

Neurological Disease Surveillance in Cameroon, a Rural and Urban-Based Inout Patient Population Study

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Abstract: Background: There is a paucity of literature on the burden of neurological diseases in sub-Saharan Africa. Objective: To create a registry and surveillance of neurological diseases from urban and rural health centers in Cameroon. Methods: Retrospective review of medical records of inpatients and outpatients from two urban public hospitals in Douala and two rural health care centers, from 2013 to 2015 was conducted. In the urban areas, the diagnosis was made by a neurologist but this was not the case in the rural areas. The following variables were analyzed: demographics, medical center characteristics, presenting neurological complaint, medical history, neurological diagnosis, death and disability. Neurological diseases were classified according to ICD-10. Results: Out of 20,131 medical charts available (13% from the rural area), 4,187 cases (20.7%) with neurological complaints were identified and reviewed, mean age 48.67 ± 18.62 years, females 54.7%, 188 children (4.4%). The most frequent neurological complaints were: paresis/weakness (G.82, 25.2%) and headache (R.51, 22.0%). The most common concurrent medical history was hypertension (I10, 40.0%), and HIV (B20, 16.45%). The most common neurological diagnoses were cerebrovascular disease (G45, G46, 51.5%), and infection (B50, G00, G04, G06, 24.13%) in adults, and epilepsy (G40, 64.0%) in children. Death due to neurological cause was recorded in 428 patients (19.1%), and disability in 1,072 (57.2%). Neurodegenerative diseases were exclusively diagnosed in urban areas. Conclusion: Population aging and changes in the distribution of risk factors have accelerated the prevalence of non-communicable diseases such as cerebrovascular disease. However, additional work to characterize the nature of diagnosis, treatment and care is important to advance quality of care in the adult and pediatric neurological disorders. A health policy geared towards prevention and neurological training for health professionals is warranted.

Keywords: Neurological Disease, Surveillance, Cameroon

1. Introduction

Over the past 25 years, the burden of neurological disorders has increased substantially and at the present time represents the leading cause group of disability, and the second-leading cause group of mortality worldwide. [1] However, despite the high prevalence and disease burden of

neurological disorders, [2] most Sub-Saharan countries have less than one neurologist per one million people and lack medically trained personnel. Furthermore, over the last decades, population aging and changes in the distribution of risk factors have accelerated an epidemic of non-communicable diseases in many developing countries. [2, 3] In this regard, in developing countries the burden of

neurological diseases, especially neurodegenerative diseases, results from being both under-recognized and under-treated. [4, 5] Much recent neuro-epidemiological work in Africa has been carried out in urban areas, [6] but data regarding disability and death associated with neurological diseases is still scarce, especially for non-communicable diseases in the rural areas. [7, 8]

Epidemiology plays an important role in elucidating patterns in the occurrence of disease and determining risk factors and etiology. [9, 10] While obtaining information on the burden of neurological diseases by region or district is important so that interventions can be specifically targeted, there is also a need to establish a reliable database system to record the prevalence of neurological diseases in larger geographical areas like the Sub-Saharan African countries.

Cameroon is located in Central Africa with a population of over 22 million people (World Bank 2013) and life expectancy 54.59 years (World Bank 2012). It has been identified by the World Health Organization (WHO) as having critical shortage of health care workers and geographic distributional inequalities that are even more severe in rural areas. [11] Although there is a small body of literature on the hospital-based prevalence of neurological disorders in Cameroon, [12, 5] little is known about the prevalence of neurological disorders in the rural area. To address this lack of information, we conducted a collaborative project in Cameroon to create a registry and surveillance of neurological disorders including neurodevelopmental disorders, stroke, epilepsy, neurodegenerative diseases, neuromuscular diseases and central nervous system infections. Our objective was to provide reliable data of neurological disease surveillance in the context of the selected urban hospitals and rural clinics, thus encompassing both urban and rural areas.

2. Methods

Design: Retrospective chart review of neurological cases with a primary neurological complaint being the main reason for admission to the hospital/visit to the outpatient clinic, seen either in an inpatient setting in 2 urban public hospitals in Douala (Laquintinie hospital and General Hospital) or in an outpatient setting in 2 rural health care centers (Tiko and Mbouda), in Cameroon, from May 1st 2013 to May 1st 2015.

Start-up & Ethics: The investigators (EC and KK) travelled to Laquintinie Hospital, Douala, Cameroon in 2014, to invite physicians to participate in the registry of neurological diseases. During an one-day course, the basis of the neurological diseases registry, project methodology and local networks between neurologists and primary care doctors were discussed. A local neurologist (JD) was designated as the principal investigator in Cameroon. This project was approved by the Internal Review Board (IRB) at Douala Laquintinie Hospital (Cameroon) and at the principal investigator IRB (Hospital Universitario Burgos, Spain).

Procedure: The in/outpatient settings covered an urban population of 3,000,000 inhabitants served by 9 neurologists

in 2 hospitals (Douala and Yaoundé), and a rural population of 380, 276 inhabitants served by 20 doctors, mainly primary care. A senior medical student from Douala University Medical School was engaged (TN) to be the primary data collector for this project. Regular teleconferences between the investigators were performed to discuss data collection and analysis issues. All medical centers provided a registry of patients with neurological complaints (paper datasheet), and each individual medical chart of patients with neurological complaints was reviewed by the primary data collector. To avoid duplicity, each patient was accounted once with the primary neurological complaint.

Outcomes: Clinical, and socio demographic information was obtained from the medical charts. Neurological diagnosed were performed by 9 neurologists in the overwhelming majority of cases in the urban areas, and by internists, general practitioners and other medical specialties in the rural areas. Medical history, neurological complaints, and neurological disorders were coded according to the International Statistical Classification of Diseases-Related Health Problems 10th revision (ICD-10) by the primary data collector.[13] The following data were collected from the patients' records: age, sex, profession, residence (rural vs. urban), the main neurological complaint upon consultation/admission, medical history (tobacco and alcohol exposure, presence of arterial hypertension, diabetes mellitus, dyslipidemia, HIV infection and malaria), district population, number of neurologists, access to laboratory services, including microbiological testing, and Computerized Tomography scan, disability (ranging from independence to partial/full dependence, based on the medical chart notes and investigator criteria), and death. Pediatric cases were classified as infants (0-2 years), toddlers (3-6 years) and children/teenagers (7-16 years).

Internal validity of neurological diagnosis. Training for entered data accuracy in the database was established by comparing the ICD-10 codes based on the clinical information obtained from the medical chart, and entered in the database by the local neurologist (JD) from 11 medical charts (as the gold standard), to the ICD-10 codes entered in the database by the primary data collector. To certify the data collector, an inter-rater agreement with a Kappa coefficient > 0.8 was necessary. The inter-rater agreement was low in the first round (kappa coefficient of 0.5), but high in the second round (kappa coefficient of 0.9) using 11 different medical charts.

To establish the accuracy of neurological diagnoses provided by non-neurologist medical doctors involved in the care of the patients included in this registry, a clinical test including the clinical vignettes of 10 general neurology cases was conducted. Three internists from the urban medical centers, and 4 primary care doctors from the rural medical centers, completed the clinical test. There were a mean number of 8.6 ± 0.5 correct answers from physicians in urban centers, and 7.4 ± 0.8 in rural centers.

Statistical analysis: Statistical analyses were performed using SPSS Version 21.0 (SPSS, Inc., Chicago, IL).

CNS tumor (3.8%), and epilepsy (2.4%). Disability data was exclusively available from urban cases. Among the 1803 survivors, dependence was observed in 810 patients (44.9%), 361 (44.5%) patients had mild-moderate dependence (need for help with Activities of Daily Life-dressing, feeding, bathing-but could do most things by themselves), and 275 patients (33.9%) patients were fully dependent.

Table 1. Demographics and medical history characteristics in Urban vs. Rural patients.

Frequency (%)	Urban N=3955	Rural N=231
Gender		
Female	2157 (54.5)	134 (58.3)
Missing information	3 (0.1)	1 (0.4)
Marital status		
Single	801 (20.3)	29 (12.6)
Married	1531 (38.7)	173 (74.9)
Widowed	345 (8.7)	29 (12.6)
Missing information	1278 (32.3)	0 (0)
Working status		
Active worker	1833 (46.3)	16 (6.9)
Retired due to age	389 (9.8)	2 (0.9)
Housewife	890 (22.5)	4 (1.7)
Unemployed	605 (15.3)	4 (1.7)
Missing information	238 (6)	205 (88.7)
Education		
Illiteracy, low level	1741 (44.0)	-
Middle-high school	939 (23.7)	1 (0.4)
University	595 (15.0)	-
Missing information	680 (17.2)	230 (99.6)
Disability		
Yes	1052 (26.2)	21 (4.1)
Missing information	2146 (54.3)	163 (70.6)
Death		
Yes	385 (9.7)	43 (18.6)
Missing information	2146 (54.3)	0
Leading Diagnosis*		
Cerebrovascular disease	1254 (31.9)	89 (39.3)
Mononeuritis	616 (15.6)	17 (7.5)
Epilepsy	544 (13.8)	18 (8.0)
Headache	501 (12.7)	5 (2.2)
Infection	478 (12.1)	62 (27.4)
Tumor	126 (3.2)	5 (2.2)

*Diagnosis frequency (%) is established based on available data. Cerebrovascular disease includes the ICD-10 codes G45,G46; Mononeuritis includes the ICD-10 codes G56,G57; Infection includes the ICD-10 codes B50,B99,G00,G04;G06; Headache includes the ICD-10 code G43; Epilepsy include the ICD-10 code G40; Tumor include the ICD-10 codes.

Table 2. Leading diagnosis, medical history, disability and deaths in adults.

Diagnosis			Medical history				Disability (yes)			Death (yes)		
Disease	Total sample	Frequency	Total sample	ICD-10 code	n	Frequency	Total sample	n	Frequency	Total sample	n	Frequency
Cerebrovas. disease	1338	34%	1286	I10	825	64%	851	651	76%	1151	239	21%
				E11	204	16%						
				F10	164	13%						
				G45.1	123	10%						
				I21	109	8%						
				F17	88	7%						
				B20	38	3%						
				E78	34	3%						

Diagnosis		Medical history					Disability (yes)			Death (yes)		
Disease	Total sample	Frequency	Total sample	ICD-10 code	n	Frequency	Total sample	n	Frequency	Total sample	n	Frequency
Mononeuritis	611	15%	525	A15	11	1%	48	1	2%	63	3	5%
				G40.1	5	0%						
				A16	1	0%						
				I10	94	18%						
				F10	29	6%						
				E11	26	5%						
				F17	9	2%						
				G45.1	9	2%						
				B20	6	1%						
				E78	3	1%						
				A15	2	0%						
				F12	1	0%						
				I21	1	0%						
Infection	601	15%	598	B20	350	59%	460	105	23%	535	152	28%
				I10	51	9%						
				A15	42	7%						
				F10	31	5%						
				E11	22	4%						
				F17	17	3%						
				G45.1	7	1%						
				G40.1	5	1%						
				F12	3	1%						
				E78	2	0%						
				B21	1	0%						
				I21	1	0%						
				I10	37	11%						
Headache	494	12%	351	F10	15	4%	42	0	0%	46	1	2%
				E11	10	3%						
				F17	9	3%						
				B20	6	2%						
				G45.1	5	1%						
				I21	2	1%						
				E78	1	0%						
				I10	102	26%						
				G45.1	51	13%						
				G40.1	37	9%						
				F10	22	6%						
				E11	20	5%						
				B20	16	4%						
Epilepsy	442	11%	397	F17	12	3%	217	37	17%	239	17	7%
				I21	5	1%						
				A15	4	1%						
				E78	4	1%						
				I10	19	16%						
				F10	11	9%						
				B20	8	7%						
				E11	8	7%						
				F17	7	6%						
				A15	3	3%						
				G45.1	1	1%						
Tumor	125	3%	120				80	36	45%	85	16	19%

Overall data was available in 3992/3998 (99.8%). Medical history was available in 3151/3955 (79.5%) urban cases, and 103/231 (44.5%) rural cases. Cerebrovasc. Disease (Cerebrovascular disease) includes the ICD-10 codes G45,G46; Mononeuritis include the ICD-10 codes G56,G57; Infection includes the ICD-10 codes B50,B99,G00,G04;G06; Headache includes the ICD-10 code G43; Epilepsy include the ICD-10 code G40; Tumor include the ICD-10 codes. Medical history include the ICD-10 codes: I10 Arterial hypertension; F83 birth injury; B20,B21 Human immunodeficiency virus; A15,A16,A17 Tuberculosis; B50 Malaria; F17 Tobacco, F10 Alcohol, F12, F14 other drugs, E10,E11 Diabetes Mellitus; E78 Dyslipemia, I21 Ischemic cardiac disease.

Table 3. Leading pediatric neurological complaints and diagnosis.

N (%)	Total sample N=188	Infant 0-2 years N=13	Toddler 3-6 years N=38	Children 7-16 years N=137
Female	104 (55)	6 (46)	21 (55)	77 (56)
Outpatient	145 (77)	11 (85)	38 (100)	96 (70)
Location				
Urban	180 (96)	13 (100)	38 (100)	129 (94)
Rural	8 (4)	0 (0)	0 (0)	8 (6)
Primary Neurological complaint				

N (%)	Total sample N=188	Infant 0-2 years N=13	Toddler 3-6 years N=38	Children 7-16 years N=137
Fever of unknown origin	18 (13)	1 (8)	1 (3)	16 (16)
Seizure	73 (50)	4 (33)	19 (63)	50 (49)
Headache	40 (27)	1 (8)	4 (13)	35 (34)
Mental status change	38 (26)	1 (8)	2 (7)	35 (34)
Diagnosis				
Epilepsy	120 (64)	5 (42)	33 (87)	82 (60)
Headache	12 (6)	0 (0)	0 (0)	12 (9)
Cerebral palsy unspecified	9 (5)	4 (31)	1 (3)	4 (3)
Infection	10 (19)	2 (17)	3 (8)	5 (4)
Tumor	7 (4)	0 (0)	1 (3)	5 (4)
Tremor/Myoclonus	5 (3)	1 (8)	0 (0)	4 (3)

Epilepsy includes the ICD-10 code G40; Fever of unknown origin include the ICD-10 code R50; Seizure the ICD-10 code R56; Headache the ICD-10 code R51, and Mental status change the ICD-10 code R40; Headache includes the ICD-10 code G43; Cerebral palsy include the ICD-10 code G80.9; Infection include the ICD-10 codes G00, G04; Tumor includes the ICD-10 codes C70,C71,C72; Tremor/Myoclonus includes the ICD-10 code G25.

4. Discussion

Our study is one of the largest retrospective reviews measuring the frequency of all serious neurological conditions, requiring either an outpatient visit or hospitalization in urban and rural areas of Sub Saharan Africa, including pediatric and adult cases. The most common diagnoses in both the inpatient and the outpatient samples were, similar to other studies involving Cameroon, with non-infectious neurological diseases being more common than infectious diseases. [8, 14, 15] In our sample we found that the major neurological disease resulting in hospitalization was cerebrovascular disease, accounting for a little more than half the cases, followed by CNS infections, accounting for almost a quarter of the cases. Among admissions for neurological reasons, 19.0% resulted in death and among the survivors, 44.5% of patients were mildly to moderately dependent and 33.9% were fully dependent. Interestingly, we have previously published a separate analysis for disability and death, and found that higher educational level was associated with decreased likelihood of disability. [16] To explain this finding several hypotheses can be taken into account such as the fact that higher educational level is usually associated with higher socioeconomic status and healthier life style, which may lead to a decrease cerebrovascular risk factors, or earlier recognition of stroke symptomatology and thus earlier intervention. However, this information cannot be extrapolated to the overall community due to selection bias with a large proportion of disability missing data especially from the rural area, lack of information regarding long-term disability, and the limited reliability of the operative definition for disability used in this study.

Traditionally, the neurological complications of infectious diseases (HIV infection, tuberculosis and malaria) have been considered to account for a significant portion of the neurological diseases found in the countries of Sub Saharan Africa.[17-19] Contrary to this belief, cerebrovascular disease accounted for 51.0% of neurological admissions in this urban sample, 55.3% of the neurological mortality and 79.7% of the neurological disability among the survivors. Other studies in Sub-Saharan Africa have reported crude

yearly stroke incidence rates of 94.5-107.9/100,000, associated with high mortality rates. [20, 21] Therefore, cerebrovascular disease represents the leading cause of mortality and disability. Cerebrovascular disease though, in the ICD-10 classification has been classified with the cardiovascular diseases in the chapter on circulatory disorders. This anomaly has made statistics on neurologic mortality and morbidity misleading. [22] A detailed analysis about the neurological diseases and risk factors associated with death and disability in this cohort has been published elsewhere. [16].

Overall, the body of literature on neurodegenerative disorders in Sub-Saharan countries is large with regards to dementia and HIV-related neurocognitive disorders, [23] but there is limited data on other neurodegenerative disorders. [24] Our results are in agreement with previous studies conducted in Africa. The prevalence of neurological disorders in those aged 70 years and over in a rural community in Tanzania, was 154.1 per 1000 (95% CI 139.2-169.1), and the most common neurological disorders were tremor (48.2), headache (41.8), stroke (23.0), peripheral polyneuropathy (18.6), upper limb mononeuropathy (6.5) and parkinsonism (5.9). [8] Moreover, based on recent estimates, it is predicted that by 2050 there will be 139 million people aged 60 and older living in Sub-Saharan Africa, [25, 26] with a subsequent increase in the prevalence and burden of neurodegenerative disorders. [25] Kuate et al, [5] also reported that the most frequent complaints and neurological diseases were low back pain (19.7%), tremor and movement disorders (13.8%) and gait disorders (4.3%) in a hospital-based sample in Yaoundé-Cameroon. Of note, neurodegenerative disorders in our sample were exclusively diagnosed in the urban areas, indicating these diseases are likely underdiagnosed and undertreated, especially in the rural areas.

Pediatric patients with neurological disorders were more often diagnosed in urban health care facilities, with the most common complaint being seizure and the most common neurological disorder being epilepsy. Previous work on epilepsy in sub-Saharan Africa has shown that many cases are severe and frequently not adequately treated. [27, 6] The prevalence of epilepsy, estimated from door to-door studies in Sub-Saharan countries, is almost double than in Asia,

Europe and North America. [28] The most commonly cited risk factors for epilepsy are birth trauma, CNS infections, tumors and traumatic brain injury. [29] These were identified up to 6% in our sample. Additional work to better characterize the nature of diagnosis and treatment in this part of the world is important in order to improve quality of care in this vulnerable population.

The main limitation of our study is that the conclusions apply to a selected in-outpatient sample and cannot be directly compared to population-based studies. Likewise, the frequency estimates obtained from these specific in-outpatient settings can be limited by several factors including selection bias because the majority of patients were inpatients, which skews the data toward stroke and underestimates degenerative diseases and dementia. Moreover, data quality in terms of recognition of neurological complaints of hospital registries, patient's access, type of health care facilities, and doctor's expertise, restricting the extrapolation of our prevalence estimates to the greater community or the whole country. Furthermore, although the accuracy of our data was examined, especially the accuracy of neurological disorders diagnosis established by non-neurologists physicians in a limited sample of doctors involved in the care of this cohort, we do not have data in terms of their specific expertise in highly specialized neurological diseases, resulting likely in diagnosis bias. Moreover, a selection bias due to small representation of the rural population, especially for the pediatric population should be taken into account.

However, despite all of these limitations, due to the lack of epidemiological data in Sub-Saharan countries, this type of publication raises awareness in neurological conditions, especially cerebrovascular disease as the leading cause of mortality and disability in this cohort. Furthermore, the information is especially valuable given that the overwhelming majority of patients in the urban areas were diagnosed by a neurologist. Finally, it is important to note that our registry included adult and pediatric population, both an urban and rural sample, which have been rarely done in previous studies. [5, 6] Accurate frequency of neurological conditions is needed to address the effects of the demographic shift that is taking place in many regions of the world, including Sub-Saharan countries to provide.

5. Conclusion

There is no doubt, that improving access to patients with neurological disorders must be coupled with increased education to raise awareness of these diseases including diagnosis and treatment, especially in the rural areas. Furthermore, high-quality neurological disorder training for health care providers is essential to tackle this increasing need. Our study findings have important health service implications including adequate planning by governments and other health care providers for prevention and management of neurological conditions.

Authors' Contribution

1. Research Project: A= Conception, B= Organization, C= Execution
2. Statistical Analysis: A= Design, B= Execution, C= Review and Critique
3. Manuscript Preparation: A= Writing the First Draft, B= Review and Critique.

JD: 1A, 1B, 1C, 2C, 3A, 3B

YNM: 1B, 1C, 2C, 3B

TN: 1C, 3B

CK: 1C, 2C, 3B

KK: 1A, 1B, 2C, 3B

BO: 2C, 3B

HS: 1B, 2C, 3B

SC: 2C, 3B

AFS: 2C, 3B

EC: 1A, 1B, 2B, 2C, 3A, 3B

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Conflict of Interest

The authors declare that they have no competing interests.

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