

**“Association of Depression with Polypharmacy among Diabetic Patients of Geriatric age group in Tertiary Care Teaching Hospital.”**

**ABSTRACT**

**Background:** Diabetes mellitus, one of the common serious conditions among elderly and depression is frequent finding as a Comorbidity. Polypharmacy is common in diabetes and In diabetic senior people, it has been linked to depression. Polypharmacy, especially among the elderly, is a consistent predictor of inappropriate medication. Approaching and analysing the prescription pattern in elderly diabetic patient will rationalize the drug utilization. The current research is to access the co-relation of polypharmacy in geriatric diabetic patient and accompanying depression.

**Objectives:** To assess depression and polypharmacy and correlate their association, in diabetic patients of geriatric age group.

**Methodology:** This is questionnaire based cross sectional observational study. This shall be conducted in geriatric population attending Medicine department (OPD and IPD) (AVBRH), DMIMS, Sawangi (M), Wardha, Maharashtra, India. The depression will be assessed using Geriatric Depression scale. The Polypharmacy will be assessed using contingency table and odds ratio.

**Expected outcome:** It is hypothesized that geriatric diabetic patients with depression (GDS $\geq$  6) are at higher risk of encountering polypharmacy.

**Conclusion:** To upgrade standards of life of senior diabetic patients, prescribing procedures should be reviewed.

**Keywords:** Depression, Diabetes, Polypharmacy, Geriatric depression scale, Geriatric.

**INTRODUCTION:**

264 million peoples all around the world suffer from Depression which is common mental illness nowadays <sup>1</sup> According to the W.H.O, depression is a mental condition marked by dejection, aloof, detachment, guilt, interrupted sleep, poor attention, and exhaustion. (2) Diabetes, a epidemic in India, now has 62 million diabetics, with the data expected to increase to 72.4 million by 2030. (3) Diabetes is termed as a metabolic disease in which blood glucose levels are elevated and by the time it damages the blood vessels, kidney, heart and nerves. Type II Diabetes, the most common one in which the body doesn't respond to insulin, it gets resisted or there is lack of insulin production. India stands 2<sup>nd</sup> highest in no. of diabetic patients in world and by the year 2025 it is expected to reach 69.9 million diabetic population in India. <sup>(4)</sup> Individual with depression and diabetes has showed poor connection to exercise and dietary restrictions, poor self-management, and poor medication adherence when compared to patients with diabetes alone (5,6). While depression may play a role in low statistics in diabetes, and its consequences may also play a role in depression. (7,8,9). Diabetes Type II is becoming more common in countries of all income levels over the last three decades. Depression affects around 43 million people worldwide. (10)

Diabetes is a global threat of the twenty-first century, according to "International Diabetic Federation." (11). In developed countries, depression and anxiety are 4<sup>th</sup> and 8<sup>th</sup> causes of disability adjusted life years, respectively. (12). Diabetes will cause 6-7 million fatalities in 2021, with one death occurring every 5 seconds. (13) Aging is an unavoidable fact of life. It is natural and uncontrollable by humans. The term "Geriatric" refers to people in their senior years, and Geriatric or Geriatric medicine is involved with the diagnosis, therapy, and prevention of disease in the elderly. (14,15). Normal ageing and aging-related disease are both a part of life. Physiological functions are typically lowered as a result of age, such as decreased bone density, osteoarthritis, and cataract lens discomfort, among other things. Decreased vital function in the older people makes them more susceptible to numerous ailments, thus they will be prescribed a variety of drugs. (16,17)

Geriatric syndromes are common in the elderly, and they can be caused by using the wrong drugs. (18). Because of neuropathy, dysautonomia, and vascular disease, diabetic patients with long-term diabetes are particularly vulnerable to all of these adverse effects.

According to one data of Medical Expenditure of Panel survey, the Diabetic patient has minimum one comorbid chronic condition, and as many as 40% have at least three. (19,20,21)

Diabetes and depression have a complicated relationship. Depression is widespread among senior diabetic people, they are at a higher risk for co-morbid depression than non-diabetic patients. (22)

Polypharmacy is routine practice among diabetic patients and is more common in geriatric population with diabetes.(23,24)

Polypharmacy is explained as concomitant use of 5 or >5 drugs for management of co-existing health related issues, and there are some evidenced based data indicating the association of polypharmacy with increased rates of drug interaction and adverse drug events (25,26,27,28)

Inappropriate prescription of drug leads to decrease in medical compliance.

According to current estimates, 350 million people worldwide go through from depression, whereas > 400 million people worldwide suffer from DM. (29)

Only about 1/3rd of persons with depression and diabetes are diagnosed and treated adequately. (30)

There are some studies about polypharmacy and its health-related hazards, but as per as our knowledge is concerned there are no study taking consideration of polypharmacy as association with depression in geriatric diabetic patients. Hence, we are planning to do present study.

### **RATIONALE:**

The relationship of depression with diabetes has been recorded years ago, but the kind of this connection remains undefined. Both diseases are serious and chronic in nature that adversely affect life quality, functional ability, and diminish overall life expectancy.(31,32)

Epidemiological evidence suggests that the prevalence of depression is higher in both type 1 and type 2 diabetic patients compared to population across the world. (33)

When compared to people who do not have diabetes, geriatric diabetic patients are at a high risk of developing co-morbid depression. (34)

Geriatric syndromes are common among the elderly, and they are often coupled with improper drugs, leading to polypharmacy. (35) Polypharmacy is evident in diabetic patients in order to maintain tight glucose control while also managing comorbid illnesses such as coronary artery disease, dyslipidemia, hypertension, and neuropathy. (36)

Evidence based data indicates the association of polypharmacy with increased rates of drug interaction and ADEs. (37,38,39)

In the years ahead, the increasing number of multi-morbid elder adults with complex treatment regimens is continued to provide a drawback to the health-care system.

As a result, it makes sense to look into the link between depression and polypharmacy among diabetic patients in their later years.

**AIM:** To find out association of depression with polypharmacy among diabetic patients of geriatric age group.

**OBJECTIVES:**

1. To assess the Depression in diabetic patient of geriatric age group
2. To observe the Polypharmacy in diabetic patient of geriatric age group
3. To correlate the association of depression and polypharmacy in diabetic people of geriatric age group.

**METHODOLOGY:**

**Study Setting:** The research will begin once the Institutional Ethical Committee has given its approval. All participants must give their informed consent.

It will be 2 months questionnaire based, cross sectional, observational study. It will be conducted in geriatric population attending Medicine department (OPD and IPD) of (AVBRH), DMIMS, sawangi (M), Wardha, Maharashtra, India.

**Study Design:** Cross Sectional study.

**Sampling Method:** Simple random sampling

**Study Population:**

Geriatric population attending Medicine department (OPD and IPD) of (AVBRH), DMIMS, Sawangi (M), Wardha, Maharashtra, India.

**Inclusion Criteria:**

1. Geriatric patients having diabetes attending Medicine department (OPD and IPD).
2. Elderly diabetic patients willing to give consent for participation in the study.

**Exclusion Criteria:**

1. Patients whose case sheets are incomplete.
2. Patients who are in Emergency and intensive care unit.

3. Patients having serious illness, malignancies and other complications which are not related.

### **Data Collection:**

The examination of the study participants' case sheets will collect all relevant data concerning demographics (age, gender), clinical information (medical diagnosis, nature of co-morbidity, number of co-morbid illnesses), and pharmacological specifics (prescription drugs, quantity of drugs).

Throughout the study period, the study investigator will check the case sheets of these study participants on a daily basis.

A total of 15 score GDS score is used to evaluate depression, with a GDS score of 6 indicating depression. (20)

Out of 15 questions, 10 signifies the presence of depression when answered positively, and the rest (1,5,7,11,13) signifies depression when answered negatively. Scores 0-4 considered normal, score 5-8 considered mild, score 9-11 considered moderate, and score 12-15 considered severe depression.

Polypharmacy will be accessed by using contingency table and odd ratio to compare clinical features of patients that whether patients use less than 5, 5 or more medications daily in the month prior to admission.

Age group are divided as 60-65 years, 65-70 years and more than 70 years.

### **Statistical Analysis:**

Contingency tables will be used for comparison of the clinical, demographic features of patients. By using SPSS software, statistical analysis will be carried out. Descriptive analyses of age, sex, associated problems will be carried out. Continuous variables which are ordinarily distributed will be depicted by using Standard, Mean error. Comparison between quantitative variables will be done using chi-square test.

### **EXPECTED RESULTS:**

Depression is expected co-morbid condition in older adult patients with diabetes. Majority of these diabetic patients can have GDS >6 score, which are strongly associated with depression.

Depression is expected to be more significant in female patients as compared to male patients and also people residing in rural area and those with presence of complications, comorbidities.

Age group of 65-70 years of patients are expected to have high degree of depression.

Many of the patients might be the consumer of 5 or more drugs and more frequently be depressed than consumer of less than 5 drugs.

Polypharmacy should be a top priority for diabetic people, especially the geriatrics. Patients having diabetes need strategies to limit the prescription of unneeded drugs, which could enhance their quality of life and adherence to therapy.

Early detection and treatment of depression can lower mortality rates.

As a result, our research can show that there is a statistically significant link between depression and polypharmacy in elderly DM patients.

## **DISCUSSION:**

This article is to assess the association between polypharmacy and depression in the diabetic people of geriatric age group.

Evidence shows the cases of depression is increased in the previously diagnosed DM patients compared to ordinary glucose metabolism fellows. (40), moderately increased in undiagnosed patient and prediagnosed diabetes patients.

Lower prevalence rates of depression is seen in DM patients according to research in UK, Germany and Canada. (41,42,43)

A research by Gendelman et.al shows remarkably higher rates of depression in both women and men with type 1 diabetes compared to those who doesn't had diabetes (44).. Similarly, increased rates of depression in people were shown by Collins et al. in their study (45)

Another study by pouwer et.al, concerning diabetic patients attending OPD in the Netherlands, found that one third of their sample of type 1 diabetes reported depression.(46)

A Study in Italy shows prevalence of polypharmacy to be 46% in the elderly aged group, i.e 65 years and older.(47)

Study by Golden et al. shows rate of depression is higher in older diabetic people as compared to younger.(48)

Most studies have suggested that numerous factors may be linked with increased prevalence rate of depression among diabetic people and out of these , persistent poor glycemic control has been to be important one<sup>(49)</sup>

A Review article by Good et al. suggests that strict control of glycaemic levels and comorbid illness mainly responsible for polypharmacy among geriatric diabetic patients.(50)

Even after accounting for the effects of obesity and adipose tissue distribution, the Baltimore study of ageing found that secretion of insulin decreased with age. (51) It demonstrates that diabetes mellitus is more prevalent in the geriatric age group.

Geriatrics use an average of 2-9 drugs every day, according to several studies conducted around the world. (52) Inappropriate medication use by geriatric patients was reported to be between 11.5 to 62.5 percent. (53)

Type 2 diabetic patients consumed much more drugs, at significantly higher costs, than non-diabetic patients matched by sex, age, and residential location in a national study in Finland.(54)

A multi-centre cross-sectional survey conducted in Italy, 57 percent of diabetic patients use five or more drugs.(55) Polypharmacy was also detected in 84 percent of diabetic individuals aged 65 and over.

Diabetes, doubles the risk of comorbid depression, which affects about 30% of patients with type 1 and DM patients.(56)

Depression found to be more in female because of estrogen level (57). It happens twice frequently in females compared to males.(58,59-62,63) reason could also be social role attributed to women.(64)

Some studies demonstrated, depression is more common in humans over the age of 60, owing to the fact that they are retired and rely on other family members; nevertheless, other studies reveal no link between age and the statistics of depression in diabetic patients. (65)

Similar studies by Raval et al, Ciechanowski et al. reveal that in rural areas depression is more prevalent than in metropolitan areas (66-75)

Depression could be a barrier to good DM treatment since it leads to patient non-adherence to medication.

Screening is mandatory for depression and other mental disorder for diabetic patients, so that it can be treated early.

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## **REFERENCES:**

1. James SL, Abate D, Abate KH, Abay SM, Abbafati C, Abbasi N, et al. Global, regional, and national incidence, prevalence, and years lived with disability for 354 Diseases and Injuries for 195 countries and territories, 1990-2017: A systematic analysis for the Global Burden of Disease Study 2017. *Lancet* [Internet]. 2018 Nov 10 [cited 2020 Sep 17];392(10159):1789–858. Available from: <https://github.com/ihmeuw/>
2. Marcus M, Yasamy MT, van Ommeren, M, Chisholm D, Saxena S. Depression: A global public health concern. Geneva, Switzerland: WHO Department of Mental Health and Substance Abuse; 2012. Available from: [http://www.who.int/mental\\_health/management/depression/who\\_paper\\_depression\\_wfmh\\_2012.pdf](http://www.who.int/mental_health/management/depression/who_paper_depression_wfmh_2012.pdf).
3. Kaveeshwar SA, Cornwall J. The current state of diabetes mellitus in India. *Australas Med J* [Internet]. 2014 [cited 2020 Sep 18];7(1):45–8. Available from: <https://www.ncbi.nlm.nih.gov/pmc/articles/PMC3920109/>
4. King H, Auburt RE, Herman WH. Global burden of diabetes 1995-2025: Prevalence, numerical estimates, and projections. *Diabetes Care* 1998;21:1414-31.
5. Lin EH, Katon W, Von Korff M, Rutter C, Simon GE, Oliver M, et al. Relationship of depression and diabetes self-care, medication adherence, and preventive care. *Diabetes Care* 2004;27:2154-60.
6. Eraker SA, Kirscht JP, Becker MH. Understanding and improving patient compliance. *Ann Intern Med* 1984;100:258-68.
7. McHale M, Hendrikz J, Dann F, Kenardy J. Screening for depression in patients with diabetes mellitus. *Psychosom Med* 2008;70:869-74
8. Al-Amer RM, Sobeh MM, Zayed AA, Al-Domi HA. Depression among adults with diabetes in Jordan: Risk factors and relationship to blood sugar control. *J Diabetes Complications* 2011;25:247-52.
9. Black SA, Markides KS, Ray LA. Depression predicts increased incidence of adverse health outcomes in older Mexican americans with type 2 diabetes. *Diabetes Care* 2003;26:2822-8.

10. Ivbijaro GO. Mental health and chronic physical illnesses: The need for continued and integrated care - World Mental Health Day 2010 [Internet]. Vol. 7, Mental Health in Family Medicine. Radcliffe Publishing and Wonca; 2010 [cited 2020 Sep 17]. p. 127. Available from: [www.wfmh.org/00WorldMental](http://www.wfmh.org/00WorldMental)
11. World health organisation, International diabetic federation.
12. International diabetic federation.
13. Varma S, Sareen H, Trivedi JK. The geriatric population and psychiatric medication. *Mens Sana Monogr* [Internet]. 2010 Jan 1 [cited 2020 Sep 17];8(1):30–52. Available from: [/pmc/articles/PMC3031932/?report=abstract](http://pmc/articles/PMC3031932/?report=abstract)
14. Nath A, Ingle G. Geriatric health in India: Concerns and solutions. *Indian J Community Med* [Internet]. 2008 [cited 2020 Sep 18];33(4):214. Available from: <http://www.ijcm.org.in/text.asp?2008/33/4/214/43225>
15. Nobili A, Garattini S, Mannucci PM. Multiple disease and polypharmacy in the elderly: Challenges for the internist of the third millennium *J comorbidity* 2011;1:28-44
16. Lavan AH, Gallagher PF, O'Mahony D. Methods to reduce prescribing error in elderly patients with multimorbidity. *Clin interv aging* 2016;11:857-66.
17. Hanlon JT, Fillenbaum GG, Schmader KE, Kuchibhatla M, Horner RD: Inappropriate medication use among community dwelling elderly residents. *Pharmacotherapy* 20:575-582,2000.
18. Druss BG, Marcus SC, Olfson M, Tanielian T, Elinson L, Pincus HA. Comparing the national economic burden of five chronic conditions. *Health Aff* [Internet]. 2001 [cited 2020 Sep 18];20(6):233–41. Available from: <https://pubmed.ncbi.nlm.nih.gov/11816664/>
19. Wolff JL, Starfield B, Anderson G. Prevalence, expenditures, and complications of multiple chronic conditions in the elderly. *Arch Intern Med* [Internet]. 2002 Nov 15 [cited 2020 Sep 18];162(20):2269–76. Available from: <https://pubmed.ncbi.nlm.nih.gov/12418941/>
20. Maddigan SL, Feeny DH, Johnson JA. Health-related quality of life deficits associated with diabetes and comorbidities in a Canadian National Population Health Survey. *Qual Life Res* [Internet]. 2005 Jun [cited 2020 Sep 18];14(5):1311–20. Available from: <https://pubmed.ncbi.nlm.nih.gov/16047506/>.
21. Anderson RJ, Freedland KE, Clouse RE, Lustman PJ. The prevalence of comorbid depression in adults with diabetes: A meta-analysis. *Diabetes Care* [Internet]. 2001 [cited 2020 Sep 18];24(6):1069–78. Available from: <https://pubmed.ncbi.nlm.nih.gov/11375373/>
22. Srivatsa S, Bose D, M S, C.R. J. A prospective cohort study to evaluate the correlates of polypharmacy and its association with depression among elderly patients. *Int J Basic Clin Pharmacol* [Internet]. 2016 Jan 10 [cited 2020 Sep 18];5(5):1984–90. Available from: [www.ijbcp.com](http://www.ijbcp.com)
23. Alwhaibi M, Balkhi B, Alhawassi TM, Alkofide H, Alduhaim N, Alabdulali R, et al. Polypharmacy among patients with diabetes: A cross-sectional retrospective study in a tertiary hospital in Saudi Arabia. *BMJ Open* [Internet]. 2018 May 1 [cited 2020 Sep 19];8(5). Available from: [/pmc/articles/PMC5988096/?report=abstract](http://pmc/articles/PMC5988096/?report=abstract)
24. Bourgeois FT, Shannon MW, Valim C, Mandl KD. Adverse drug events in the outpatient setting: An 11-year national analysis. *Pharmacoepidemiol Drug Saf* [Internet]. 2010 Sep [cited 2020 Sep 18];19(9):901–10. Available from: [/pmc/articles/PMC2932855/?report=abstract](http://pmc/articles/PMC2932855/?report=abstract)
25. Burgess CL, Holman CDAJ, Satti AG. Adverse drug reactions in older Australians, 1981- 2002. *Med J Aust* [Internet]. 2005 Mar 21 [cited 2020 Sep 18];182(6):267–70.

Available from: <https://www.mja.com.au/journal/2005/182/6/adverse-drug-reactions-older-australians-1981-2002>

26. Johnell K, Klarin I. The relationship between number of drugs and potential drug-drug interactions in the elderly: A study of over 600 000 elderly patients from the Swedish prescribed drug register. *Drug Saf* [Internet]. 2007 [cited 2020 Sep 18];30(10):911–8. Available from: <https://pubmed.ncbi.nlm.nih.gov/17867728/>
27. Haider SI, Johnell K, Thorslund M, Fastbom J. Trends in polypharmacy and potential drug-drug interactions across educational groups in elderly patients in Sweden for the period 1992 - 2002. *Int J Clin Pharmacol Ther* [Internet]. 2007 [cited 2020 Sep 18];45(12):643–53. Available from: <https://pubmed.ncbi.nlm.nih.gov/18184532/>
28. WHO Diabetes fact sheet 2017. WHO Depression fact sheet 2012. Available at: <http://www.who.int/mediacentre/factsheets/fs369/en/>. Updated February 2017. Accessed November 8, 2017.
29. Lustman PJ, Harper GW. Nonpsychiatric physicians' identification and treatment of depression in patients with diabetes. *Compr Psychiatry* 1987;28:22-7.
30. Goetzel RZ, Hawkins K, Ozminkowski RJ, Wang S. The health and productivity cost burden of the "top 10" physical and mental health conditions affecting six large U.S.employers in 1999. *J Occup Environ Med* [Internet]. 2003 Jan 1 [cited 2020 Sep 20];45(1):5–14. Available from: <https://pubmed.ncbi.nlm.nih.gov/12553174/>
31. O'Connor PJ, Crain AL, Rush WA, Hanson AM, Fischer LR, Kluznik JC. Does diabetes double the risk of depression? *Ann Fam Med* [Internet]. 2009 [cited 2020 Sep 20];7(4):328–35. Available from: <https://pubmed.ncbi.nlm.nih.gov/19597170/>
32. Leshner EL, Berryhill JS. Validation of the geriatric depression scale- short form among inpatients. *J Clin Psychol* [Internet]. 1994 [cited 2020 Sep 20];50(2):256–60. Available from: <https://pubmed.ncbi.nlm.nih.gov/8014251/>
33. Anderson RJ, Freedland KE, Clouse RE, Lustman PJ. The prevalence of comorbid depression in adults with diabetes: A meta-analysis. *Diabetes Care* [Internet]. 2001 [cited 2020 Sep 18];24(6):1069–78. Available from: <https://pubmed.ncbi.nlm.nih.gov/11375373/>
34. Saraf AA, Petersen AW, Simmons SF, Schnelle JF, Bell SP, Kripalani S, et al. Medications associated with geriatric syndromes and their prevalence in older hospitalized adults discharged to skilled nursing facilities. *J Hosp Med* [Internet]. 2016 Oct 1 [cited 2020 Sep 19];11(10):694–700. Available from: <https://pubmed.ncbi.nlm.nih.gov/27111111/>
35. Good CB. Polypharmacy in Elderly Patients With Diabetes. Vol. 15, *Diabetes Spectrum*. 2002.
36. Bourgeois FT, Shannon MW, Valim C, Mandl KD. Adverse drug events in the outpatient setting: An 11-year national analysis. *Pharmacoepidemiol Drug Saf* [Internet]. 2010 Sep [cited 2020 Sep 18];19(9):901–10. Available from: <https://pubmed.ncbi.nlm.nih.gov/20311111/>
37. Burgess CL, Holman CDAJ, Satti AG. Adverse drug reactions in older Australians, 1981- 2002. *Med J Aust* [Internet]. 2005 Mar 21 [cited 2020 Sep 18];182(6):267–70. Available from: <https://www.mja.com.au/journal/2005/182/6/adverse-drug-reactions-older-australians-1981-2002>.
38. Johnell K, Klarin I. The relationship between number of drugs and potential drug-drug interactions in the elderly: A study of over 600 000 elderly patients from the Swedish prescribed drug register. *Drug Saf* [Internet]. 2007 [cited 2020 Sep 18];30(10):911–8. Available from: <https://pubmed.ncbi.nlm.nih.gov/17867728/>

39. Chen S, Zhang Q, Dai G, Hu J, Zhu C, Su L, Wu X. Association of depression with pre-diabetes, undiagnosed diabetes, and previously diagnosed diabetes: a meta-analysis. *Endocrine*. 2016
40. Brown LC, Majumdar SR, Newman SC, Johnson JA. Type 2 diabetes does not increase risk of depression. *CMAJ* [Internet]. 2006 Jul 4 [cited 2020 Sep 20];175(1):42–6. Available from: </pmc/articles/PMC1482744/?report=abstract>
41. Icks A, Kruse J, Dragano N, Broecker-Preuss M, Slomiany U, Mann K, et al. Are symptoms of depression more common in diabetes? Results from the Heinz Nixdorf Recall study. *Diabet Med* [Internet]. 2008 Nov [cited 2020 Sep 20];25(11):1330–6. Available from: <https://pubmed.ncbi.nlm.nih.gov/19046224/>
42. Paddison CAM, Eborall HC, French DP, Kinmonth AL, Prevost AT, Griffin SJ, et al. Predictors of anxiety and depression among people attending diabetes screening: A prospective cohort study embedded in the ADDITION (Cambridge) randomized control trial. *Br J Health Psychol* [Internet]. 2011 Feb [cited 2020 Sep 20];16(1):213–26. Available from: <https://pubmed.ncbi.nlm.nih.gov/21226792/>
43. Gendelman N, Wadwa RP, Snell-Bergeon JK, Bishop F, McFann K, Rewers M, et al. Prevalence and correlates of depression in individuals with and without type 1 diabetes. *Diabetes Care* [Internet]. 2009 Apr [cited 2020 Sep 20];32(4):575–9. Available from: <https://pubmed.ncbi.nlm.nih.gov/19171719/>
44. Collins MM, Corcoran P, Perry IJ. Anxiety and depression symptoms in patients with diabetes: Original Article: Psychology. *Diabet Med* [Internet]. 2009 Feb [cited 2020 Sep 20];26(2):153–61. Available from: <https://pubmed.ncbi.nlm.nih.gov/19236618/>
45. Pouwer F, Geelhoed-Duijvestijn PHLM, Tack CJ, Bazelmans E, Beekman AJ, Heine RJ, et al. Prevalence of comorbid depression is high in out-patients with Type 1 or Type 2 diabetes mellitus. Results from three out-patient clinics in the Netherlands. *Diabet Med* [Internet]. 2010 [cited 2020 Sep 20];27(2):217–24. Available from: <https://pubmed.ncbi.nlm.nih.gov/20546267/>
46. Kroenke K, Spitzer RL, Williams JB. The PHQ- 9: Validity of a brief depression severity measure. *J Gen Intern Med* 2001;16:606- 13.
47. Golden SH, Lazo M, Carnethon M, Bertoni AG, Schreiner PJ, Diez Roux A V., et al. Examining a bidirectional association between depressive symptoms and diabetes. *JAMA - J Am Med Assoc* [Internet]. 2008 Jun 18 [cited 2020 Sep 20];299(23):2751–9. Available from: </pmc/articles/PMC2648841/?report=abstract>
48. Knol MJ, Heerdink ER, Egberts ACG, Geerlings MI, Gorter KJ, Numans ME, et al. Depressive symptoms in subjects with diagnosed and undiagnosed type 2 diabetes. *Psychosom Med*. 2007;69(4):300–5.
49. Good CB. Polypharmacy in Elderly Patients With Diabetes. Vol. 15, *Diabetes Spectrum*. 2002.
50. Meneilly GS, Ryan AS, Veldhuis JD, et al. Increased disorderliness of basal insulin release, attenuated insulin secretory burst mass and reduced ultradian rhythmicity of insulin secretion in older individuals. *J Clin Endocrinol Metab* 1997; 82: 4088-4093.
51. Saraf AA, Petersen AW, Simmons SF, Schnelle JF, Bell SP, Kripalani S, et al. Medications associated with geriatric syndromes and their prevalence in older hospitalized adults discharged to skilled nursing facilities. *J Hosp Med* [Internet]. 2016 Oct 1 [cited, 2020 Sep 19];11(10):694–700. Available from: </pmc/articles/PMC5048583/?report=abstract>
52. Good CB. Polypharmacy in Elderly Patients With Diabetes. Vol. 15, *Diabetes Spectrum*. 2002.

53. Reunanen A, Kangas T, Martikainen J, Klaukka T: nationwide survey of comorbidity, use, and costs of all medications in Finnish diabetic individuals. *Diabetes care* 23:1265-1271,2000
54. Noale M, Veronese N, Cavallo Perin P, et al. Polyparmacy in elderly patients with type 2 diabetes receiving oral antidiabetic treatment. *Acta Diabetol* 2016;53:323-30.
55. Anderson RJ, Lustman PJ, Clouse RE, de Groot M, Freedland KE. Prevalence of depression in adults with diabetes: a systematic review [Abstract]. *Diabetes* 2000;49: A64.
56. Archer JS. NAMS/Solvay resident essay award. Relationship between estrogen, serotonin, and depression. *Menopause* 1999;6:71-8.
57. Al-Amer RM, Sobeh MM, Zayed AA, Al-Domi HA. Depression among adults with diabetes in Jordan: Risk factors and relationship to blood sugar control. *J Diabetes Complications* 2011;25:247-52.
58. Nasser J, Habib F, Hasan M, Khalil N. Prevalence of depression among people with diabetes attending diabetes clinics at primary health settings. *Bahrain Med Bull* 2009;31:1-7.
59. Rahman M, Rahman MA, Flora MS, Rakibuz-Zaman M. Depression and associated factors in diabetic patients attending an urban hospital of Bangladesh. *Int J Collaborat Res Intern Med Public Health* 2011;3:65-76.
60. Sotiropoulos A, Papazafiropoulou A, Apostolou O, Kokolaki A, Gikas A, Pappas S. Prevalence of depressive symptoms among noninsulin treated Greek type 2 diabetic subjects. *BMC Res Notes* 2008;1:101.
61. Téllez-Zenteno JF, Cardiel MH. Risk factors associated with depression in patients with type 2 diabetes mellitus. *Arch Med Res* 2002;33:53-60.
62. Roupoulou P, Sotiropoulou P, Makriniakou E, Marneras X, Lahana , et al. Anxiety and depression in patients with type 2 diabetes mellitus, depending on sex and body mass index. *Health Sci J* 2009;3:32-40.
63. Shobhana R, Rama Rao P, Lavanya A, Padma C, Vijay V, Ramachandran A. Quality of life and diabetes integration among subjects with type 2 diabetes. *J Assoc Physicians India* 2003;51:363-5.
64. Lloyd CE, Dyer PH, Barnett AH. Prevalence of symptoms of depression and anxiety in a diabetic clinic population. *Diabet Med* 2000;17:198-202.
65. Raval A, Dhanaraj E, Bhansali A, Grover S, Tiwari P. Prevalence and determinants of depression in type 2 diabetes patients in a tertiary care centre. *Indian J Med Res* 2010;132:195- 200.
66. Ciechanowski PS, Katon WJ, Russo JE. Depression and diabetes: Impact of depressive symptoms on adherence, function, and costs. *Arch Intern Med* 2000;160:3278- 85.
67. Ashfaq, Aaliya Rukhsar Mohammad, Najnin Khanam, Farhan Khan, Rutuj Narendra Waghmare, and Shobha Kanhaiyalal Joshi. "Assessment of Self-Care Practices among Type 2 Diabetes Patients at a Tertiary Care Hospital - A Cross-Sectional Study." *JOURNAL OF EVOLUTION OF MEDICAL AND DENTAL SCIENCES-JEMDS* 9, no. 36 (September 7, 2020): 2630–35. <https://doi.org/10.14260/jemds/2020/572>.
68. Damke, Smita, Dhruva Chandi, and Ramesh Fule. "Study of Bacterial Vaginosis among Women of Reproductive Age Using Contraceptive Methods in a Tertiary Care Hospital." *JOURNAL OF KRISHNA INSTITUTE OF MEDICAL SCIENCES UNIVERSITY* 9, no. 2 (June 2020): 22–27.
69. Damke, Smita Sachin, and Sachin Gajanan Damke. "Seroprevalence of Brucella Agglutinins in Patients with Pyrexia of Unknown Origin Attending a Tertiary Care

- Rural Hospital.” JOURNAL OF EVOLUTION OF MEDICAL AND DENTAL SCIENCES-JEMDS 9, no. 3 (January 20, 2020): 162–65. <https://doi.org/10.14260/jemds/2020/36>.
70. Dubey, Aishwarya, Babaji Ghewade, Keerthan Ganapathi, Diti Gandhasiri, and Dadasaheb Sherekar. “Study of Clinico-Radiological Profile of Mediastinal Masses in a Tertiary Care Centre.” MEDICAL SCIENCE 24, no. 102 (April 2020): 664–72.
  71. Nagdive, Amit, R. U. Zaman, Himanshu Deepak Mansharamani, Prakash B. Behere, and Rouchelle Fernandes. “A Study of Perceived Stress & Coping in Interns in a Tertiary Care Hospital in a North Eastern State of India.” JOURNAL OF EVOLUTION OF MEDICAL AND DENTAL SCIENCES-JEMDS 9, no. 52 (December 28, 2020): 3950–55. <https://doi.org/10.14260/jemds/2020/864>.
  72. Patil, Praful S., Dhruva Hari Chandi, Smita Damke, Shital Mahajan, R. Ashok, and Silpi Basak. “A Retrospective Study of Clinical and Laboratory Profile of Dengue Fever in Tertiary Care Hospital, Wardha, Maharashtra, India.” JOURNAL OF PURE AND APPLIED MICROBIOLOGY 14, no. 3 (September 2020): 1935–39. <https://doi.org/10.22207/JPAM.14.3.32>.
  73. Waghmare, Rutuj, Shobha Joshi, and Pramita Muntode. “Patient’s Rights-Awareness among Indoor Patients of a Tertiary Care Teaching Hospital in Wardha.” JOURNAL OF EVOLUTION OF MEDICAL AND DENTAL SCIENCES-JEMDS 9, no. 8 (February 24, 2020): 570–75. <https://doi.org/10.14260/jemds/2020/127>.
  74. Agrawal, Rajat Kumar, and Shailesh Nagpure. “A Study on Polypharmacy and Drug Interactions among Elderly Hypertensive Patients Admitted in a Tertiary Care Hospital.” INTERNATIONAL JOURNAL OF HEALTH AND ALLIED SCIENCES 7, no. 4 (December 2018): 222–27. [https://doi.org/10.4103/ijhas.IJHAS\\_152\\_17](https://doi.org/10.4103/ijhas.IJHAS_152_17).
  75. Shaheed, Raza, Samarth Shukla, Sourya Acharya, Uplabdh Gopal, and Neema Acharya. “Journey from Fighters to Survivors: Quality of Life and Mental Status in Cancer Patients in a Rural Tertiary Care Hospital.” JOURNAL OF CLINICAL AND DIAGNOSTIC RESEARCH 13, no. 8 (August 2019): XC1–4. <https://doi.org/10.7860/JCDR/2019/41982.13065>.