the established guidelines for stroke treatment Seizures, new ischemic strokes, cerebral oedema, and brain herniation have all been associated with the use of alteplase for ischemic stroke"[11].

4. MEDICINES OPTIMISATION FOR PRIMARY PREVENTION OF STROKE

"Three main prevention studies looked at the safety and effectiveness of adjusting anticoagulant and antithrombotic drugs for people at risk of stroke".[12,13,14] A pharmacist intervention identified 78 of 218 (35.6%) needed adjustments to optimize the current antithrombotic medication for patients hospitalized at risk of stroke (e.g., with AF), in accordance with locally developed evidence-based guidelines. To reduce the risk of stroke, prophylactic medications[14] that may be less effective but safer had to be prescribed in 60 (76.9%) of the instances. Similar to this, a hospital's 134 patients with AF participated in a pharmacist-led stroke risk assessment program

that increased warfarin use from 74% at admission to 98% at discharge. "Of the 50 recommendations for medication change, 44 (80%) were accepted and put into practice. More effective therapy was needed in 30 out of 44 instances (or 68%)" [15]. "According to the clinical audit, 382 anticoagulant and antithrombotic optimization suggestions for patients with AF received approval by general practitioners in 77% of cases" [13].

Men whose risk is sufficiently high (10-year risk > 10%) for the benefits to outweigh the risks associated with treatment can use aspirin daily (50-100 mg) for cardiovascular events, including stroke prophylaxis. The 10-year risk can be calculated using a cardiovascular risk calculator; however, there is no calculator designed specifically for India. Utilizing clinical judgment and the WHO calculator, risk can be assessed

Because aspirin has been associated with an increased risk of bleeding, it should only be taken when absolutely required. Utilizing clinical judgment and the WHO calculator, risk can be assessed. Through ongoing Indian studies, risk calculators customized for Indians will be created in the upcoming years.

Post-menopausal women, including those with diabetes mellitus, whose risk [14] is sufficiently high for the benefits to outweigh the risk associated with treatment can benefit from taking aspirin (50 mg to 100 mg daily). Its use should be clinically examined for draining potential.

The therapeutic outcome will be improved by clinical pharmacist involvement in the discharge process, computerized drug order entry, monitoring drug-drug interactions prior to administration, and patient counseling [16].

Various studies have shown that pharmaceutical interventions improve patient outcomes. Patients with dual antiplatelet therapy and a high bleeding risk may benefit from drugs such as the addition of omeprazole, a reduction in the dose of piperacillin or tazobactam from 4.5 g IV 6 h to 2.25 g IV 6 h in patients with creatinine clearance less than 20 ml/min. increasing the dose of atorvastatin from 20 mg to 40 mg for better LDL cholesterol reduction Physicians agreed with 84% of the top clinical pharmacists [17].

Most of the side effects were in the WHO "probable" category. None of the side effects identified were "probably preventable" or "eventual" and most were "mild" in severity. Involving the pharmacist in the evaluation of medication-related issues improves the patient's quality of life, especially for elderly multi-pharmacy patients. "The primary goal of the pharmacy should be to locate potential drug-related problems. In the majority of cases, the drug therapy is being altered, and these recommendations were widely accepted. Polypharmacy was found to be a potential risk factor for drug-related issues. The therapeutic outcomes are enhanced by involving pharmaceutical care for the early detection of drug-related issues" [18].

Table 1. The indicative List of Drugs under NPCDCS for emergency care is as follows:[16]

ACE Inhibitors	Enalapril, Ramipril, Lisinopril, Captopril
Calcium channel blocker	Amlodipine
Diuretics	Indapamide, Chlorthalidone, Frusemide, Hydrochlorothiazide
Aldosterone antagonist Beta blocker	Atenolol, Metoprolol, Labetalol
Oral Hypoglycemic	Metformin, Gliclazide, Glibenclamide
Insulin	Short, Intermediate, Long-acting
Fibrinolytics	Streptokinase, rTPA (alteplase and Tenecteplase)
Anti-platelet	Clopidogrel
Lipid-lowering	Statins

Table 2. Adverse drug reactions to drugs [16]

Drugs	Adverse drug reaction
Aspirin plus dipyridamole, Cilostazol	Headache, Palpitation, Tachycardia
Nicardipine	Phlebitis
Warfarin	Upper gastrointestinal bleeding
Hydralazine, Carvedilol	Hypotension

Table 3. Different drug-related problems

Drug use without indication	Domperidone, Piperacillin+tazobactam, Ondansetron, Paracetamol, Pantoprazole, Tramadol
Drug Duplication	Clopidogrel, Pantoprazole
Subtherapeutic Dose	Amlodipine, Telmisartan, 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

5. CONCLUSION

Patients' health-related quality of life (HRQoL) has been shown to improve as a result of hospital care, as shown by various studies. Research also shows that men are more likely to have a stroke because of their social habits and that those aged 41-70 are at the highest risk. The most important drug interactions observed in this study increased both the likelihood of hospitalization and healthcare costs. Antiplatelet agents, anticoagulants, and antihypertensive medications were the most common interacting medications. The use of antihypertensive drugs significantly increases the probability of side effects. Most drug-related problems occur during the prescribing process and DRP due to the wrong indication and dosage of the drug. Age, aphasia, high comorbidity, polytherapy, and the intravenous (IV) route of administration are additional factors that increase the risk of medication errors due to impaired oral administration. A patient receiving a combination of heparin, warfarin, blood pressure medication, and platelets is at high risk for drug interactions and prescribing errors. Therapeutic outcomes are improved through clinical pharmacist participation in the prescribing process, computerized entry of medication orders, monitoring of drug interactions prior to administration, and patient counseling. Patients who do not receive intensified drug therapy are more likely to experience a decline in health-related quality of life (HRQoL). Drug treatment has clear benefits and positive effects on patient HRQoL.

CONSENT

It is not applicable.

ETHICAL APPROVAL

It is not applicable.

COMPETING INTERESTS

Authors have declared that no competing interests exist.

REFERENCES

1. Hatano S. Experience from a multicentre stroke register: a preliminary report. *Bulletin of the World Health Organisation*. 1976;54((5)):541-553.. [PMC free article] [PubMed] [Google Scholar]
2. Caplan LR. *Caplan's Stroke: A Clinical Approach*, 4th ed, Saunders Elsevier, Philadelphia:. Basic pathology, anatomy, and pathophysiology of stroke. 2009;22. [Google Scholar]
3. Lozano R, Naghavi M, Foreman K, Lim S, Shibuya K, Aboyans V, et al. Global and regional mortality from 235 causes of death for 20 age groups in 1990 and 2010: a systematic analysis for the Global Burden of Disease Study 2010. *Lancet*. 2012;380:2095.. [PubMed] [Google Scholar]
4. Krishnamurthi RV, Feigin VL, Forouzanfar MH, Mensah GA, Connor M, Bennett DA, et al. Global and regional burden of first-ever ischaemic and hemorrhagic stroke during 1990-2010: findings from the Global Burden of Disease Study 2010. *Lancet Glob Health*. 2013;1:e259. [PMC free article] [PubMed] [Google Scholar]
5. Benjamin EJ, Blaha MJ, Chiuve SE, Cushman M, Das SR, Deo R, et al. Heart Disease and Stroke Statistics-2017 Update: A Report From the American Heart Association. *Circulation*. 2017;135:e146. [PMC free article] [PubMed] [Google Scholar]
6. Adams HP, Jr., Davis PH, Leira EC, Chang KC, Bendixen BH, Clarke WR, et al.

- Baseline NIH Stroke Scale score strongly predicts outcome after stroke: A report of the Trial of Org 10172 in Acute Stroke Treatment (TOAST). *Neurology*. 1999; 13;53(1):126—131.. [PubMed] [Google Scholar]
7. Scott JS, Sampson EF, D'Amato T. Self-medication in hospitalized stroke patients: effectiveness of medication counselling for right and left hemiplegics [abstract]. Presented at the Annual Meeting of the American Society of Health-System Pharmacists; 1989:32E.
 8. Jackson SL, Peterson GM. Stroke risk assessment for atrial fibrillation: Hospital-based stroke risk assessment and intervention program. *J. Clin. Pharm. Ther.* 2011;36:71–79. [Google Scholar] [CrossRef] [PubMed]
 9. del Zoppo GJ, Saver JL, Jauch EC, Adams HP Jr. Expansion of the time window for treatment of acute ischemic stroke with intravenous tissue plasminogen activator: a science advisory from the American Heart Association/American Stroke Association. *Stroke*. 2009;40:2945-2948.
 10. Adams HP Jr, del Zoppo G, Alberts MJ. Guidelines for the early management of adults with ischemic stroke: a guideline from the American Heart Association/American Stroke Association Stroke Council, Clinical Cardiology Council, Cardiovascular Radiology and Intervention Council, and the Atherosclerotic Peripheral Vascular Disease and Quality of Care Outcomes in Research Interdisciplinary Working Groups: the American Academy of Neurology affirms the value of this guideline as an educational tool for neurologists. *Stroke*. 2007;38:1655-1711.
 11. Activase (alteplase) package insert. South San Francisco, CA: Genentech, Inc; April 2011.
 12. Bajorek BV, Krass I, Ogle SJ, Duguid MJ, Shenfield GM. Optimizing the use of antithrombotic therapy for atrial fibrillation in older people: A Pharmacist-Led Multidisciplinary Intervention. *J. Am. Geriatr. Soc.* 2005;53:1912–1920. [Google Scholar] [CrossRef] [PubMed]
 13. Virdee MS, Stewart D. Optimizing the use of oral anticoagulant therapy for atrial fibrillation in primary care: A pharmacist-led intervention. *Int. J. Clin. Pharm.* 2017;39: 173–180. [Google Scholar] [CrossRef]
 14. Bajorek BV, Krass I, Ogle SJ, Duguid MJ, Shenfield GM. Optimizing the use of antithrombotic therapy for Atrial Fibrillation in Older People: A Pharmacist-Led Multidisciplinary Intervention. *J. Am. Geriatr. Soc.* 2005;53:1912–1920. [Google Scholar] [CrossRef] [PubMed]
 15. Jackson SL, Peterson GM. Stroke risk assessment for atrial fibrillation: Hospital-based stroke risk assessment and intervention program. *J. Clin. Pharm. Ther.* 2011;36:71–79. [Google Scholar] [CrossRef] [PubMed]
 16. Carina Hohmann, Tobias Neumann-Haefelin, Juergen M. Klotz, Annette Freidank, Roland Radziwill J; Drug-related problems in patients with ischemic stroke in hospital *Clin Pharm.* 2012; 34:828–831. DOI 10.1007/s11096-012-9690-7
 17. Kannikar Semcharoen, Sajja Supornpun, Surakit Nathi suwan, Junporn Kongwat charapong 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>.
 18. Celin A.T, Seuma J, Ramesh A* Assessment of drug related problems in stroke patients admitted to a South Indian Tertiary care teaching hospital. *Indian Journal of Pharmacy Practice* .2012;5(4).

© 2023 Gannoji et al.; This is an Open Access article distributed under the terms of the Creative Commons Attribution License (<http://creativecommons.org/licenses/by/4.0>), which permits unrestricted use, distribution, and reproduction in any medium, provided the original work is properly cited.

Peer-review history:
The peer review history for this paper can be accessed here:
<https://www.sdiarticle5.com/review-history/106669>