

1 Correlation between Serum uric acid and Blood Pressure in Plateau Central

5 Abstract

6 **Aim:** To determine the Correlation between Serum uric acid (SUA) and Blood
7 Pressure in Plateau Central

8 Background

9 Serum uric acid, an end product of purine metabolism, has been shown to be
10 associated with an increased risk of hypertension cardiovascular⁴ and chronic kidney
11 disease in previous epidemiological studies. Elevated SUA has been shown to
12 predict the development of high blood pressure and may contribute to increasing BP
13 by several mechanisms. Clinical trials in young patients have supported this
14 mechanism but do not yet agree with pharmacologic reduction of SUA as first-line
15 therapy for hypertension.

17 Materials and Method

18 200 (males and females) subject of Panyam district of Mangu L. G. C. in Plateau
19 central formed the study population with aged 40 - 90 years. The intervention groups
20 were 120 and control group was 80. Five ml of blood was aliquoted for assay uric
21 acid for each subject using the enzymatic uricase method.

23 Results

24 The data obtained were coded into Stata Software for analysis. The data was
25 presented as mean ± S.D. comparison was done by student's t-test for continuous
26 variables. A BMI of >30kg/m² was found in 31 individuals giving a prevalence 15.5%
27 with SUA prevalence of 41% with 32% being the prevalence of hypertension.

29 Conclusion

30 The rural farmers were found to have high SUA and positively correlated to
31 hypertension.

32 With the high level of SUA and high BP, advocacy should be made on regular BP
33 checks and possible provision of personal portable BP machine for self-monitoring.

35 Introduction

36 One of the worldwide public health challenges is hypertension and remains a leading
37 cause of morbidity and mortality. It is the most common form of cardiovascular
38 disease and its prevalence growing higher with age and other risk factors like
39 obesity, physical inactivity, diabetes mellitus and race^{1, 2}. The end product of purine
40 metabolism is uric acid which has been shown to be associated with an increased
41 risk of hypertension³, cardiovascular, and chronic kidney disease in previous
42 epidemiological studies. Also, elevated levels of uric acid may progress to peripheral
43 arterial disease, insulin resistance and components of the metabolic syndrome⁴.
44 Hyperuricemia defined as 7mg/dL in males and 6mg/dL in females. It is a common

45 metabolic disorder occurring in 2.18% population varying in age, sex and other
46 factors like excessive alcohol⁵. SUA concentration could determine the outcome of
47 hypertension and may also initiate hypertension, though it is less clear that
48 hyperuricaemia can be regarded as independent risk factor given its association with
49 well-recognised factors. At physiologic concentration, uric acid is antioxidant and
50 beneficial, but at a high level, it becomes a free radical leading to pathological
51 processes⁶. The result of pathophysiological processes of SUA increases such as
52 impaired renal sodium handling but may also contribute to renal vascular damage
53 particularly endothelial dysfunction. This may cause subtle endothelial damage that
54 may stimulate the renni-angiotensin-aldosterone system (RAAS) causing
55 hypertension.⁷ More than 20-40% of patients with untreated hypertension and 80%
56 or more patients with malignant hypertension have high SUA levels. Hyperuricaemia
57 common in primary HTN especially in patients with HTN of recent onset is
58 associated with micro albuminuria⁸
59 The Framingham heart study showed that each increase in SUA by 1.3mg/dl was
60 associated with the development of HTN with an odd ration of 1.17⁹. The reduction
61 in SUA to less than 5mg/dL with allupurinol was associated to the reversal of HTN in
62 86% of the patients¹⁰. A 12 years Italian study – (The PIUMA study) involving 1720
63 previously untreated hypertensive patients, Verdechia and colleagues also found
64 that serum acid was a powerful predictor of cardiovascular disease and all-cause
65 mortality¹¹. Because of eminent cardiovascular disease morbidity, it has become
66 pertinent to determine the prevalence of hyperuricaemia and hypertension and the
67 association between SUA and the blood pressure¹²⁻¹⁴.

68

69 **Materials and methods**

70 **Research setting and Design**

71 This is a prospective study covering all neighbouring villages. 200 famers and retired
72 civil servants of Panyam district of Mangu L.G.C in Plateau central formed the study
73 population aged 40 - 90 years. Both males and females were recruited into the
74 study. The intervention groups were 120 and control group was 80.

75 **Sample and Sampling Method**

76 This study was carried out at Panyam Primary Health Care.

77 Permission was obtained from the district head. He mobilised his subjects to have an
78 overnight fast for the study. Blood samples was collected from the subjects into plain
79 specimen tubes between 8:00 to 10:00am. This blood was spun at 3000 rpm for 5
80 minutes using gallenkamp bench-centrifugal, after clot retraction. The supernatant
81 (serum) extracted using Pasteur pipettes into storage tubes and stored frozen at -
82 20°C before analysis at Jos University Teaching Hospital Chemical Pathology
83 Laboratory.

84 **Data Collection and Analysis**

85 Weight was measured using a portable weighing scale (standiometer) while a meter
86 rule used in the measurement of height. Both were taken in a standing position with
87 shoes, and heavy clothing removed. The body mass index (BMI) was calculated as
88 follows:

89 $BMI = \text{Weight (kg)}/\text{height (m}^2\text{)}$.

90 BMI was categorised using the WHO definitions.

91 The aneroid sphygmomanometer was used in the measurement of blood pressure.
92 Blood pressure was measured in the right arm after at least 15 min of rest and while
93 participants were sitting down. The cuff was applied evenly and snugly around the
94 bare arm, with the lower edge 2.5 cm above the antecubital fossa. The participants
95 must not have eaten, smoked tobacco or taken alcoholic beverages for at least 30
96 minutes before the measurements. The first and fifth Korotkoff sounds were taken as
97 the systolic blood pressure (SBP) and diastolic blood pressures (DBP) respectively.
98 Hypertension was noted if systolic blood pressure 130mmHg, or upon self-report of a
99 medical diagnosis of hypertension or current treatment for hypertension with
100 prescription medication.

101 Five ml of blood was aliquoted for the assay. Blood glucose and uric acid were
102 determined by glucose oxidase and uricase enzymatic method respectively. The
103 reference interval for glucose is 3.5-5.9 and for uric acid 120µmmol/L - 420µmmol/L.

104 **Ethics**

105 Individual consent was obtained from all recruited individuals. Ethical clearance was
106 obtained from the research and ethical committee of Jos University Teaching
107 Hospital.

108

109 **Statistical Analysis**

110 The stata software was employed for the analysis. Pearson chi-square **were** used for
111 nominal and the independent samples –test for continuous variables. A value below
112 0.05 was considered significant.

113

114

Results

115 **Characteristics of study participants according to their quartiles**

116 The analysis was conducted on 200 individuals who had both BMI, GLU, Uric
117 acid and blood pressure assessment. The mean age of the individuals was
118 63.63 years and approximately 43% were men. SUA prevalence of 41% and
119 32% prevalence of hypertension was noted.

120 Table 1: **Characteristics of study participants according to their quartiles**

Characteristics	1 ST	2 ND	3 RD	4 TH	P- VALUE
	QUARTILE	QUARTILE	QUARTILE	QUARTILE	
	≤0.2	0.2-0.24	0.25-0.29	≥0.30	
AGE	47.5±7.5	59±3	85±5	110±10	<0.0001
BMI	19.58±2.64	23.40±1.05	26.52±2.06	33.73±4.84	<0.0001
GLU(F/R)	2.04 – 3.85	3.93 – 4.76	4.81 – 6.5	6.6 – 11.6	<0.0001
URIC ACID	59-246	265-436	450-750	793-1963	<0.0001
SBP	110±10		135±5	160±20	<0.0001
DBP	75±5		85±5	110±10	<0.0001

121

122

Associations

123

Partial Spearman correlation analysis demonstrated the strongest association between Uric acid and body mass index (BMI). It also signifies a low correlation between uric acid and blood pressure. SBP and DBP had a very strong Correlation from the data analyzed.

124

125

126

127

Table 2: **Partial Spearman correlation coefficients among Uric acid, blood pressure, Body mass index and**

128

	URIC ACID	BMI	SBP	DBP
BMI	0.1453			
SBP	-0.0394	-0.0740		
DBP	-0.0758	-0.0767	0.7561	
GLU(F/R)	0.1265	0.0108	0.0972	0.0250

129

130

131

132

133

134

135

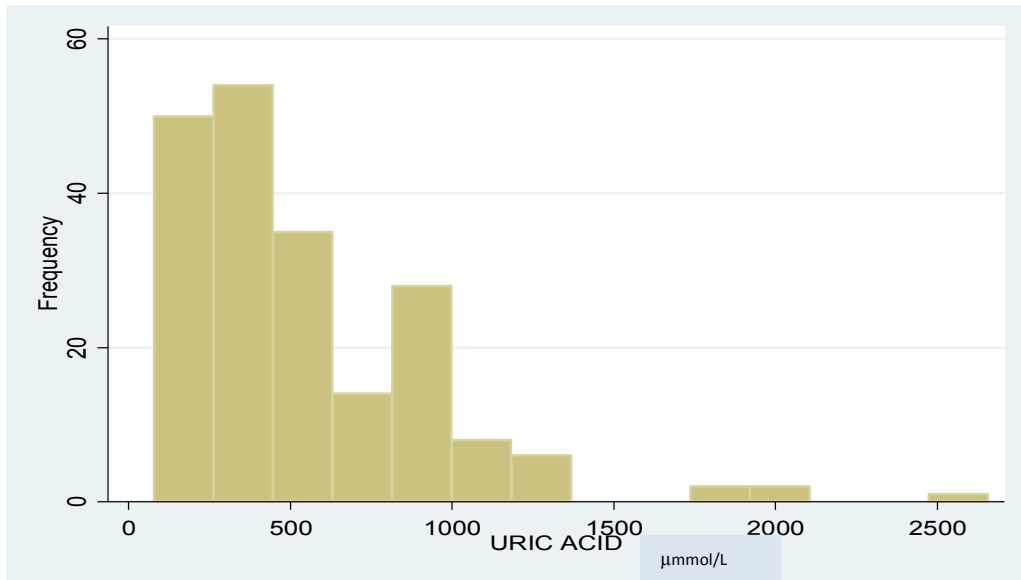
136

137

138

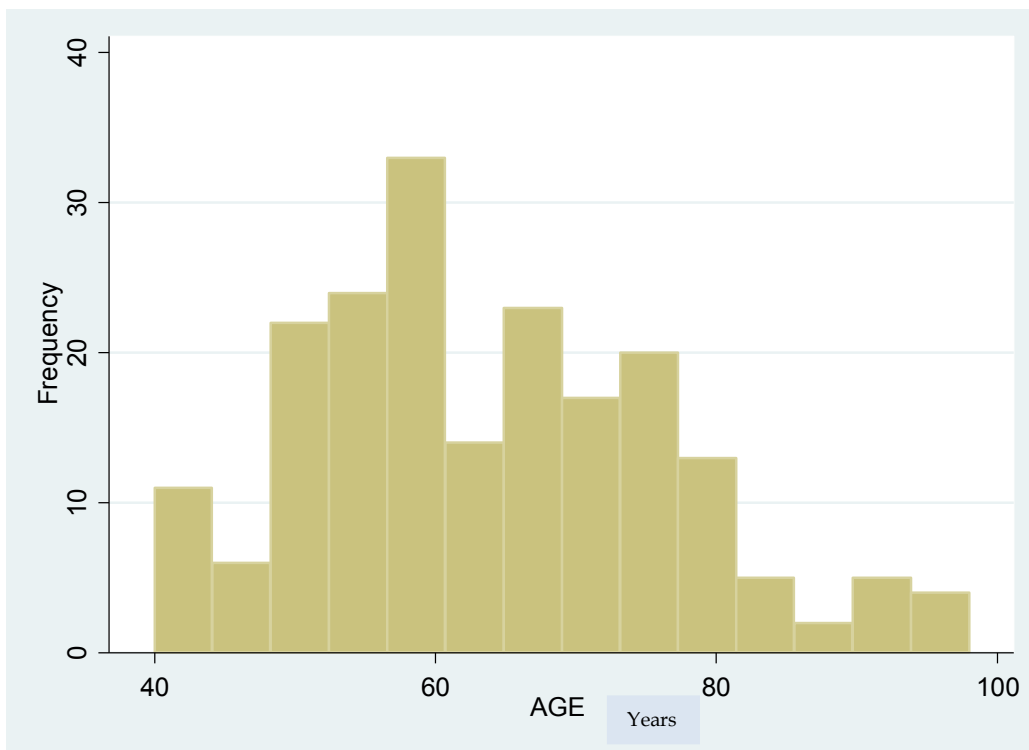
139

140 Fig1: uric acid levels and percentage population
141



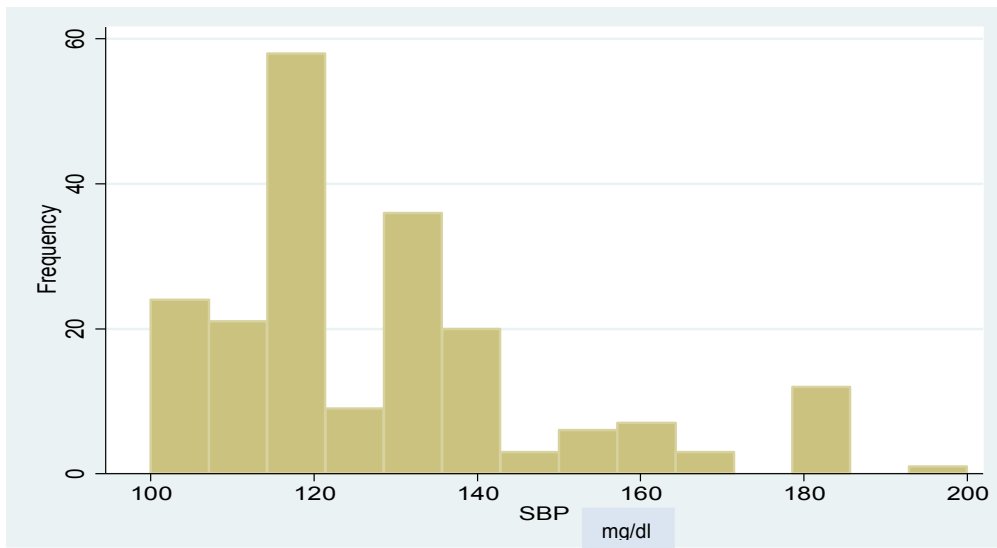
142
143

144 Fig 2 : age and percentage population



145
146
147

148 Fig 3 : SBP and percentage population



149

150

151 Discussion

152 This study was set-put to examine the prevalence of hypertension and
153 hyperuricaemia among middle age and elderly rural farmers. The second aim was to
154 determine any association between SUA levels and hypertension.

155 A high-risk group for the development of hypertension is the Africa population and
156 identifying risk factors is therefore important for preventive actions against
157 cardiovascular diseases¹⁵.

158 Based on the seventh report of the Joint National Committee on Prevention,
159 Detection, Evaluation and Treatment of high blood pressure¹⁶, normal BP is defined
160 as less than 120/80mmHg. The SBP was obviously higher in elderly intensive than
161 the control. It is known that BP increases with age so it is consistent with previous
162 work done¹⁶ in HTN association with age; this only further proves that point. From
163 this study, it was found that a mean SUA of 569 μ mol/L compared with the control
164 484 μ mol/L from studies¹⁷, it showed that the development of HTN has generally
165 been consistent. The SUA was found to be significant, higher in the intensives than
166 the control. Previous work revealed SUA to be higher and cause HTN in the young.
167 We discovered in this elderly population that SUA was significantly higher than the
168 control for the reason of advanced age and the elevated BP. This work was in
169 consistency with the Framingham Heart Study¹⁸⁻¹⁹ that reported SUA was not casual
170 risk factors for CVD events because uric acid was not independent of HTN. Logical
171 data on the independent prognostic role of SUA might be accounted for the complex
172 interrelations between SUA and a variety or risk markers for cardiovascular diseases
173 including male gender epidemiological cardiovascular events.

174 This work showed a consistency with systolic HTN in the elderly program and
175 Chicago studies in several individuals²⁰.

176 In our study, the relationship of SUA to CV HTN is and also apparent from inspection
177 of studies by Alderman *et al*²¹ in subjects with hypertension. Longitudinal studies are
178 needed to clarify the potential value of SUA to reflect and predict the vicious cycle
179 leading to progressive renal damage and elevated blood pressure. Increased activity
180 of the sympathetic nervous system has also been associated with reduced renal
181 excretion of uric acid but the basic mechanisms are unknown²².

182 The present study demonstrates a strong independent association between SUA
183 and HTN, initially untreated and asymptomatic adult subjects with essential
184 hypertension, but it is unable to answer the question of whether SUA exerts direct
185 toxic effects.

186 Under most conditions, an elevated SUA is in equilibrium with intracellular level²³.
187 However, one confounding aspect is that SUA levels are known to fall in diabetic
188 subjects, as glycosuria can lead to proximal tubular dysfunction and uricosuria. SUA
189 levels have been also reported to be higher in those with better diabetes control.

190 A possible explanation for high-level SUA in the population is the high indulgence in
191 local alcohol beverage which is known to cause hyperuricaemia. Another reason for
192 high hyperuricemia and hypertension is African population is known that blood
193 pressure was more pronounced in the African men²⁴.

194 The adoption of more western seed diet, because industrialisation had a great effect
195 on the latter mentioned. Due to the high content of sugar (fructose) in western seed
196 diet, the prevalence of obesity and diabetes increased resulting in concomitant
197 increases in SUA levels. A confounding factor such as BMI could explain this
198 association²⁵.

199 The finding this work relating SUA to hypertension incidence confirmed several
200 previous reports. The strength of the association was modest in our study compared
201 to other reports, for example, in the Olivet Study a high increment in SUA was
202 associated an OR of 1.23 for hypertension incidence during 12 years follow up²⁶.

203

204 **Conclusion**

205 It is concluded that rural farmers were found to have high SUA and significantly
206 correlated with hypertension. With the high level of SUA and high BP, advocacy
207 should be made on regular BP checks and possible provision of personal portable
208 BP machine for self-monitoring. There is a need for further health awareness on the
209 control in the consumption of local alcoholic beverage and other lifestyle modification
210 such as regular exercise and personal hygiene.

211

212

213 **Ethics & Consent:**

214 Individual consent was obtained from all recruited individuals. Ethical clearance was
215 obtained from the research and ethical committee of Jos University Teaching
216 Hospital.

217

218

219 Abbreviation :

220 **SUA = Serum uric acid**

221 **RAAS = Renni-angiotensin-aldosteron system**

222 **HTN = Hypertension**

223 **SBP = Systolic blood pressure**

224 **DBP = Diastolic blood pressures**

225 **BMI = Body mass index**

226 **CVD = Cardio-vascular diseases**

227

228

229

230

231

232 **References;**

- 233 1. Kaminer B and Lutz WP. Blood pressure in Bushmen of the Kalahani Desert,
234 Circ Res. 1960;12:289 – 295.
- 235 2. Shaper AG, Wright DH and Kyobe J. Blood pressure and body build in three
236 nomadic tribes of Northern Kenya E. Afr. Med J. 1969, 46:273 – 281.
- 237 3. Johnson T. O Prevalence of overweight and obesity among adult subjects of
238 an urban African population sample. Br. J. Prev. Soc Med. 1970, 24:105 –
239 **109**
- 240 4. Johnson T. O. America blood pressure and hypertension in an urban America
241 population sample Br. J. Prev. Soc Med. 1971 25:26, 33
- 242 5. F Tesfaye, H van minh, P Byass, Y Berhane, R Bonita, NG Nawi et al.
243 Association between BMI and blood pressure across three populations in
244 Africa and asia. Am J heart assoc. 2017; (3): 501-504.
- 245 6. J. stevens, J.Cai, E. R. Pamuk, D. F. Williamson, M. J. Thun, and J. L. Wood,
246 “the effect of age on the association between body mass index and mortality”,
247 N Engl J Med 1998; 338(1): 1-7.
- 248 7. P Bovet, A. G. Ross, J. P. Gervasoni et al., “Distribution of blood pressure,
249 body mass index, and smoking habits in the urban population of Dar es
250 Salaam, Tanzania, and associations with socioeconomic status”, Intl J Epid.
251 2002; 31(1): 240-247.
- 252 8. R. B. Singh, S. S. Rastogi, V. Rastogi et al, “Blood pressure trends, plasma
253 insulin levels and risk factors in rural and urban elderly populations of north
254 India,” Cor. Art. Dis, 1997; 8(7): 463-468.
- 255 9. S. K. Kumanyika, J. R. Landis, Y. L. Matthews, S. L. Weaver, L. C. Harlan, and
256 W. R. Harlan, “Secular trends in blood pressure among adult blacks and
257 whites aged 18-34 years in two body mass index strata, United States, 1960-
258 1980,” Am J of Epid. 1994;139(2) :141-154.
- 259 10. A. G. Shaper and P. H. Whincup, “Annotation: hypertension in populations of
260 African origin,” Am J of Pub Health 1997; 87(2) :155-156.
- 261 11. Michael L Ganz, Neil Wintfeld, Qian Li, Veronica Alas, Jakob Langer and
262 Mette Hammer. The association of body mass index with the risk of type 2
263 diabetes: a case-control study nested in an electronic health records system
264 in the United states. Diabetology and Metabolic Syndrome 2014, 6:50(10) 17-
265 58.
- 266 12. Johan, S., Lisa S., Ralph, B. et al Relations of Serum Uric Acid to Longitudinal
267 Blood Pressure Tracking and Hypertension. Hypertension 2005; 45:28-33.
- 268 13. Jossa, F., Fainaro, E., Panico, S., Krogh, V., Clenetano, E., Galasso, R.,
269 Mancini, M., Trevisan, M. Serum Uric Acid and hypertension: the Olivetti
270 Heart study. J Hum Hypertens. 1994; 8:667-618.
- 271 14. Selby, J.V., Friedman, G.D., Quesenberry, C. P., Jr. Precursors of essential
272 hypertension: pulmonary function, heart rate, uric acid, serum cholesterol, and
273 other serum chemistries. Am J Epidemiol. 1990; 131:1017-1027.

- 274 15. Kahn, H. A., Medalie, J H., Neufeld H.N., Riss, E. Goldbourt U. The incidence
275 of hypertension and associated factors: the Israel ischemic heart disease
276 study. *Am heart J.* 1972; 84:171-182
- 277 16. Nakanishi, N., Okamoto, M., Yoshida, H., Matsuo, Y., Suzuki, K, Tatara, K.
278 serum uric acid and risk for development of hypertension and impaired fasting
279 glucose or type II diabetes in Japanese male office workers. *Eur. J Epidemiol.*
280 2003; 18:583-530
- 281 17. Hunt, S.C., Stephenson, S, H., Hopkins, P. N., Willimas, R. R. Predictors of
282 an increased risk of future hypertension in Utah. A screening analysis.
283 *Hypertension* 1991; 17:969-976.
- 284 18. Dawber T. R., Meadors, G. F., Moore, F. E. J. Epidemiological approaches to
285 heart disease: the Framingham study. *Am J public health* 1951; 41:279-286.
- 286 19. Clarke, R., Shipley, M., Lewington, S., Younman, L., Collins, R., Mar,ot, M.,
287 Peto, R. Underestimation of risk association due to regression dilution in
288 longterm follow-up of prospective studies. *Am J Epidemiol.* 1999; 150:341-
289 353.
- 290 20. Y Tamiguchi , T. Hayashi, K. Tsumura, G. Endo, S. Fuji, and K. Okada,
291 "Serum uric acid and the risk for hypertension and type 2 diabetes in
292 Japanese men: The Osaka health survey," *Journal of Hypertension*, 2001; 19
293 (7): 1209-1215.
- 294 21. Oviasu VO, O Okupa FE Arteria blood pressure and hypertension in Benin in
295 the equatorial forest zone of Nigeria. *Trop Geogr. Med.* 1980,32.241 – 44
- 296 22. JE Renail and cardiovascularText Google Scholar mechanism of hypertension
297 in obesity *Hypertension* 1994 – 23:381, Abstract Full
- 298 23. Gamison R. J. Healthy adiposity in women the Framingham offspring study:
299 *J. Am Coll Nutr* 1993;12:357.362.
- 300 24. A. Shankar, R. Klein, B.e.K. Klein and F.J. Nieto, "The association between
301 serum uric acid level and long-term incidence of hypertension 2006; 20 (12):,
302 937-945.