Original Research Article

2 PREVALENCE OF INTESTINAL PARASITOSIS AMONG MENTALLY ILL PERSONS 3 IN NEUROPSYCHIATRIC HOSPITAL, CALABAR, NIGERIA

4

5 ABSTRACT

Aim: The prevalence of intestinal parasitosis in relation to mental illness among inpatients of Federal Neuropsychiatric Hospital, Calabar was investigated with a view to improving the quality of their medicare.

Study design: This was cross sectional study. Ethical approval and patients' informed consent were sought and obtained before collection and processing of samples.

Place and Duration of Study: Federal Neuropsychiatric Hospital, Calabar, between February and August, 2016.

Methodology: We included 246 (126 mental, 120 apparently normal) subjects. Intestinal parasites were detected by direct stool microscopy and formol ether concentration technique and the cellophane anal swab method for *Enterobius vermicularis*. Chi square analysis was used to compare percentages.

Results: The prevalence of intestinal parasitosis among the 126 studied mental subjects was 49.2% compared to 11.7% recorded among the 120 apparently healthy (control) subjects examined (P < 0.0001). There was no significant association between intestinal parasites prevalence and gender in mental as well as normal subjects 60.0% v. 45.1%, respectively, p = 0.2022) and 14.3% v. 8.8%, respectively, P = 0.3113). Subjects with chronic mental cases insignificantly harbored more intestinal parasites than those with acute cases (53.8% v. 47.1%, respectively; P = 0.5699). Parasites detected among mental subjects were Hookworm 34.6%, *Entamoeba histolytica/dispar* 25%, *Ascaris lumbricoides* 17.3%, *Trichuris trichiura* 9.6%, *Enterobius vermicularis* 5.8%, *Giardia intestinalis* 3.8%, *Taenia* species and *Schistosoma mansoni* 1.9% each. Hookworm ranked highest 34.6% among the parasites encountered in the test subjects while *Ascaris lumbricoides* 33.3% took the lead in the control group. People diagnosed with Variance Psychosis disorder recorded highest infection rate (68.4%) while those with substance abuse had the least (28.0%).

Conclusion: Intestinal parasitosis may possibly play aetiologic or enhancement role in mental health problems. For effective management of the mental challenge, periodic parasitic investigations and appropriate therapy before and after admittance should be effected in all psychiatric institutions.

6 7 8

Keywords: Mental Illness, Psychiatric disorder, Intestinal parasitosis, Prevalence, Calabar.

9

10 **1. INTRODUCTION**

11

Intestinal parasites are associated with significant morbidity and mortality worldwide, particularly in

12 developing countries [1] where an estimated 3.5 billion people are affected, the majority being children [2],

13 the aged, and institutionalized persons [3]. Socioeconomic and environmental factors including poor

14 personal, food hygiene, and sanitation, poverty, and overcrowding have been documented to promote

15 intestinal parasitic infections [4]. These infections may lead to low nutritional and immune status, impaired

16 growth, and poor cognitive performance [5] which complicate and contribute to other illnesses [6, 7, 8, 9].

Mental illness refers collectively to all of the diagnosable mental and emotional conditions and are characterized by abnormalities in cognition, emotion or mood, or the highest integrative aspects of behavior, such as social interactions or planning of future activities [10, 11]. When the mental illness significantly interferes with the performance of major life activities such as learning, working and communicating, among others, the term "psychiatric disability" is applied [11]. It is a core tenet of modern science that behavior and our subjective mental lives reflect the overall workings of the brain as the mental functions are all mediated by the brain [10].

Mental illnesses are disabling and common globally but underestimated and under-treated in many developed and developing countries with estimated global burden between 21.2% - 32.4% of years lived with disability (YLDs) [12] and around 450 million people currently suffering from such conditions worldwide [13].

An estimated 20%–30% of the Nigerian population are believed to suffer from mental disorders [14]; unfortunately, inadequate attention is given to this health problem as only about three percent of the government's budget on health, according to the World Health Organization estimates, goes to mental health [15]. This, coupled with misconceptions and poor level of awareness of the Nigerian public on mental health issues [16], has compounded this problem. Despite several previous reports of high prevalence of intestinal parasitosis in many mental facilities elsewhere, very limited studies have been documented in Nigeria.

The cause of mental illness and psychiatric disorders is controversial but a combination of factors, including biological, psychological, environmental, social and spiritual factors have been incriminated [17, 18]. Genetic and biological factors have been associated with schizophrenia, depression, and alcoholism [17]. There are records of accumulated evidence showing Schizophrenia and Bipolar Disorder as complex diseases in which many predisposing genes interact with one or more environmental agents to cause symptoms, *Toxoplasma gondii* being an example of infectious agent that has been linked to schizophrenia and in which genes an infectious agent interact. Such infections may occur early in life and 42 are thus consistent with neurodevelopmental as well as genetic theories of psychosis [19]. 43

Risk factors of mental disorders include traumatic brain injury [20], substance abuse, viral 44 45 infections [21], and general physical health. Researches show that a number of mental disorders occur in 46 people suffering from other diseases more often than would be expected by chance as in infection, 47 Coeliac disease, etc and this is often never investigated. Currently, science believes that mental illness is 48 linked to genetic flaws. However, recent medical research also points to immune system dysregulation, 49 most likely originating from gastrointestinal dysfunction as another factor. Psychiatrist Dr C.M. Reading, 50 after over 30 years of practice believed that a significant percentage of those with mental illness suffer due 51 to gastrointestinal & physical problems manifesting as Coeliac or latent Coeliac disease, food allergies, 52 infections, auto-immune disease and malabsorbtion [22]. The of role the 53 gastrointestinal system in the development of many illnesses, especially mental illness and neurological disorders has often been overlooked. Recent research findings have linked mental disorders to 54 microbiota - brain- gut interaction (Brain-Gut Axis) through adjustment in the gut microbiota and activation 55 of certain immune system cells in response to an infection, or on an ongoing basis (chronic inflammation) 56 57 and studies are under way to evaluate the use of anti-inflammatories in treating depression and 58 schizophrenia [23].

59 Repeated immune response due to infection or allergy may result in inflammation, particularly in 60 the area of the small bowel and over time this may lead to damage of the mucosal villi and in turn increase 61 mucosal permeability. Recurrent gastrointestinal infection, gastritis, post antibiotic infection (colonization 62 of bad bacteria), tropical sprue and inherited gastro-immunological disorders such as coeliac sprue, non-63 coeliac sprue and food intolerances may lead to the development of mental illness and disease [22]. 64 Inflammation plays a key role in mood disorders and mental illness. When inflammatory antibodies cross 65 the blood-brain barrier, it interferes with the brain's ability to function. The immune system secretes 66 antibodies that are distributed in the blood to help fight the infection or repair the problem. The blood-brain 67 barrier is supposed to protect the brain from those antibodies. But for yet unknown reasons, when

68 inflammation reaches the brain, the cytokines wreak havoc on the neurotransmitters, interfering with the 69 brain function [24]. Auto-immune response can lead to symptoms like anxiety, depression, and 70 hallucinations [25] and some studies have even found higher levels of inflammation in patients with 71 depression and suicidal thoughts, PTSD, and chronic fatigue [26, 27].

The relationship between mental and medical illnesses further emphasizes the need for continued mental health research for proper diagnosis of psychiatric illness [28].

The impact of mental illness is grave on the victims, their family members and the communities [29-31]. In addition to the health and social costs, those suffering from mental illnesses are also victims of human rights violations, stigmatization and discrimination, both inside and outside psychiatric institutions. Many psychiatric institutions have inadequate, degrading and even harmful care and treatment practices, as well as unhygienic and inhuman living conditions [32]. With memory loss, poor reasoning, low education, poverty, and low hygiene level, mentally ill persons are likely to acquire and spread infectious agents which in turn promotes the already underlying health problem.

81 Although a relationship between infectious diseases and psychiatric disorders has been 82 suggested, this relationship is yet to be well demonstrated and be considered important by many health 83 care providers [8, 17]. A better understanding of the role of infection may speed treatment and prevention efforts and reduce the degree of disability and stigma associated with mental illness. This study therefore, 84 85 aimed to investigate the prevalence of intestinal parasites among hospitalized psychiatric patients in the 86 Federal Neuropsychiatric Hospital, Calabar and its relationship with demographic variables of interest. 87 Related risk factors and consequences of intestinal parasitic infections among the study subjects were 88 also assessed and action initiated for treatment of the infected persons.

89 90

91

2. MATERIALS AND METHODS

This study was carried out at the Federal Neuropsychiatric Hospital, Calabar situated in the tropical rain forest of Southern Nigeria between February and August, 2016. It is the only psychiatric institution in the state and serves as a home as well as hospital for people with mental illnesses from across the state

95 and neighboring states. At the time of the study, the institution had a 235 bed capacity with 181 96 inmates/inpatients, mostly adults. With ethical approval from the hospital research ethics 97 committee and informed consent of participants, a total of one hundred and twenty six (126) patients of the hospital who complied and one hundred and twenty (120) apparently 98 99 normal subjects from the general population without any history of anti-parasitic medication 100 in last preceding month were recruited for the study. Demographic data and health status of each studied 101 subjects were obtained through the instrument of interviewer- administered questionnaire, with the help of 102 the hospital/faculty workers, based on medical records. Subjects were also questioned for recent 103 abdominal discomforts, diarrhoea, anal itching or emergence of nematodes from the anus, and anti-104 parasitic medication.

All studied subjects were screened for intestinal parasites based on stool and anal swab 105 106 examination. The subjects or their informants were each given a sterile screw-capped wide mouth 107 universal container for collection of stool sample and cellophane anal swab for collection of anal swab. 108 Anal swabs of the subjects were obtained early in the morning (before defecating/bathing) using the 109 cellophane anal swab (8 by 2- cm of transparent adhesive tape), held sticky side out, over the end of a 110 glass microscope slide [33]. Samples were processed and examined in the University of Calabar 111 Teaching Hospital laboratory. The stool samples were macroscopically examined for appearance, 112 consistency, presence of blood, mucus, worm segments and worms. Stool samples were further 113 processed by direct smear microscopy and formol ether concentration technique. The preparations and the anal swabs were examined microscopically using the 10x and 40x 114 115 objectives for the presence of larva, cysts, and eggs of parasites [6]. Percentages were compared using 116 Chi square analysis. All infected subjects were recommended for appropriate treatment.

117 3. RESULTS

118 The results of this study are as shown in Figures 1-4 and Tables 1-2.

Fig. I displays the prevalence of intestinal parasites among psychiatric and normal subjects in Calabar. Sixty two (49.2%) of the 126 test subjects versus 14 (11.7%) of 120 normal (control) subjects examined significantly tested positive for parasites (P < 0.0001).



123 Mental as well as normal female subjects were insignificantly more infected than their male counterpart







134	Chronic	39	21 (53.8)		
	Total	126	62 (49.2)		
135 136 137	Key: Acute - N	ewly admitted patients who spent less than 4 w	with sudden onset, high de	egree but short-term	condition
138 139 140 141	Chronic - P wl ha	atients with long-lasting no sometimes acted no allucination, delusions a	condition (who spent mor rmally, but at other times s nd the like.	e than 4 weeks at th suffered from periods	e care center), s of rage,
142	The most fre	equently detected para	site among the psychiatric	c patients was hook	worm 34.6% followed
143	by Entamoeba h	istolytica/dispar 25.0%.	Other parasites detected	d in descending ord	er of frequency were
144	Ascaris lumbrico	ides 17.3%, Trichuris	trichiura 9.6%, Enterobu	iis vermicularis 5.8%	6 (in one sampling),
145	Giardia intestina	lis 3.8%, Taenia spec	ies and Schistosoma ma	ansoni 1.9% each,	whereas among the
146	controls, Ascaris	ranked highest 33.3%	followed by hookworm and	d E. histolytica/dispa	r 22.2% each,

147 *Enterobius* and *Giardia* 11.1% each (Fig. 3).



148

149 Fig. 3: Frequency of intestinal parasites among mental and normal subjects

150 Table 2 shows occurrence of mixed parasitic infections among psychiatric patients. The rate of single

151 infection was higher than that of mixed infection 68.1% and 31.9%, respectively. Mixed infections with 2

152 parasites were more often encountered 86.4% than those with 3 parasites 13.6%.

153 Table 2: Occurrence of Mixed parasitic infections among psychiatric patients

	Parasites grouping/No.	Occurrence	Frequency (%)	
155	Mixed Infections			
156	2 Parasites	19	(27.5)	
157	3 Parasites	3	(4.3)	
158				
159	Subtotal	22	(31.9	
160	Single Infection		·	
161	1 Parasite	47	(68.1)	4
	Total	69	(100.0)	
162				1

The highest prevalence of infection was recorded among persons diagnosed with Variance Psychosis 68.4%, closely followed by those with Mania disorder 63.6%, Acute Schizophrenia 56.2%, Chronic Schizophrenia 48.2%, Depression 46.7%, Organic Source 45.4%, while those with Substance Abuse recorded the least prevalence 28.0% (Fig. 4).



180 against 11.4% recorded among the controls (P < 0.0001). These findings are similar to those of 181 Sirivichayakul et al. [34] in Thailand where a higher prevalence was also observed among 182 institutionalized mentally handicapped than non-institutionalized normal individuals (57.6% v. 7.5 %, respectively, p < 0.05). The 49.4% prevalence recorded in this study is high compared to 8.4% recorded 183 in mental hospitals in North Taiwan [35], 7.3% in New York [36], 20.5% in Urmia, Iran [37], 35.7% in 184 Korea [38] but is related to 52.3% reported in Puerto Rico [39], 53.8% in Italy [8], and 76.7% in Egypt 185 186 [40]. The higher prevalence observed among our studied subjects compared to controls may be 187 attributed to the poor state of hygiene, usually observed among this (institutionalized) group of persons, 188 coupled with their abnormal behavior, limited access to anti-parasitic therapy, poor environmental 189 conditions and poor sanitary practices within the facility.

190 The prevalence rate observed in the study area, being hospital environment, may not necessarily 191 reflect the endemicity level of intestinal parasitoses in the general population of Calabar as Calabar wide 192 prevalence data on intestinal parasitoses is lacking and most of the mentally ill persons are victims of 193 displaced homes (may not be Calabar residents), poverty, low levels of education, poor hygiene and feeding problems which are known risk factors of parasitoses. It is believed that those persons were 194 195 harboring these infectious agents before admittance into the hospital, since no routine medical 196 examination was done on their admission. However, the possibilities of hospital-acquired infection and 197 inter-hospital transfer are inevitable.

The higher prevalence of parasites (49.4%) noted among the test subjects in this study may not necessarily attribute their mental illnesses to intestinal parasitoses but our results being consistent with those reported elsewhere, Puerto Rico 52.3% [39], Italy 53.7% [8], and Egypt 76.7% [40] is suggestive that intestinal parasitosis may possibly play etiologic or enhancement role in mental disorders.

Mental as well as the control females were insignificantly more infected with intestinal parasites than their males counterpart (60.0% v. 45.1%, respectively, P = 0.2021) and (14.4% v. 8.8%, respectively, P = 0.3113). This differs from the findings in Iran [36] where infection rates among males and females were similar (20.3% v. 20.5%, respectively). The prevalence of intestinal parasites among the psychiatric patients insignificantly increased with duration of cases as subjects with chronic (long-lasting) condition showed a higher parasites prevalence (53.8%) than those with acute (short-term) mental problems (47.4%) (P = 0.5699). This may be attributed to difference in the degrees of chronic stresses and immune system deficiency (which might be higher in the chronic cases) usually associated with mental disorders which are also known risk factors of parasitosis [41].

Hookworm ranked highest among the parasites encountered in the test subjects 34.6% followed by Entamoeba histolytica/dispar 25%, Ascaris lumbricoides 17.3%, Trichuris trichiura 9.6%, Enterobius vermicularis 5.8% (in one sampling), and Giardia intestinalis 3.8%, Taenia species and Schistosoma mansoni 1.9% each whereas, among the controls, Ascaris ranked highest 33.3% followed by hookworm and *E. histolytica/dispar 22.2*% each, *Enterobius* and *Giardia* 11.1% each.

The high prevalence of hookworm here may be attributed to the habit of walking bare foot, observed among some of the test subjects, and poor hygiene related to faeces. *Taenia* species and *Schistosoma mansoni* encountered here probably represent imported infection and have been related to organic sources of mental disorder and their associated symptoms [8, 9]. According to various authorities, taeniasis is thought to be the cause of psychiatric symptoms due to its neural and psychological effects. These claims have been confirmed in a taeniasis case study of a 36 year-old woman whose psychiatric symptoms (obsessive and compulsive neurosis and depression) decreased after the taeniasis treatment [11].

Entamoeba histolytica/dispar was the leading protozoan infection (25%) in this study. Although this study was limited to stool examination, this organism has been previously reported to be capable of causing extra intestinal infection in other parts of the body including the brain [42]. Amoebic brain abscesses which have been rarely reported [43], result when trophozoites invade the central nervous system [44] leading to headache, altered mental confusion, focal neurologic signs and seizures.

230 Extra intestinal migration of *Enterobius vermicularis* has been documented to result in severe health 231 disorders, including nervousness or even death, especially in population dense areas and institutionalized persons [45, 46]. Among the parasite positive persons, single infections recorded 65.9%, mixed infections
34.1% (31.7% double and 2.4% triple) prevalence.

The highest prevalence of parasites was noted among subjects diagnosed of Variance Psychosis 68.4% closely followed by those having Mania 63.6%, Acute Schizophrenia 56.2% while those with Substance Abuse had the least prevalence 28.0%. Poor mental health in association with parasitic infections may produce extreme anxiety with recurrent attacks of Mania and Depression [10]. The least parasites prevalence, 28.0% observed among Substance Abuse subjects suggests the need to investigate hard substances (cocaine, cannabis, etc.) and their lethal effect on intestinal parasites.

240 A number of mental disorders have often been tentatively linked with microbial pathogens [47]. 241 particularly viruses and parasites [48]. Taenia solium, Naegleria fowleri, and Toxoplasma gondii are all 242 parasites that have been documented to infect the human brain resulting in symptoms such as headaches, 243 fever, confusion, nausea, seizures, loss of balance, and hallucinations with Toxoplasma being the cause of 244 most cases of schizophrenia and bipolar disorder [21]. Acute infection with Toxoplasma gondii has been 245 shown to produce personality changes and psychosis; incidence of infection in schizophrenic patients being 246 twice that of control subjects (42 % versus 11 %, respectively). T. gondii is usually spread to humans from 247 cats, exposure to cats in childhood revealed in two studies as a risk factor for the development of 248 schizophrenia [8]. The parasite has been documented might play a role in the development of these 249 disorders by affecting the production of dopamine -- the chemical that relays messages in the brain 250 controlling aspects of movement, cognition and behavior [49]. The Toxoplasma gondii parasite has been 251 linked, in another study, to the brain cells damage leading to suicide attempts [50] while meningitis or 252 encephalitis was found in 24 % of 1300 cases of trichinosis reported from Germany [8].

Psychiatric Disorders are illnesses of the brain and parasitic infections have been documented, could alter normal functions by depleting the host's essential nutrients, interfering with enzyme and neuroimmune functions, and releasing massive amounts of waste products, enteric poisons, and toxins which may disable brain metabolism [8]. Previous reports show that tape worms have been associated with direct brain invasion (as in neurocysticerccosis) leading to depression and psychosis [8]. These tapeworms could produce cysts, swelling, and encephalitis in brains of patients. Pittella [51] linked neurocysticercosis with seizures, increased intracranial pressure, ischemic cerebrovascular disease, dementia, and signs of
 compression of the spinal roots/cord.

There are several reports [51, 52, 53] on neuroschistosomiasis caused by *Schistosoma mansoni* infection. These worms can evoke granulomatous inflammatory reaction when eggs are being transmitted to the spinal cord or brain via the vascular system, or by inadvertent adult worms' migration to these organs resulting in psychiatric symptoms, including seizures and increased intracranial pressure [51].

Other parasitic infections (giardiasis, ascariasis, etc.) may produce mental symptoms indirectly through
 brain – gut - axis (BGA) which may clear after effective therapy [8].

The relationship between mental health problems and parasitic infections, although yet to be well demonstrated, is real and needs to be given concrete consideration by health care providers. The results obtained here further stresses the need for continued investigation on intestinal parasitosis and mental health.

271 5. CONCLUSION

272

This study has revealed a high prevalence of intestinal parasitic infections among institutionalized mentally ill patients in the Federal Neuropsychiatric Hospital compared to apparently healthy persons in the general population in Calabar, Nigeria. There is no prevalence data on intestinal parasitoses among mentally ill people in Calabar and in Nigeria, such information is sparsely documented. The study strongly suggests that intestinal parasites may play aetiologic or enhancement role in mental health problems. Periodic parasitic investigations and appropriate therapy before and after admittance should be effected in all psychiatric institutions.

280 6. SIGNIFICANCE OF THE STUDY

The findings of this study will improve the quality of medicare of patients treated at Federal Neuro-Psychiatric Hospital, Calabar as intestinal parasitosis would be considered during clinical and laboratory diagnosis.

284

286 CONSENT

- 287 All authors declare that written informed consent was obtained from the patients/guardians after
- 288 details of the study was explained to them, before recruitment into the study.

289 INCLUSION AND EXCLUSION CRITERIA

- 290 Participation in the study was strictly voluntary. Only those subjects who gave their consent with
- 291 compliance and were not on any antiparasitic and hematinic medication were included in the study. On the
- 292 other hand, those who refused to give their consent were left out of the study.

293 ETHICAL ISSUE

- 294 Ethical approval for the study was obtained from Committee on Research Ethics, Federal Neuro-
- 295 Psychiatric Hospital, Calabar.
- 296 **CONFLICT OF INTERESTS:** None declared.

297

298

300

299 **REFERENCES**

- 301 1. Hotez PJ. Reducing the global burden of human parasitic diseases. *Comp Parasitol* 2002; 69: 140-5.
- 302
 303
 304
 305
 305
 306
 307
 308
 309
 309
 309
 3000
 3000
 3000
 3000
 3000
 3000
 3000
 3000
 3000
 3000
 3000
 3000
 3000
 3000
 3000
 3000
 3000
 3000
 3000
 3000
 3000
 3000
 3000
 3000
 3000
 3000
 3000
 3000
 3000
 3000
 3000
 3000
 3000
 3000
 3000
 3000
 3000
 3000
 3000
 3000
 3000
 3000
 3000
 3000
 3000
 3000
 3000
 3000
 3000
 3000
 3000
 3000
 3000
 3000
 3000
 3000
 3000
 3000
 3000
 3000
 3000
 3000
 3000
 3000
 3000
 3000
 3000
 3000
 3000
 3000
 3000
 3000
 3000
 3000
 3000
 3000
 3000
 3000
 3000
 3000
 3000
 3000
 3000
 3000
 3000
 3000
 3000
 3000
 3000
 3000
 3000
 3000
 3000
 3000
 3000
 3000
 3000
 3000
 3000
 3000
 3000
 3000
 3000
 3000
 3000
 3000
 3000
 3000
 3000
 3000
 3000
 3000
 3000
 3000
 3000
 3000
 3000
 3000
 3000
 3000
 3000
 3000
 3000
 3000
 3000
 3000
 3000
 3000
 3000
 3000
 3000
 3000
 3000
 3000
 3000
 3000
- 307 3. Chan MS. The global burden of intestinal nematodes infections- fifty years on. *Parasitol Today* 1997;
 308 13: 436-43.
- 309

- Al-Mohammed HI, Amin TT, Aboulmagd E, Hablus HR, Zaza BO. Prevalence of intestinal parasitic infections and its relationship with socio-demographics and hygienic habits among male primary school children in Al-Ahsa, Saudi Arabia. Asian Pac J Trop Med. 2010; 3:906–12.
- 314 5. de Silva NR, Guyatt HL, Bundy DAP. "Morbidity and mortality due to Ascaris-induced intestinal obstruction," *Transactions of the Royal Society of Tropical Medicine and Hygiene*. 1997; vol. 91, no. 1, pp. 31–36.
- 317
- Arora DR, Arora B. Medical Parasitology. 2nd Ed. New Delhi, Bangalore, India: CBS Publishers and Distributors; 2005; p. 213-6.
- Read CP. The vertebrate small intestine as an environment for parasitic helminthes. *The Rice Institute Pamphlet* 1990; 37: 1-94.
- 323
 324 8. Howenstine Dr. J. The overlooked relationship between infectious diseases and mental symptoms.
 325 *http://www.newswithviews.com/Howenstine/james16.htm#_ftn1.*2004 [accessed 6 May 2015].
 326

- Inceboz T, Yalçin G, Aksoy U. Case report: taeniasis, is it a cause of psychiatric and neural symptoms?
 Turkiye Parazitol Derg 2006; 30(3): 187-9.
- 329
- Mental Health: A Report of the Surgeon General, Chpt 2. The Fundam- entals of Mental Health and
 Mental Illness. *http://www.Surgeongeneral. gov/library/mentalhealth/chapter8/sec1.html.* [accessed 15
 May 2015].
- 334 11. Zuckerman D, Debenham K, Moore K. The ADA and People with Mental Illness. A Resource Manual
 for Employers. Available from the National Mental Health Association, 1021 Prince Street, Alexandria,
 VA22314-2971: 1993; (703): 684-722.
- 337

Vigo D, Thornicroft G, Atun R. Estimating the True Global Burden of Mental Illness. Lancet Psychiatry.
 2016; 3 (2): 171-78 Vigo D, Thornicroft G, Atun R. Estimating the True Global Burden of Mental Illness.
 Lancet Psychiatry. 2016; 3 (2): 171-78.

- WHO, 2001: The world health report 2001 Mental Health: New Understanding, New Hope WHO,
 2001: The world health report 2001 Mental Health: New Understanding, New Hope.
- 345 14. Onyemelukwe C. Stigma and mental health in Nigeria: Some suggestions for law reform. J Law Policy
 346 Glob 2016; 55:63-8
- WHO-AIMS Report on Mental Health System in Nigeria, WHO and Ministry of Health, Ibadan, Nigeria,
 [2006]. https://www.who.int/mental_health/evidence/nigeria_who_aims_report.pdf. Retrieved 7 January
 2016.
- 352 16. Suleiman DE. Mental health disorders in Nigeria: A highly neglected disease. Ann Nigerian Med 2016;
 353 10:47-8.
- 354 355

361

- Torrey EF, Miller J. The Invisible Plague: The Rise of Mental Illness from 1750 to the Present.
 Piscataway, NJ 08854- n8099: Rutgers University Press, 2002; 314.
- 35918. Study Finds Serious Mental Illness Often Dismissed by Local Church. 360 *http://newswise.com/articles/view/545316/.* [accessed 15 May 2015].
- 36219. Yolken RH and Torrey EF. Are some cases of psychosis caused by microbial agents? A review of the evidence. *Molecular Psychiatry* 2008; 13, 470–479; doi:10.1038/mp.2008.5
- 364
 36520. Jesse R. Fann, MD, MPH; Bart Burington, MS; Alexandra Leonetti, MS; Kenneth Jaffe, MD; Wayne J.
 366 Katon, MD; Robert S. Thompson, MD. Psychiatric illness following brain injury in an adult health
 367 maintenance organization population. *Arch Gen Psychiatry* 2004; 61(1): 53-61.
- 368
 369 21. Yolken RH, Torrey EF. Viruses, schizophrenia, and bipolar disorder. *Clin Microbiol Rev* 1995; 8(1):
 370 131-45.
- 370 371
- 372 22. www.GutandMentalillness.com. The Gastrointestinal origin of mental Illness and complimentary
 373 treatment strategies. *Based on the writings of Psychiatrist Dr C.M. Reading and Jordan Rubin* 374 [accessed 17 February 2016].
- 375 23. Haapakoski R, Mathieu J, Ebmeier KP, Alenius H, Kivimäki M. Cumulative meta-analysis of
 interleukins 6 and 1β, tumour necrosis factor α and C-reactive protein in patients with major depressive
 disorder. Brain Behav Immun. 2015; 49:206-15. doi: 10.1016/j.bbi.2015.06.001.

378		
379	24.	Rege, S. (2016). Autoimmune Diseases Masquerading as Psychiatric Disorders – A Paradigm Shift.
380		https://psychscenehub.com/psychinsights/autoimmune-diseases-masquerading. Accessed 28/1/2017
381		
382	25	Souhel Najjar Danjel M Pearlman Kenneth Alper Amanda Najjar and Orrin Devinsky
383	20.	Neuroinflammation and psychiatric illness. I Neuroinflammation 2013 10: 43 doi: 10.1186/1742-
384		2004_10_43
385		2004-10-43.
386	26	Holmes SE Hinz R Conen S Gregory CI Matthews IC Anton-Rodriguez IM et al Elevated
387	20.	Translocator Protein in Anterior Cinculate in Major Depression and a Role for Inflammation in Suicidal
388		Thinking: A Positron Emission Tomography Study, Biological Psychiatry, Journal 2018, Volume 83
380		Issue 1. Pages 61_69. DOI: https://doi.org/10.1016/j biopsych.2017.08.005
300		133de 1, 1 dges 01-03. DOI: https://doi.org/10.1010/j.biopsych.2017.00.003
301		
392	27	Montova JG Holmes TH Anderson JN Maecker HT Rosenberg-Hasson Y Valencia JJ et al
302	21.	Cytokine signature associated with disease severity in chronic fatigue syndrome natients PNAS 2017:
394		Volume 114 (34) E7150-E7158 https://doi.org/10.1073/pnas.1710519114
395		
396	28	Linda C. Mental Disorders Secondary to General Medical Condition –overview
397	20.	http://emedicine_medscape_com/article/294131-overview [accessed 1 March 2016]
398		
399	29	Veggeberg SK. The big story in depression: What isn't happening? <i>Brainwork-The Neuroscience</i>
400	20.	Newsletter, Vol. 7, http://www.dana.org/articles/bwn_1997, cfm, [accessed 28 August_2017].
401		
402	30.	National Institute of Mental Health. The impact of mental illness on society.
403		http://www.nimh.nih.gov/publicat/2001b/burden.cfm. [accessed 27 June 2015].
404		
405	31.	U.S. Department of Health and Human Services. Mental health: A report of the surgeon general.
406		http://www.surgeongeneral.gov/library/2000/mentalhealth/home.html. [accessed 27 December 2016].
407		
408	32.	World Health Organization. Investing in Mental Health. http://www.who.
409		int/mental_health/en/investing_in_mnh_fin-al. Pdf. 2003. [accessed 30 June 2016].
410		
411	33.	Faust EC, Russel PF, Jung RC. Craig and Faust's Clinical parasitology, 8 th Ed. Philandelphia PA: Lea &
412		Febiger 1970; 525-7.
413		
414	34.	Sirivichayakul C, Pojjaroen-anant C, Wisetsing P, Siripanth C, Chanthavanich P and Pengsaa K.
415		Prevalence of intestinal parasitic infection among Thai people with mental handicaps. Southeast Asian
416		J Trop Med Public Health 2003; 34(2):259-63.
417		
418	35.	Cheng HS, Wang LC. Intestinal parasites may not cause nosocomial infections in psychiatric hospitals.
419		Parasitol Res. 2005; 95(5): 358-62.
420	~~	
421	36.	Schupt N, Ortiz M, Kapell D, Kiely M, Rudelli RD. Prevalence of intestinal parasitic intections among
422		individuals with mental retardation in New York State. <i>Ment Retard.</i> 1995; 33(2):84–9.
423	27	Herroti Tenneh //h Mehemmedredek H. Neled Dekim D. Drevelence of Justic Keyl Deverifier
424	37.	Hazrati Tappen Kn, Monammadzaden H, Nejad Ranim R. Prevalence of Intestinal Parasitic
420		infections among Mentally Disabled Children and Adults of Ormia, Iran J Parasitol. 2010, 5(2).
420 427		00-4.
421 190	38	Lee I Park GM Lee DH Park SI Vong TS Intestinal parasite infections at an institution for the
+20 120	50.	handicanned in Korea. Korean I Parasital 2000: 38/3):170, 81
729 430		$\frac{1}{2}$
.00		

431 432 433	39.	Ferrer-Rodríguez I and Kozek W J. Prevalence of intestinal parasitoses among patients and staff of an institution for the mentally retarded. <i>J Parasitol and Vector Biol</i> 2011; 3(5): 69-74.
434 435 436	40.	Mohamed NH, Salem SA, Azab ME, Bebars MA, Khattab HM, Kamal AM. Parasitic infections associated with mental retardation in Egypt. <i>J Egypt Soc Parasitol</i> 1991; 21(2): 319–31.
437 438 439	41.	Beaton DB. Effects of Stress and Psychological Disorders on the Immune System. Rochester Institute of Technology at http://www.personalityresearch.org/papers/beaton.html. [accessed 9 October 2017] .
440 441 442	42.	Entamoeba and Amebiasis at http://www.garlandscience.com/res/pdf/9780815365006ch03. pdf. [accessed 18 July 2016].
443 444 445	43.	Banerjee AK, Bhatnagar RK, Bhusnurmath SR. Secondary cerebral amebiasis. <i>Trop Geogr Med</i> 1983; 35(4): 333-6.
446 447 448	44.	Maldonado-Barrera CA, Campos-Esparza Mdel R, Muñoz-Fernández L, Victoria-Hernández JA, Campos-Rodríguez R, Talamás-Rohana P, <i>et al.</i> Clinical case of cerebral amebiasis caused by <i>E. histolytica. Parasitol Res</i> 2012; 110(3): 1291-6.
450 451 452	45.	Markell EK, John DT, Krotoski WA. <i>Markell and Voge's Medical Parasitology</i> . Philadelphia: W.B. Saunders Company; 2006.
453 454 455 456	46.	Schmunis GA, Lopez-Antunano FJ. In: Cox FEG, Wakelin D, Gillespie SH, Despommier DD, Editor (s). <i>Toply and Wilson Microbiology and Microbial infections, Parasitology.</i> p. London: Edward Arnold. 2005; 24.
457 458 459	47.	Pearce BD. Modeling the role of infections in the aetiology of mental illness. <i>Clinical Neuroscience Research</i> 2003; 3 (4-5): 271-82.
460 461 462 463	48.	Thomas HV, Thomas DR, Salmon RL, Lewis G and Smith AP. Toxoplasma and Coxiella infection and psychiatric morbidity: a retrospective cohort analysis. <i>BMC Psychiatry</i> 2004; 4: 32. DOI: 10.1186/1471-244X-4-32
464 465 466 467	49.	University of Leeds (2009, March 11). Toxoplasmosis Parasite May Trigger Schizophrenia and Bipolar Disorders. <i>Science Daily</i> . Retrieved August 26, 2013, from <i>http://www.sciencedaily.com/releases/2009/03/090311085151.htm</i> .
468 469 470	50.	Michigan State University (2012, August 16). Common parasite may trigger suicide attempts: Inflammation from T. gondii produces brain-damaging metabolites. <i>Science Daily</i> . Retrieved August 26, 2013, from <i>http://www.sciencedaily.com</i> /releases/2012/08/120816170400.htm.
472	51.	Pittella, José EH. Neurocysticercosis. Brain Pathology 1997; 7: 681-93.
473 474 475 476 477	52.	Nascimento-Carvalho CM, Moreno-Carvalho OA. Neuroschistosomiasis due to <i>Schistosoma mansoni</i> : a review of pathogenesis, clinical syndromes and diagnostic approaches. Rev Inst Med Trop Sao Paulo 2005; 47(4):179-84.
478 479	53.	Houdon L, Flodrops H, Rocaboy M, Bintner <i>M</i> , Fériot JP, Tournebize P <i>et al.</i> Two Patients with Imported Acute Neuroschistosomiasis Due to <i>Schistosoma mansoni. J Travel Med</i> 2010; 17(4): 274–7.
480		