1	Association between ABO Blood Group Phenotype and Reaction to Academic stress
2	and coping strategies among college students
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5	Abstract:
6	Introduction: Proper stress response and use of ideal coping strategy can reduce
7	stress-related death. However, whether individual's stress response/coping method
8	is related to the blood type has not been empirically examined.
9	Objectives: Given this, this study examined the relationship between ABO blood
10	type and reaction to academic stress and coping strategies among college students
11	in Southern Nigeria.
12	Methods: A cross-sectional study was conducted on 319 (142 male and 77 female)
13	students. They were examined using Perceived stress scale-10 (PSS-10) which
14	measures stress level. Reaction to stress and coping strategy questionnaires (CSQ)
15	were used to measure four types of stress reactions and coping methods.
16	Results: Prevalence of perceived stress was 79.6%. Most of the stressed
17	participants (46.5%) were of blood group O. Most prevalent reactions to stress
18	versus (vs) coping methods according to blood type were emotional (99.3%) vs
19	religious (95.0%), emotional (88.4%) vs active distracting (90.9%), behavioral
20	(82.5%) vs active practical (95.0%), and physiological (91.5%) vs active distracting
21	(98.3%), for blood groups A, B, AB and O respectively. Compared to those with
22	blood group A, participants with blood groups B and AB had higher odds of
23	emotional reaction to stress (Odds Ratios (ORs); 2.13, (95% Confidence intervals
24	(CI):1,459-2.799) and 1.47(95% CI: 0.555-3.871) respectively. ORs and CIs for
25	cognitive reaction were 5.68(1.866-36.486) and 1.71(0.429-6.773) for blood groups ${\rm B}$
26	and AB respectively. For behavioral reaction, ORs and CIs were 1.25(0.494-3.174),
27	2.08(1.684-6.309) and 1.54(0.575-4.09) for blood groups B, AB and O respectively,
28	while ORs and CIs for physiological reaction was 2.75(1.302-10.877) for blood group

Comment [U1]: Verify the sum

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O. Active practical, active distracting, religious and active practical coping methods
were mostly used by individuals with blood groups A, AB, B and O respectively.

Conclusion: There is a relationship between individuals' ABO blood type and the risk of being stressed, type of stress response and coping methods employed to combat stress. Interventions to reduce stress and associated disorders should among other factors consider individuals' blood group.

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36 Introduction:

Stress is universal and inevitable part of life created by excessive environmental 37 and internal demands. It needs to be adequately responded to and coped with to 38 avoid the associated adverse health effects which are enormous and debilitating. 39 Although every one experiences stress as a result of normal daily activities, prior 40 studies have shown a unique stress response among individual to combat the 41 perceived stress-imposed threat. Each person has a different reaction and coping 42 strategy to his or her stress. Stress response is part of the homeostatic balance and 43 is necessary for evolution and survival. Dysfunction of stress response system leads 44 to poor physical, metabolic and physiological endpoints and consequently, to several 45 diseases including diabetes mellitus [1], peptic ulcer [2], obesity [3], sleep disorders, 46 hypertension [4], ischaemic heart disease [5], osteoporosis [6], reduced mental 47 48 alertness, increased anxiety, risk of depression, poor academic performance and several cancers. Interestingly, people and groups differ in sensitivity and 49 vulnerability to these stress-induced disorders probably due to differences in their 50 interpretation, reaction and coping strategies adopted to combat stressors [7]. 51

The central nervous system may response to stressful situation by altering the endocrine, neural and/or immune activities. However, the pattern of biological response depends on the situation. Even so, there are individual differences in stress response to similar situation [8]. This inherent stress adaptive ability has been shown to have some genetic predisposition [9] and thus, largely depends on our body's natural biochemical response which is determined by our

deoxyribonucleic acid (DNA). The composition of our ABO blood type is determined 58 by the inheritance of gene on chromosome 9(9q34) [10]. Interestingly, the same 59 DNA locus (9g34) that determines our blood type also controls our response to 60 stress. Previous studies had attempted to establish a link between individuals' 61 62 blood type and personality trait with no scientific consensus, however recent evidence implicates the role of a catecholamine metabolizing enzyme (dopamine 63 beta hydroxylase (DBH)) which is known to be involved in the metabolism of 64 dopamine into norepinephrine. It is now known that ABO/DBH linkage exists that 65 determine the dopamine: norepinephrine ratio that influences individual's behavior 66 67 [11, 9].

It is posited that our blood type is important with regards to the type and concentration of these stress hormones, their resting levels, the way we respond to stress, how quickly these level revert back after stress. In the university environment, students are continuously exposed to high level of stress which may be academic or self imposed kind of stressors or both. These may include psychosocial, physical or individual stressors, and they must respond to it on a daily basis [7], though differently [12].

Whether the response pattern has any relationship with individual's blood group 75 phenotype has not been investigated. Understanding the ABO group phenotypic 76 77 correlate of stress, stress response and coping strategy may suggest genetic related risk factors that increase individual's susceptibility to stress and predicts ideal 78 stress response and coping methods to reduce the risk of stress-related diseases. 79 Therefore, the aim of this study was to assess the relationship between ABO blood 80 group phenotype and reaction to academic stress and coping strategies among 81 82 undergraduates in a Nigerian University.

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87 Subjects and Methods

This cross-sectional study was conducted between the month of March and October, 88 2018 in a Nigerian University. Students were invited to participate in the survey 89 through several media including announcement during lecture and practical 90 91 classes. Participation was free and voluntary. Of the initial 528 students who responded to the invitation, 319 (142 male and 77 female) undergraduates from 92 different faculties in the university met the inclusion criteria and were assessed. 93 This number represented 60.42% of the initial number invited to participate. Others 94 (39.58%) were excluded for not meeting the inclusion criteria. 95

96 The exclusion criteria were inappropriate age (<18years), missing data, decline
97 participation, inadequate completion of questionnaire, and poor mental status.

All participants signed written informed consent and the study protocol wasapproved by the Institutional Research Ethics Committee.

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101 Instruments of survey/Assessment measures

Two survey instruments were used in this study, including a 4-section semi structured questionnaire which assessed the demographic characteristics of participants, stress level, reaction to stress and coping strategies employed by the participants and determination of ABO blood group phenotype. The questionnaire was adapted from previous studies with similar hypothesis [13, 14].

Part A of the questionnaire recorded the socio-demographic characteristics of
participants including age (years), gender, marital status, year of study and faculty
of study.

Part B of the questionnaire consisted of items adapted from Perceived Stress Scale-10 (PSS-10) and Student-life Stress Inventory (SLSI), and measures perceived stress, reaction to stress and coping methods used by respondents as described elsewhere [13, 14]. Briefly, participants were asked to respond to each question on a five point scale ranging from 0 (never), to 4(very often). Scores ranged from 0-40 with higher scores indicative of higher stress. A score of ≥ 20 was suggestive of high stress level.

Reaction sub-scale consisted of 28 items and measures four categories of reactions 117 to stressors (physiological, 14 items, emotional, 4 items, behavioral, 8 items and 118 119 cognitive, 2 items). The reaction sub-scale was assessed on a 4-point scale (i.e., 1=never, 2=occasionally, 3=often, 4=most of the times) and dichotomized (1and2 vs 120 3 and 4). It was summed to produce total scores ranging from 1-14 for physiological 121 reaction, 1-4, 1-8 and 1-2 for emotional, behavioral, and cognitive reactions 122 respectively. Low versus (vs) high reactions were 1-7 vs 8-14, 1-5 vs 6-8, 1 vs 2 for 123 124 physiological, behavioral, and cognitive reactions respectively.

The coping strategy sub-section consisted of 63 items and measures 4-types of coping including active practical, active distracting, avoidance and religious coping methods. This section was assessed on a 5-point scale (i.e., 1=never, 2=seldom, 3=occasionally, 4=often, and 5=most of the time). None, low and high usages of a specific coping method were represented by scores of 1, 2-3 and ≥ 4 respectively.

Blood samples were obtained from all participants through a finger prick with a sterile disposable lancet. Anti-A, anti-B and anti-D monoclonal blood group reagents were used to determine participants' ABO blood group phenotype by slide agglutination method.

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135 **Results**:

The socio-demographic variables of 319 undergraduates who participated in this survey revealed that 254 (79.6%) participants were stressed, while 65 (20.4%) were not stressed. In addition, 177 (55.5%) were female while 142 (44.5%) were males. Other demographic variables (age, level of study and faculty of study) showed no significant difference between the stressed and not stressed participants with the exception of marital status which significantly (P<0.05) differed between the two groups (Table 1). 143 Table 2 shows the distribution of stress by blood group as follows; 29.1%, 8.7%,

144 15.7% and 46.5% for blood groups A, B, AB and O respectively. The highest
prevalence of stress was found among individuals with blood group O, followed by

146 A, AB and B

147 Table 3 shows that most participants with blood group A showed emotional reaction (97.37%), followed by behavioral (68.9%) and physiological (51.4%) reactions to 148 academic stressors. Majority of those with blood group B (86.4%) displayed 149 emotional reaction followed by behavioral (59.1%) and physiological (54.5%) 150 reactions to stress. A greater number of participants with blood group AB showed 151 behavioral (82.5%) reaction followed by cognitive (75%) and physiological reactions 152 to stress. Blood group O individuals mostly reacted physiologically (91.5%) followed 153 by behavioral (64.4%) and cognitive reaction (47.5%). 154

Furthermore, majority (93.2%) of participants in group A blood adopted religious 155 mode of coping with stress, blood group B (90.9%) made use of distracting coping 156 strategy while blood group AB (95.0%) employed practical coping strategy. 157 Similarly, most (98.3%) participants with blood group O blood group adopted 158 distracting coping method. Interestingly, it was further observed that although 159 there was a slight variation in the use of practical, distracting, religious coping 160 methods within each of the blood group phenotype, there were no significant 161 162 differences in the number of respondents who employed these coping strategies to 163 cope with academic stressors.

Table 5 shows that the odds of being stressed when faced with stressful situation were higher among individuals with blood group B. Individuals with blood group B have the highest likelihood of reacting cognitively (OR:5.86, CI:1.866-36.486, P=0.006), followed by emotional reaction (OR:2.13, CI: 1.459-2.799, P=0.007) and behavioral reaction (OR:1.25, CI:0.494-3.174, P=0.635) and less likelihood of reacting physiological (OR:0.88, CI:0.338-2.286, P=0.792).

Individuals in blood group AB had higher likelihood of exhibiting behavioral
reaction (OR:2.08, CI:0.266-2.141, P=0.754), cognitive (OR:1.71, CI:0.429-6.773,

172	P=0.448) and emotional (OR:1.47, CI:0.555-3.873, P=0.439) and less likelihood of
173	reacting physiologically (OR:0.754, CI:0.266-2.141, P=0.754). Blood group O
174	individuals have increased likelihood of exhibiting physiological reaction (OR:2.75,
175	CI:1.302-10.877, P=0.042) followed by behavioral reaction (OR:1.54, CI:0.575-4.09,
176	P=392) and less likelihood of exhibiting emotional and cognitive reactions (Table 6).
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Table 1: Socio-demographic Characteristics of the Respondents

	With Stress	Without	Total	
Variables	(n=254)	Stress	(n=319)	p-value
	(11-254)	(n=65)		
Sex				
Male	108(42.5)	34(52.3)	142(44.5)	0.202
Female	146(57.5)	31(47.7)	177(55.5)	
Age (Years)				
16-19	39(15.4)	12(18.5)	51(16.0)	
20-25	170(66.9)	47(72.3)	217(68.0)	0.385
26-30	33(13.0)	5(7.7)	38(11.9)	
31 and above	12(4.7)	1(1.5)	13(4.1)	
Marital Status				
Single	211(83.1)	64(98.5)	275(86.2)	
Married	35(13.8)	1(1.5)	36(11.3)	0.016*
Divorced/Separated	4(1.6)	0(0.0)	4(1.3)	
Widowed	4(1.6)	0(0.0)	4(1.3)	
Level of Study				
100L	41(16.1)	9(13.8)	50(15.7)	
200L	53(20.9)	17(26.2)	70(21.9)	0.165
300L	50(19.7)	19(29.2)	69(21.6)	0.100
400L	71(28.0)	17(26.2)	88(27.6)	
500L	22(8.7)	1(1.5)	23(7.2)	

600L	17(6.7)	2(3.1)	19(6.0)	
Faculty				
Arts	50(19.7)	18(27.7)	68(21.3)	
Engineering	36(14.2)	11(16.9)	47(14.7)	
Environmental	15(5.9)	7(10.8)	22(6.9)	
Law	3(1.2)	3(4.6)	6(1.9)	
Pharmacy	19(7.5)	0(0.0)	19(6.0)	
Science	32(12.6)	7(10.8)	39(12.2)	0.255
Social Science	14(5.5)	3(4.6)	17(5.3)	0.200
Basic Medical	19(7.5)	3(4.6)	22(6.9)	
Education	12(4.7)	3(4.6)	15(4.7)	
Clinical Science	20(7.9)	3(4.6)	23(7.2)	
Business	18(7.1)	4(6.2)	22(6.9)	
Administration	16(6.3)	3(4.6)	19(6.0)	
Agriculture				

182 *significant at 5%(p<0.05).

Table 2: Distribution of academic stress by blood group phenotype of the respondents

Blood Phenotype	Stress (n=254)	Not Stress (n=65)	Total (n=254)	Chi-square statistic	p-value
А	74(29.1)	20(30.8)	94(29.5)	0.467	0.926^{NS}
В	22(8.7)	5(7.7)	27(8.5)		
AB	40(15.7)	12(18.5)	52(16.3)		
0	118(46.5)	28(43.1)	146(45.8)		

NS= not significant at 5 %(p>0.05).

Table 3: Distribution of Reaction to Stress by Blood Phenotype

Reaction to	Α	В	AB	0
Stress	(n=74)	(n=22)	(n=40)	(n=118)
Emotional	72(97.3)	19(86.4)	17(42.5)	40(33.9)
Cognitive	28(37.8)	8(36.4)	30(75.0)	56(47.5)
Behavoural	51(68.9)	13(59.1)	33(82.5)	76(64.4)
Physiologica	38(51.4)	12(54.5)	19(47.5)	108(91.5)
1				
χ^2 value	22.92	4.77	7.63	36.80
p-value	< 0.0001	0.189	0.054	< 0.0001

Coping Strategies	A (n=74)	B (n=22)	AB (n=40)	0 (n=118)
Practical	69(93.2)	19(86.4)	38(95.0)	116(98.3)
Distracting	64(86.5)	20(90.9)	34(85.0)	95(80.5)
Religious	62(83.8)	18(81.8)	37(95.5)	114(96.6)
χ^2 value	0.400	0.286	0.018	2.480
p-value	0.8129	0.867	0.991	0.289

Table 4: Distribution of Coping Strategies by Blood Phenotype.

Table 5: Logistic Regression showing Association between Stress and Blood Phenotype

Variable	В	S.E	Wald	OR(95% C.I)	p-value
Blood Phenotype			1.491		
А	-	-		1.00(reference)	0.684
В	-0.155	0.577	0.07	1.17[0.377 - 3.623]	0.788
AB	-0.189	0.584	0.11	0.83[0.264 - 2.598]	0.746
0	-0.262	0.631	0.172	0.77[0.223 - 2.652]	0.679

205 Adjusted for socio-demographics of the respondents.

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Table 6: Logistic Regression showing association between reaction to academic
 stress and blood phenotype

Reaction to Stress	Blood Group	OR unadjusted (95% C.1)	p-Value
	А	1.00 (reference)	
Emotional	В	2.13[1.459 - 2.799]	0.007**
reaction	AB	1.47[0.555 - 3.873]	0.439
	0	0.95[0.397 - 2.271]	0.786
Cognitive			
reaction	А	1.00 (reference)	

	В	5.68[1.866 - 36.486]	0.006**
	AB	1.71[0.429 - 6.773]	0.448
	0	0.74[0.172 - 3.223]	0.693
Behavioral	А	1.00 (reference)	
reaction	В	1.25[0.494 - 3.174]	0.635
reaction	AB	2.08[1.684 - 6.309]	0.019*
	0	1.54[0.575 - 4.09]	392
Dh	А	1.00 (reference)	
Physiological	В	0.88[0.338 - 2.286]	0.792
reaction	AB	0.754[0.266 - 2.141]	0.754
	0	2.75[1.302 - 10.877]	0.042*

Adjusted for Socio-demographics characteristics.

*Significant at 5% (p<0.05),

**Significant at 1% (p<0.01).

217	Table 7: Logistic Regression and 95% confidence Interval showing Relationship
218	between Blood Phenotype and Coping Strategies

Coping Strategies	Blood Phenotype	OR (95% C.1)	p-value
	А	1.00 (reference)	
Drastical Coming	В	1.25[0.321 - 4.113]	0.632
Practical Coping	AB	2.28[1.011 - 6.885]	0.032*
	0	1.41[0.933 - 10.892]	0.632
Distruction	A	1.00 (reference)	
Distracting Coping	В	3.22[1.282 - 10.882]	0.036*
Coping	AB	2.12[0.952 - 6.683]	0.583
	0	1.20[0.833 - 4.021]	0.933
	А	1.00 (reference)	
Religious Coping	В	1.95[1.358 - 10.588]	0.044*
	AB	1.82[1.252 - 4.049]	0.049*
	0	0.95[0.178 - 2.393]	0.519

Adjusted for socio-demographics characteristics.

*Significant at 5%(p<0.05), **Significant at 1%(p<0.01).

225 Discussion:

The results of this study showed that undertaking a university education is a 226 highly stressful venture, and how this stress is perceived, responded to, and coped 227 with could be influenced by individual's blood type. Evidence for this notion is 228 twofold; first, similar to the results of the present study, previous investigators have 229 consistently recorded high prevalence of stress among college students. Here we 230 found that 79.6% of students were stressed. In a similar study at the University of 231 Calabar, Oku et al [15] reported 94.2% prevalence of stress among undergraduate 232 students. In 2015, Melaku [16] reported 52.4% prevalence of stress among college 233 students in Ethiopia. Likewise studies in other populations [17-23] have 234 demonstrated similar high frequency of stress among college students, although 235 variations in frequency across population have been observed, likely due to the 236 237 notion of individual and ethnic diversity in stress perception.

Second, perceived stress was more prevalent (46.5%) among individuals with blood 238 group O compared with other blood groups. This is consistent with the findings of 239 Chaudhuri et al [24] in a similar study in eastern India. In this study, perceived 240 stress score was significantly higher in group O compared to group A individuals, as 241 observed in the present study. This observation is despite the fact that individuals 242 of blood group O have higher threshold limit for stress than individuals of blood 243 244 group A. They secret less stress hormones (Cortisol and adrenaline) and hence less likelihood of being stressed compared with blood group A individuals known to 245 secret high levels of these stress hormones, and consequently have higher tendency 246 for stress. 247

A plausible explanation for this observation is partly due to the fact that blood group O is the most prevalent blood group in the general population of Nigeria and elsewhere. The preponderance of blood group O over other blood groups have been reported in several studies conducted in different parts of Nigeria and elsewhere, although with varying frequencies likely due to the underlying genetic and ethnic diversity of human populations [10, 25]. For instance, in a recent review to establish

a reference data base on the distribution of ABO and Rh blood groups from studies 254 conducted in different parts of Nigeria, Anifowoshe et al[10] reported a consistent 255 pattern of blood group O preponderance (52.93%). In a retrospective study among 256 blood donors in Sokoto, Nigeria, Musa et al [26] recorded a consistent pattern of 257 258 blood group O predominance in 51.91% of the participants. Medugu et al [27] found that 47.7% of pregnant women in Yola, Adamawa state, Nigeria were of blood group 259 260 O. Likewise Hassan et al [28] reported the prevalence of 49.2% for blood group O in a 5 year retrospective survey of ABO blood group distribution among blood donors 261 in Ahmadu Bello University Teaching Hospital, Kaduna, Nigeria. 262

263 A similar multi-race/ethnic study in United State reported a consistent pattern of blood group O preponderance [25]. Therefore, individuals of blood group O 264 constituting the largest proportion of stressed participants could partly be corollary 265 266 of the higher prevalence of blood group O in the general population of Nigeria. Also, there is delayed release of stress hormones (cortisol and catecholamines) following 267 268 stressful encounter in individuals of blood group O, however, when released, the metabolism of these hormones is delayed because the enzyme of monoamine oxidase 269 (MAO) which is working among others to enhance the metabolism of catecholamines 270 such as adrenaline and nor-adrenaline is less active in platelets of blood group O 271 individuals. Therefore, stress episodes/effects tend to be more sustained in 272 individuals with blood group O than those with blood group A. Group A individuals 273 respond to stress by releasing a large amount of adrenaline, which is also 274 eliminated quickly to reduce the effect of stress. 275

Physiological reaction to stress was more prevalent in blood group O than A, while emotional was more prevalent in blood group A than O. The most common physiological reactions among blood group O individuals were feeling of uncoordination, headache, lower back pain, sweating and chills, stomach upset and changes in appetite, probably due to the effect of the secreted catecholamines. The behavioral and emotional reactions also observed in some group O individuals may be due to the longer time of stress in these individuals which is associated with exhaustion, and subsequently to anger and aggression. Most prevalent reports were the feeling of fear, guilt, apathy and decreased efficiency and effectiveness. Common coping strategies adopted by these individuals were religious, active practical and active distracting. Some of them used reading of bible, meditation, praying for guidance to cope with their stresses. Others watched movies, involved in physical activities or embarked on a short visit to friends.

Blood group A individuals reacted mostly emotionally, behaviorally and 289 physiologically to stress, and least of cognitive reaction. Most participants in this 290 group agreed that they forget things easily. They have limited attention span and 291 292 difficulty in concentrating. Some of them reported having problem with sleep and are easily annoved. Few others suffered from headache and poor appetite. Group A 293 individuals employed mainly active practical, active distracting and religious coping 294 295 methods to lessen their stress levels. They naturally have a higher serum cortisol level than other blood groups, and when stressed, serum level of adrenaline rises 296 more than in other blood groups. They are designated as the worst group in 297 reducing their stress levels. Blood group B displayed marked emotional and 298 moderate behavioral and physiological reactions to stress. They employed mostly 299 active distracting, active practical and religious methods to allay their stresses. 300 They naturally have large amount of cortisol and release more adrenaline during 301 302 stress although not as much as blood group A individuals. However, blood levels of these stress hormones are reduced immediately, thereby enhancing faster recovery 303 from stressful encounter similar to blood group A. 304

Individuals with blood group AB showed marked behavioral and cognitive reaction to stress and employed mainly practical, religious and distracting coping strategies to manage their stresses and reduce stress hormone slowly almost similar to blood group O. This study has few limitations especially those related to the crosssectional nature of the study design. Also, self reporting bias should be considered in the interpretation of results.

312 Conclusion	n
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312	Conclusion:
313	There is a correlation between individual's blood type and the odds of being
314	stressed, reaction to stress and coping strategy employed to cope with stress.
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316	Conflict of interest: None declared
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