

ORIGINAL RESEARCH

IDENTIFYING RISK FACTORS FOR CARDIOVASCULAR DISEASE DURING A MEDICAL MISSION OUTREACH

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ABSTRACT

Background

Cardiovascular diseases (CVD) contribute 17.3million to global mortality annually. With increasing urbanization, sedentary lifestyle, obesity and other worsening social indices, incidence of CVD has continued to rise in Nigeria and other third world countries, ultimately worsening already poor economic and clinical outcomes. Medical missions when properly designed and executed can serve as an important avenue for identifying gaps in patient care and promotion of pharmacist-cognitive services.

The objective of this study was to identify CVD risks in volunteers during a medical mission outreach.

Method

The study was an exploratory study. Volunteers were sampled from respondents at a medical mission at Festac town, Amuwo Odofin, Lagos State. A total of 224 volunteers were interviewed

by pharmacists to obtain demographic data and medical history. The weights, height and blood pressure were measured in accordance with JNC7 guidelines. Laboratory tests (Random Blood Sugar or Fasting Blood Sugar and total cholesterol) were conducted by licensed laboratory scientists. Informed consent was taken as decision to participate after the participants were addressed.

Results

The mean weight was $75.4\text{Kg} \pm 12.9$) with 42.9% of the total sample obese and 35.7% overweight. Mean BMI between sexes was different (Male 26.9 ± 4.5 and Female 30.92 ± 4.6) and higher in females (t-test, $p < 0.0001$). Using the waist circumference measure, 54% of patients had a higher risk of CVD. Both BMI and waist circumference were significantly related to age (F-test, $p = 0.001$). More than half of the volunteers (57.1%) had systolic blood pressure (SBP) elevated above 140mmHg and 19.6% having diastolic blood pressure elevated above 90mmHg. SBP was significantly higher in males ($151.4\text{mmHg} \pm 20$) than females ($140.8\text{mmHg} \pm 20.4$) (t-test, $p < 0.0001$).

Conclusion

Medical missions are an important resource in identifying gaps in the healthcare sector and solutions to such problems can be designed at that level. There is currently a huge gap in patient care in Nigeria and pharmaceutical care can be the platform for bridging the gap in care.

KEY WORDS: Medical missions, Hypertension, Pharmaceutical Care, BMI and Waist Circumference.

INTRODUCTION

Cardiovascular diseases (CVDs) are the number one cause of death globally¹. More than three quarters of deaths from CVDs occur in low- and middle- income countries like Nigeria^{2,3}.

Some of the risks for CVD are modifiable others are not modifiable. Non-modifiable risk factors include: age (>55 years), family history of early heart disease or stroke and ethnicity⁴.

Modifiable risk factors include raised blood pressure (BP), body Mass Index (Overweight/Obese category), current smoking, increasing sedentary lifestyle, diet, excessive alcohol consumption, high low density lipoproteins (LDL_C), low high density lipoproteins (HDL_C), high total cholesterol (Chol_t), impaired blood glucose and diabetes^{1,3,4}. The risk of a cardiovascular disease event is dependent on the blood pressure level, presence or absence of sub-clinical target organ damage and other associated cardiovascular risk factors in individual patients⁵⁻⁷. The relationship between cardiovascular disease risk and hypertension is graded, predictable and independent of any other risk factors^{1,8}. Lewinton et al⁸ also showed that hypertension increased the aggregate risk in the presence of other risk factors.

The World Health Organization (WHO) has reported that for countries like Nigeria, low- and middle-income countries, the healthcare system does not offer integrated primary health care programmes that allow prevention, early detection, management of chronic disease and amelioration of risk factors compared to people in high-income countries³. The WHO has identified two types of interventions: population-wide and individual, which are recommended to be used in combination to reduce the greatest cardiovascular disease burden³. At the individual level, for prevention of first heart attacks and strokes, individual health-care interventions need to be targeted to those at high total cardiovascular risk or those with single risk factor levels above traditional thresholds, such as hypertension and hypercholesterolemia. The former

approach is more cost-effective than the latter and has the potential to substantially reduce cardiovascular events^{3, 5}. These interventions are feasible at primary healthcare level in low-resource settings like Nigeria, and can be facilitated by non-physician health workers³.

In many societies in Nigeria, medical missions have been undertaken severally to ameliorate the health system gaps⁹⁻¹². Medical missions when properly designed and executed can serve as an important avenue for promotion of pharmacist-cognitive services.

The objective of this study was to identify CVD risks in volunteers during a medical mission outreach.

METHOD

Study was an exploratory study. Volunteers were sampled from respondents at a medical mission to commemorate the 25th anniversary of Victory Drugs (an independent community pharmacy) located in Festac town, Lagos State, Nigeria.

Sampled respondents were seated in a welcoming area and consent was obtained by decision to participate in the study. Volunteers were then interviewed by adequately trained pharmacists to obtain demographic data and medical history of the volunteers. The weights and heights were measured. The blood pressure was measured three times, at least five minutes apart.

Laboratory tests (Random Blood Sugar - RBS, Fasting Blood Sugar - FBS and total cholesterol Chol_t) were conducted by licensed laboratory scientists.

No identifying data was collected during the study and referrals were made for further evaluation when necessary.

RESULTS

A total of 224 adults participated in the outreach. Half of the volunteers seen were aged 60-69 years (50.4% of sample) and more than half were female (53.1%). The mean weight in the sample was 75.4Kg (\pm 12.9) with 42.9% of the total sample obese and 35.7% overweight. Using the waist circumference measure, 54% of volunteers had a higher risk of CVD.

More than half of the volunteers (57.1%) had systolic blood pressure elevated above 140mmHg and 19.6% having diastolic blood pressure elevated above 90mmHg.

Most of volunteers (86.4%) who had a RBS test had a normal/expected blood sugar level while 35% of volunteers that had a Fasting Blood Sugar test (FBS) had impaired FBS. About 52% of volunteers required intervention in their total cholesterol levels.

Among the volunteers that presented, 52.7% were diagnosed hypertensive, 12.1% had diabetes and 15.2% had been diagnosed with hypercholesterolemia.

Systolic blood pressure was found to be higher in males [151.4 mmHg (\pm 20)] than females [140.8mmHg (\pm 20.4)] volunteers. For those with prior diagnosis of hypertension; more than 80% and 55% of those on antihypertensives had their BMI and waist circumference (respectively) above recommended levels for reduction of CVD risks. About 70% of those not on medication had their blood pressures elevated above pre-hypertensive values. Less than 30% of those on medication had their systolic blood pressure controlled (lower than 140) while 76% of those on medication had achieved recommended diastolic BP goals. Less than 5% had their RBS higher than recommended levels. About half of the volunteers (49.3%) had their total cholesterol above recommended levels.

TABLE 1: MEASURES AND SUMMARY STATISTICS

MEASURES	SUMMARY STATISTICS
AGE	Modal age (50.4%) 60-69 years

SEX	Male 46.1% female 53.1%	
WEIGHT	75.4kg (\pm 12.99)	
HEIGHT	1.612m (sd 0.081)	
BMI*	29.02 (\pm 4.95) Male 26.9 (\pm 4.5), Female 30.92 (\pm 4.6) t-test $p < 0.0001$	Age 50-59 had highest BMI of 30.7 (\pm 5.9)
WAIST**	Mean 94.57 (\pm 10.91cm) Male 93.1cm (\pm 11.8), Female 95.9cm (\pm 9.9) t-test $p > 0.05$	54% have higher risk (greater than recommended levels) of CVD
BLOOD PRESSURE (SYSTOLIC)	Mean 145.8mmHg (\pm 20.9mmHg) Male 151.4 (\pm 20mmHg), Female 140.8 (\pm 20.4mmHg) t-test $p < 0.0001$	57.1% >140 mmHg 20.1% prehypertensive 120-139mmHg 22.8% normal range
BLOOD PRESSURE (DIASTOLIC)	Mean 81mmHg (\pm 10.7mmHg) Male 80.7 (\pm 11.1mmHg) Female 81.3 (\pm 10.2mmHg) t-test $p > 0.05$	48.2% normal range, 19.6% >90 mmHg 32.1% 80-89mmHg
PULSE	Mean 74.5 (\pm 12.4)	
FBS	Mean 98.11g/L (\pm 12.1g/L) Male 96 (\pm 8.4) Female 99.6 (\pm 14.4) t-test $p > 0.05$	
RBS	Mean 99.5g/L (\pm 32.1g/L) Male 97.8 (\pm 27.4), Female 100.9 (\pm 35.7) t-test $p > 0.05$	86.4% normal RBS, 2.5% at significant risk, 3.4% had impaired RBS, 7.6% low RBS
TOTAL CHOL (Chol_t)	Mean 204.63g/L (\pm 48.3g/L) Male 195.6 (\pm 41.5) Female 212.1 (\pm 52.4) t-test $p < 0.05$	48.2% normal, 51.8% requiring intervention, 29.2% borderline high risk, 22.6% high risk

*Decline noted from 50-59 (not less than 1.1 points for each decade, F-test $p = 0.001$)

**Increased up to 60-69years and reduces afterwards (F-test $p = 0.001$)

DISCUSSION

The weight of volunteers was significantly related to age (F-test, $p < 0.0001$). Mean BMI between sexes (Male $26.9 (\pm 4.5)$, Female $30.92 (\pm 4.6)$) was different and significantly higher in females (t-test, $p < 0.0001$). This finding is similar to studies conducted in other states in Nigeria¹³⁻¹⁶ and among urban-dwelling Africans^{17, 18} this is particularly worrying since in another study, it has been shown¹⁴ that obese non-hypertensive Nigerian women may show signs of heart structure modifications that further increase risk of CVDs. Some of these modifications in obese non-hypertensive women include; higher left ventricular mass (LVM), left ventricular mass index (LVMI), wall thickness above normal weight than in non-hypertensive women and women who are obese and hypertensive seem to present more with eccentric hypertrophy significantly higher than in male obese hypertensive subjects¹⁴. This phenomenon is however absent in non-hypertensive obese men. These cardiac remodeling features are an added risk factor in hypertension, heart failure and cardiovascular morbidity and mortality^{5, 19-21} and when the duo of the heart remodeling and obesity occur together, this may predispose to additive risk¹⁴ and may be responsible for noted increases in incidence of other comorbidities²². To reverse this trend, measures must begin to take account of important evidence-based interventions⁵. Important benefits usually result when the interventions made take into account multiple risk factors present²³⁻²⁵.

BMI and waist circumference (WC) were also found to be significantly related to age (F-test, $p = 0.001$). This was correlated to low High Density Cholesterol by Iloh *et al*¹³ and to Triglyceride levels by Fasanmade *et al*²⁶ in similar sample. Okosun *et al* also found a correlation between increasing waist circumference quartiles to blood pressure and fasting blood sugar in a cohort of Nigerian women with gestational diabetes²⁷. Kerenyi²⁸ in their comment to Okosun *et al*²⁷, also

found in this cohort that multiple linear regression showed a statistically significant association between WC, Waist-Hip ratio and BMI to systolic Blood pressure (SBP); but when adjusting for age and SBP, only the BMI had a significant statistical relationship with HBA_{1C}. Pre-pregnancy BMI higher than 25.2kg/m² in females without hypertension or diabetes had an OR of 3.45 (95% CI 1.25 – 9.55) for diabetes and OR of 4.29 (95% CI 1.74 – 10.62) for hypertension ²⁸. Hence interventions to improve BMI may prove to be significantly important in reduction of risk in these cohort and outcomes of treatment may also be improved with interventions targeting patients with BMI above cut-off points indicated in these studies ⁵. Pharmacist intervention has been shown in RCTs to be effective in reducing BMI and other CVD risk factors associated with diabetes ²⁹.

Systolic blood pressure was significantly higher in male than female volunteers (t-test, p< 0.0001). this difference was reversed in studies done in mild hypertensives (excluding obese volunteers and those with compelling indications ^{30, 31} but after adjusting for lean body weight, the observation was as in this study ³¹. Similarly, blood pressure was also found to be significantly different for each age group and this is supported by findings in the WHO STEPs report ¹⁶.

For those diagnosed hypertensive; most of those on antihypertensives had their BMI and waist circumference (respectively) above recommended levels for reduction of CVD risks. This finding of multiple risk factors for CVD is similar to results in Benin ³² and studies in other urban population in Africa ³³. Pharmaceutical care interventions have been shown to improve outcomes through interventions targeting adherence to pharmacological ³⁴⁻³⁷ and some non-pharmacological therapies ^{34, 38-42}.

The observed indices of fewer than 35% receiving treatment for their hypertension are supported by the PURE study⁴³. Less than 30% of those on medication had their systolic blood pressure controlled (lower than 140) while most of those on medication had achieved recommended diastolic BP goals. A significant number however, have their total cholesterol above recommended levels. These are gaps that have been shown to be significantly ameliorated by pharmaceutical care interventions⁴⁴⁻⁴⁶.

CONCLUSION

This study underscores the prevailing under-diagnosis of hypertension and the issues of unmitigated risk for CVDs in the population. Most of the volunteers that participated in the outreach had more than one associated risk factors for CVD and were not currently taking steps to correct their level of risk.

There is currently a yawning gap in patient care in Nigeria and pharmaceutical care can be the platform for bridging the gap in care especially in patients with two or more associated risk factors. Medical Outreach is an important resource in identifying this gap with a view to designing appropriate interventions.

It is, therefore, important that healthcare providers should regard as top priority the need to intensify their outreach programmes and aggregate data therefrom for the much-needed interventions.

REFERENCES

1. Lim SS, Vos T, Flaxman AD, Danaei G, Shibuya K, Adair-Rohani H, et al. (2012). A comparative risk assessment of burden of disease and injury attributable to 67 risk factors and risk factor clusters in 21 regions, 1990-2010: a systematic analysis for the Global Burden of Disease Study 2010. *Lancet*.380(9859):2224-60. Epub 2012/12/19.
2. World Health Organization. (2012). World Health Statistics. 2015 5/03/2015]; Available from: www.who.int/gho/publications/world_health_statistics/2012/en/.
3. World Health Organization. (2014). Global status report on noncommunicable diseases 2014.
4. Thayer JF, Yamamoto SS, Brosschot JF. (2010). The relationship of autonomic imbalance, heart rate variability and cardiovascular disease risk factors. *International journal of cardiology*.141(2):122-131.
5. Chobanian AV, Bakris GL, Black HR, Cushman WC, Green LA, Izzo JL, Jr., et al. (2003). The Seventh Report of the Joint National Committee on Prevention, Detection, Evaluation, and Treatment of High Blood Pressure: the JNC 7 report. *JAMA*.289(19):2560-72. Epub 2003/05/16.
6. James PA, Oparil S, Carter BL, et al. (2014). 2014 evidence-based guideline for the management of high blood pressure in adults: Report from the panel members appointed to the eighth joint national committee (jnc 8). *JAMA*.311(5):507-520.
7. National Clinical Guideline Centre. (2011). Aug. Hypertension: The Clinical Management of Primary Hypertension in Adults: Update of Clinical Guidelines 18 and 34. London: National Clinical Guideline Centre.
8. Prospective Studies C, Lewington S, Whitlock G, Clarke R, Sherliker P, Emberson J, et al. (2007). Blood cholesterol and vascular mortality by age, sex, and blood pressure: a meta-analysis of individual data from 61 prospective studies with 55,000 vascular deaths. *Lancet*.370(9602):1829-39. Epub 2007/12/07.
9. Adebayo AM, Ige OK, Ilesanmi OS, Ogunniyan TB, Ojo T. (2011). Making a case for community screening services: findings from a medical outreach in ibadan, Nigeria. *Ann Ib Postgrad Med*.9(1):14-8. Epub 2011/06/01.
10. Nwankwo KC, Aniebue UU, Aguwa EN, Anarado AN, Agunwah E. (2011). Knowledge attitudes and practices of cervical cancer screening among urban and rural Nigerian women: a call for education and mass screening. *Eur J Cancer Care (Engl)*.20(3):362-7. Epub 2010/03/30.
11. Ramsey K, Iliyasu Z, Idoko L. (2007). Fistula Fortnight: innovative partnership brings mass treatment and public awareness towards ending obstetric fistula. *Int J Gynaecol Obstet*.99 Suppl 1:S130-6. Epub 2007/09/18.
12. Akinremi T, Adeniyi A, Olutunde A, Oduniyi A, Ogo C. (2014). Need for and relevance of prostate cancer screening in Nigeria. *Ecancermedicallscience*.8:457.
13. Iloh G, Amadi AN, Njoku PU, Ofoedu JN, Awa-Madu J. (2012). The magnitude of abdominal adiposity and atherogenic dyslipidemia among geriatric Nigerians with arterial hypertension in a rural hospital in South-eastern Nigeria. *Niger J Clin Pract*.15(4):462-8. Epub 2012/12/15.
14. Akintunde AA, Oladosu Y, Opadijo OG. (2013). Gender specific pattern of left ventricular cardiac adaptation to hypertension and obesity in a tertiary health facility in Nigeria. *Afr Health Sci*.13(3):595-600. Epub 2013/11/20.

15. Akarolo-Anthony SN, Odubore FO, Yilme S, Aragbada O, Odonoye G, Hu F, et al. (2013). Pattern of dietary carbohydrate intake among urbanized adult Nigerians. *Int J Food Sci Nutr.*64(3):292-9.
16. World Health Organization. (2003). World Health Organization STEPS Country Report. Geneva, Switzerland. Geneva, Switzerland 2015 6th May]; Available from: <http://www.who.int/chp/steps/reports/en/>.
17. Okafor CI, Gezawa ID, Sabir AA, Raimi TH, Enang O. (2014). Obesity, overweight, and underweight among urban Nigerians. *Niger J Clin Pract.*17(6):743-9. Epub 2014/11/12.
18. Abubakari AR, Lauder W, Agyemang C, Jones M, Kirk A, Bhopal RS. (2008). Prevalence and time trends in obesity among adult West African populations: a meta-analysis. *Obes Rev.*9(4):297-311. Epub 2008/01/09.
19. Cohn JN, Ferrari R, Sharpe N. (2000). Cardiac remodeling--concepts and clinical implications: a consensus paper from an international forum on cardiac remodeling. Behalf of an International Forum on Cardiac Remodeling. *J Am Coll Cardiol.*35(3):569-82. Epub 2000/03/15.
20. de Simone G, Devereux RB, Roman MJ, Alderman MH, Laragh JH. (1994). Relation of obesity and gender to left ventricular hypertrophy in normotensive and hypertensive adults. *Hypertension.*23(5):600-6. Epub 1994/05/01.
21. Morse SA, Bravo PE, Morse MC, Reisin E. (2005). The heart in obesity-hypertension. *Expert Rev Cardiovasc Ther.*3(4):647-58. Epub 2005/08/04.
22. Iloh G, Amadi AN, Nwankwo BO, Ugwu VC. (2011). Obesity in adult Nigerians: a study of its pattern and common primary co-morbidities in a rural Mission General Hospital in Imo state, South-Eastern Nigeria. *Niger J Clin Pract.*14(2):212-8. Epub 2011/08/24.
23. Buse JB, Ginsberg HN, Bakris GL, Clark NG, Costa F, Eckel R, et al. (2007). Primary prevention of cardiovascular diseases in people with diabetes mellitus: a scientific statement from the American Heart Association and the American Diabetes Association. *Diabetes Care.*30(1):162-72. Epub 2006/12/29.
24. Gaede P, Lund-Andersen H, Parving HH, Pedersen O. (2008). Effect of a multifactorial intervention on mortality in type 2 diabetes. *N Engl J Med.*358(6):580-91. Epub 2008/02/08.
25. American Diabetes A. (2015). (8) Cardiovascular disease and risk management. *Diabetes Care.*38 Suppl:S49-57. Epub 2014/12/30.
26. Fasanmade OA, Odeniyi IA, Amira CO, Okubadejo NU. (2013). Association of body mass index and abdominal adiposity with atherogenic lipid profile in Nigerians with type 2 diabetes and/or hypertension. *Niger Med J.*54(6):402-7. Epub 2014/03/26.
27. Okosun IS, Cooper RS, Rotimi CN, Osotimehin B, Forrester T. (1998). Association of waist circumference with risk of hypertension and type 2 diabetes in Nigerians, Jamaicans, and African-Americans. *Diabetes Care.*21(11):1836-42. Epub 1998/11/05.
28. Kerenyi Z, Stella P, Bosnyak Z, Tabak AG, Tamas G. (1999). Association between central adiposity and multimetabolic syndrome in a special cohort of women with prior gestational diabetes. *Diabetes Care.*22(5):876-7. Epub 1999/05/20.
29. Santschi V, Chiolero A, Paradis G, Colosimo AL, Burnand B. (2012). Pharmacist interventions to improve cardiovascular disease risk factors in diabetes: a systematic review and meta-analysis of randomized controlled trials. *Diabetes Care.*35(12):2706-17. Epub 2012/11/23.

30. Eison H, Phillips RA, Ardeljan M, Krakoff LR. (1990). Differences in ambulatory blood pressure between men and women with mild hypertension. *J Hum Hypertens*.4(4):400-4. Epub 1990/08/01.
31. Gleim GW, Stachenfeld NS, Coplan NL, Nicholas JA. (1991). Gender differences in the systolic blood pressure response to exercise. *Am Heart J*.121(2 Pt 1):524-30. Epub 1991/02/01.
32. Houehanou YC, Lacroix P, Mizehoun GC, Preux PM, Marin B, Houinato DS. (2015). Magnitude of cardiovascular risk factors in rural and urban areas in benin: findings from a nationwide steps survey. *PLoS One*.10(5):e0126441. Epub 2015/05/07.
33. Mathenge W, Foster A, Kuper H. (2010). Urbanization, ethnicity and cardiovascular risk in a population in transition in Nakuru, Kenya: a population-based survey. *BMC Public Health*.10:569. Epub 2010/09/24.
34. Morgado M, Rolo S, Castelo-Branco M. (2011). Pharmacist intervention program to enhance hypertension control: a randomised controlled trial. *Int J Clin Pharm*.33(1):132-40. Epub 2011/03/03.
35. Morgado MP, Morgado SR, Mendes LC, Pereira LJ, Castelo-Branco M. (2011). Pharmacist interventions to enhance blood pressure control and adherence to antihypertensive therapy: Review and meta-analysis. *Am J Health Syst Pharm*.68(3):241-53. Epub 2011/01/25.
36. Fischer MA, Choudhry NK, Bykov K, Brill G, Bopp G, Wurst AM, et al. (2014). Pharmacy-based interventions to reduce primary medication nonadherence to cardiovascular medications. *Med Care*.52(12):1050-4. Epub 2014/10/17.
37. Carter BL, Doucette WR, Franciscus CL, Ardery G, Kluesner KM, Chrischilles EA. (2010). Deterioration of blood pressure control after discontinuation of a physician-pharmacist collaborative intervention. *Pharmacotherapy*.30(3):228-35. Epub 2010/02/26.
38. Tremblay M, Comtois D, O'Loughlin J. (2013). Pharmacists' smoking cessation counseling practices: a comparison between 2005 and 2010. *Nicotine Tob Res*.15(12):2114-9. Epub 2013/08/15.
39. Saba M, Diep J, Saini B, Dhippayom T. (2014). Meta-analysis of the effectiveness of smoking cessation interventions in community pharmacy. *J Clin Pharm Ther*.39(3):240-7. Epub 2014/04/23.
40. Marin Armero A, Calleja Hernandez MA, Perez-Vicente S, Martinez-Martinez F. (2015). Pharmaceutical care in smoking cessation. *Patient Prefer Adherence*.9:209-15.
41. Aguwa CN, Ukwue CV, Ekwunife OI. (2008). Effect of pharmaceutical care programme on blood pressure and quality of life in a Nigerian pharmacy. *Pharm World Sci*.30(1):107-10. Epub 2007/08/19.
42. Lenz TL, Monaghan MS. (2008). Lifestyle modifications for patients with hypertension. *J Am Pharm Assoc* (2003).48(4):e92-9; quiz e100-2. Epub 2008/07/26.
43. Chow CK, Teo KK, Rangarajan S, Islam S, Gupta R, Avezum A, et al. (2013). Prevalence, awareness, treatment, and control of hypertension in rural and urban communities in high-, middle-, and low-income countries. *JAMA*.310(9):959-68. Epub 2013/09/05.
44. Pande S, Hiller JE, Nkansah N, Bero L. (2013). The effect of pharmacist-provided non-dispensing services on patient outcomes, health service utilisation and costs in low- and middle-income countries. *Cochrane Database Syst Rev*.2:CD010398. Epub 2013/03/02.

45. Wal P, Wal A, Bhandari A, Pandey U, Rai AK. (2013). Pharmacist involvement in the patient care improves outcome in hypertension patients. *J Res Pharm Pract.*2(3):123-9. Epub 2014/07/06.

46. Amariles P, Sabater-Hernandez D, Garcia-Jimenez E, Rodriguez-Chamorro MA, Prats-Mas R, Marin-Magan F, et al. (2012). Effectiveness of Dader Method for pharmaceutical care on control of blood pressure and total cholesterol in outpatients with cardiovascular disease or cardiovascular risk: EMDADER-CV randomized controlled trial. *J Manag Care Pharm.*18(4):311-23. Epub 2012/05/03.