

Global Epidemiology of Stroke with Special Reference to Sub - Saharan Africa

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Learning Objectives

- In this session we will:
- **Remind** ourselves of the recent definition of stroke.
- Appreciate the trend of global and African epidemiology of stroke
- Understand the drivers of the changing stroke epidemiology
- Be **inspired** to identify where/how we can fit in, to tackle the growing burden of stroke in Africa.

Outline

- Stroke definitions
- Epidemiology of Stroke _ Global
- Epidemiology of Stroke _ Africa
- Risk factors for Stroke
- Drivers of Changing Stroke Epidemiology
- Is Genetics important to Stroke in Africa?
- Tackling the growing Stroke burden in Africa
- Conclusion

Stroke Definition

(Before Now)

- Global or focal disturbance of cerebral function resulting only from a vascular cause with dysfunction lasting more than 24 hours or resulting in death.
- Included in this definition are: cerebral infarction, intracerebral haemorrhage, subarachnoid haemorrhage

WHO, 1988.

Revised Definition

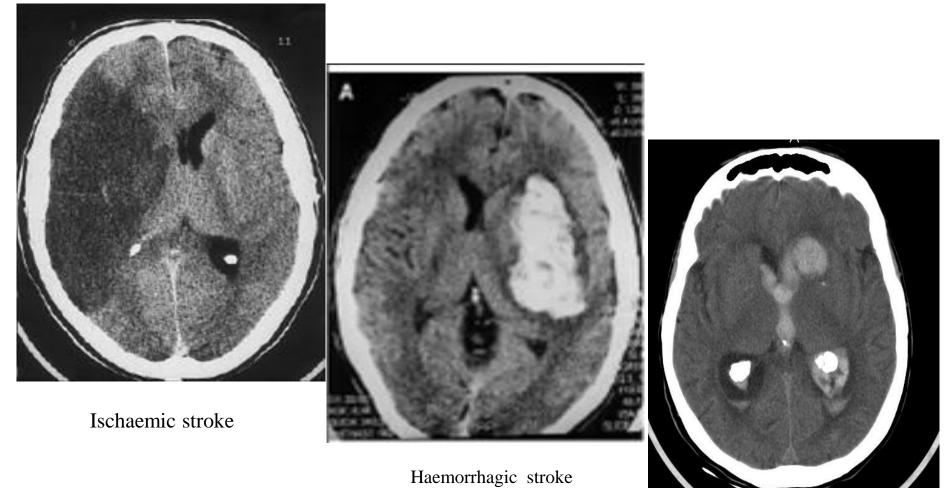
- Stroke is defined as a sudden global or focal neurological deficit resulting from spontaneous hemorrhage or infarction of the central nervous system with objective evidence of infarction/hemorrhage irrespective of duration of clinical symptoms. AHA/ASA 2009.
 - CT/MRI necessary to increase diagnostic accuracy.

Current concepts

- TIA: New definition classifies TIA as a transient episode of neurologic dysfunction caused by focal brain, spinal cord, or retinal ischemia, without evidence of acute infarction. No objective evidence of acute infarction in the affected region of brain or retina. Sorensen et al Neuroimaging Clin N Am. 2011; 21(2): 303–313.
- Stroke : Stemming from the new tissue-based definition of transient ischemic attacks (TIA), stroke can be defined as a sudden global or focal neurological deficit resulting from spontaneous hemorrhage or infarction of the central nervous system with objective evidence of infarction irrespective of duration of clinical symptoms. CT/MRI necessary to increase diagnostic accuracy. Sacco RL et al Stroke. 2013;44:00-00.

(cf Angina pectoris vs Myocardial infarction).

Major Stroke Types



Subarachnoid haemorrhage

Trends in Global Stroke Epidemiology

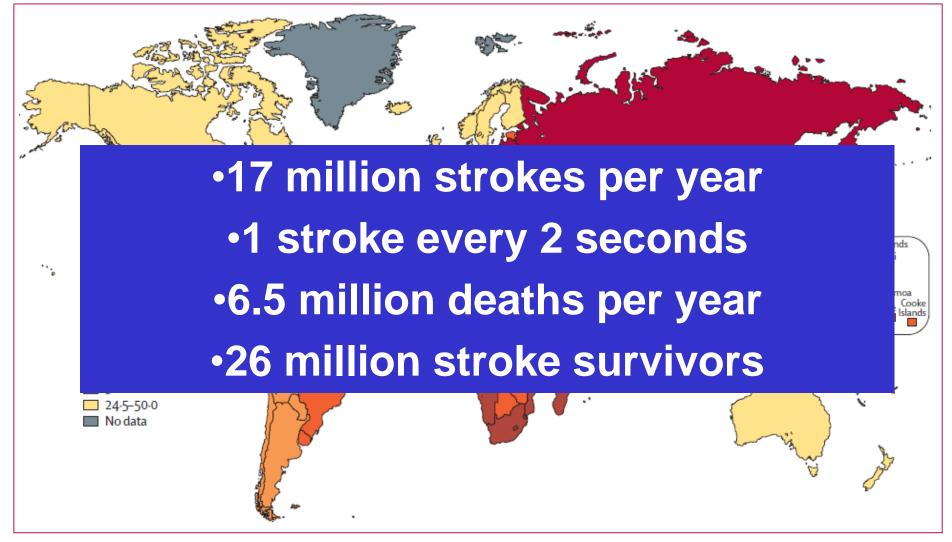
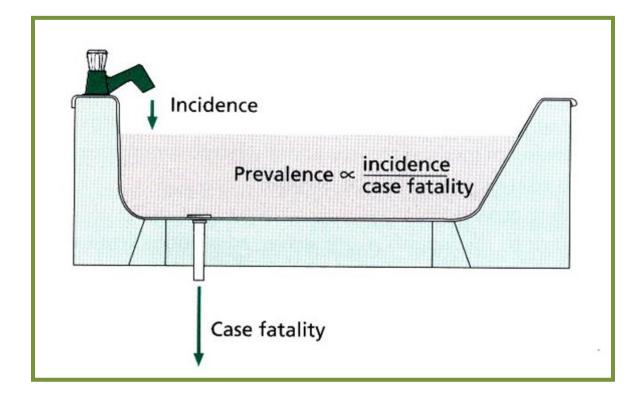


Figure 1: Age-adjusted and sex-adjusted stroke mortality rates Rates are highest in eastern Europe, north Asia, central Africa, and the south Pacific.

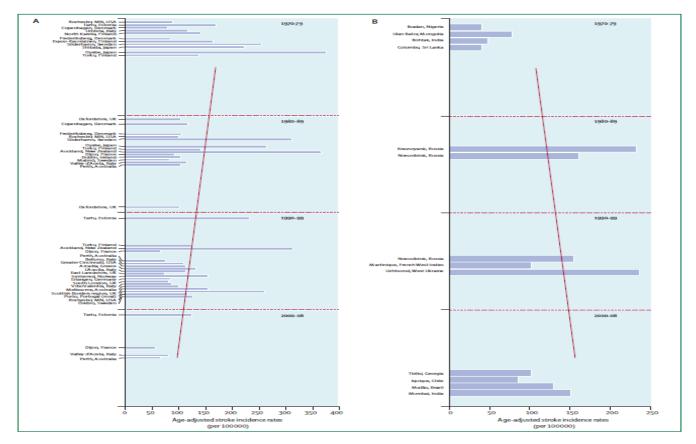
The burden of stroke in population



Worldwide stroke incidence and early case fatality reported in 56 population-based studies: a systematic review



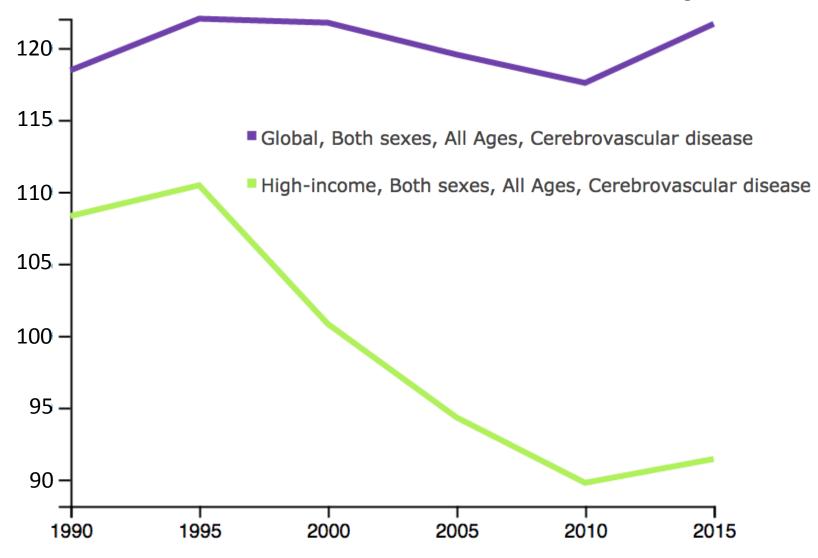
Lancet Neural 2009; 8: 355-69



Stroke incidence rates over four decades (1970- 2010), with a 42% ↓ in HIC and ↑ 100% LMIC . In 2000–08, overall stroke incidence rates in LMIC, for the first time, exceeded HIC by 20% . - Feign et al, 2009

Global trends in stroke incidence rates per 100,000 person-years (1990-2015)

-Feigin et al, 2016



Global and regional burden of first-ever ischaemic and haemorrhagic stroke during 1990–2010: findings from the Global Burden of Disease Study 2010

Rita V Krishnamurthi, Valery L Feigin, Mohammad H Forouzanfar, George A Mensah, Myles Connor, Derrick A Bennett, Andrew E Moran, Ralph L Sacco, Laurie M Anderson, Thomas Truelsen, Martin O'Donnell, Narayanaswamy Venketasubramanian, Suzanne Barker-Collo, Carlene M M Lawes, Wenzhi Wang, Yukito Shinohara, Emma Witt, Majid Ezzati, Mohsen Naghavi, Christopher Murray, on behalf of the Global Burden of Diseases, Injuries, and Risk Factors Study 2010 (GBD 2010) and the GBD Stroke Experts Group*

Krishnamurthi RV Lancet Glob Health 2013;1: e259-81

Global and regional burden of stroke during 1990–2010: findings from the Global Burden of Disease Study 2010

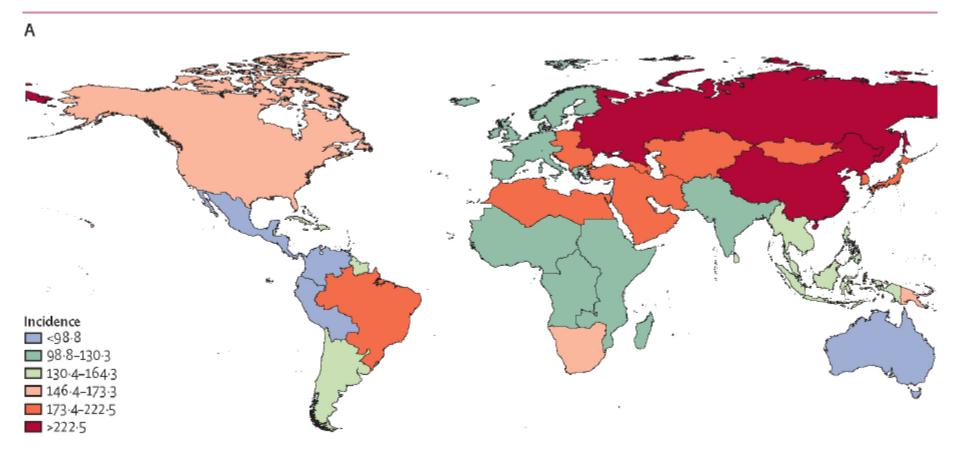
Valery L Feigin, Mohammad H Forouzanfar, Rita Krishnamurthi, George A Mensah, Myles Connor, Derrick A Bennett, Andrew E Moran, Ralph L Sacco, Laurie Anderson, Thomas Truelsen, Martin O'Donnell, Narayanaswamy Venketasubramanian, Suzanne Barker-Collo, Carlene M M Lawes, Wenzhi Wang, Yukito Shinohara, Emma Witt, Majid Ezzati, Mohsen Naghavi, Christopher Murray, on behalf of the Global Burden of Diseases, Injuries, and Risk Factors Study 2010 (GBD 2010) and the GBD Stroke Experts Group*

Feigin VL Lancet. 2014 ;383:245-54.

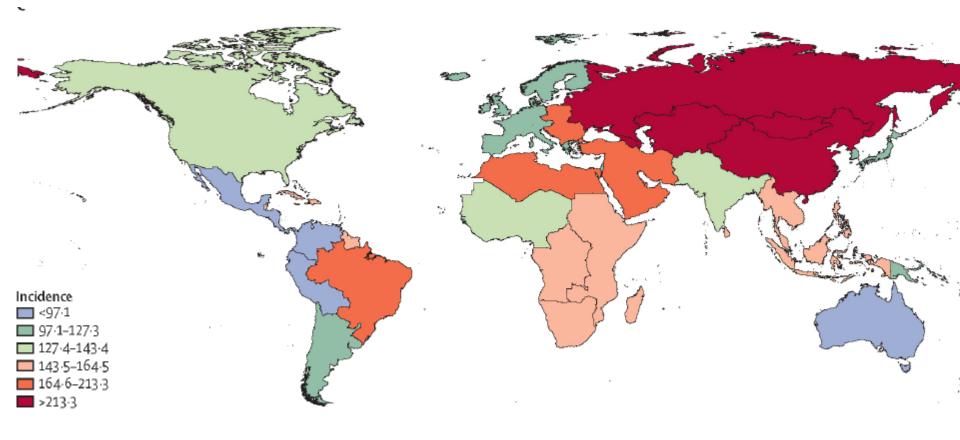
The global burden of disease used multi-state models implemented in the software program DisMod III. Neuroepidemiology 2012;38:30–40, Mathers CD British Medical Bulletin 2009; 92: 7–32



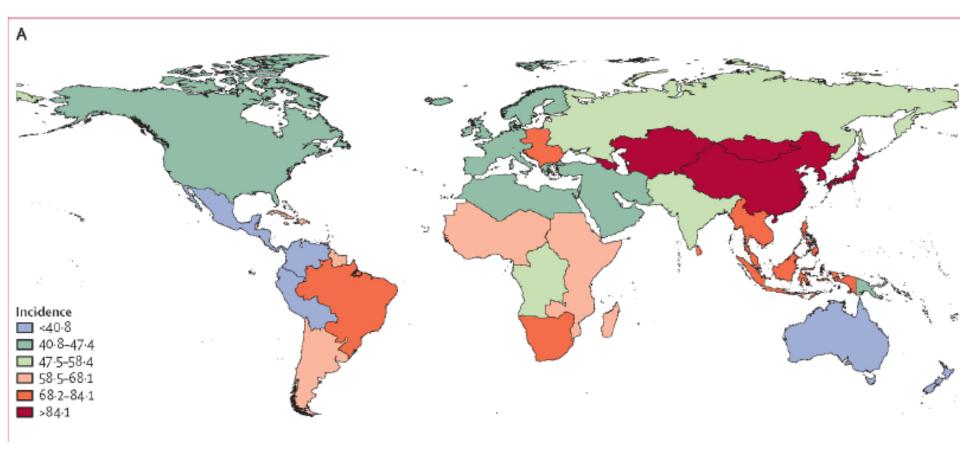




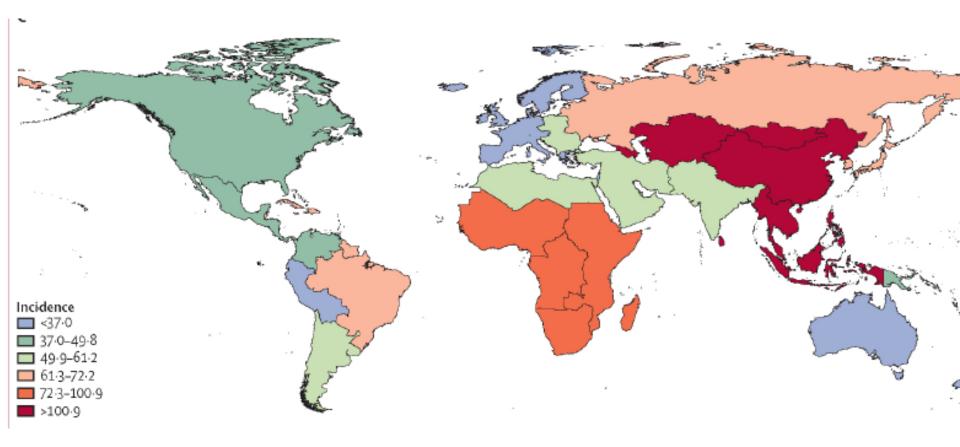
Age standardized Incidence of Ischemic Stroke 1990



Age standardized Incidence of Ischemic Stroke 2010

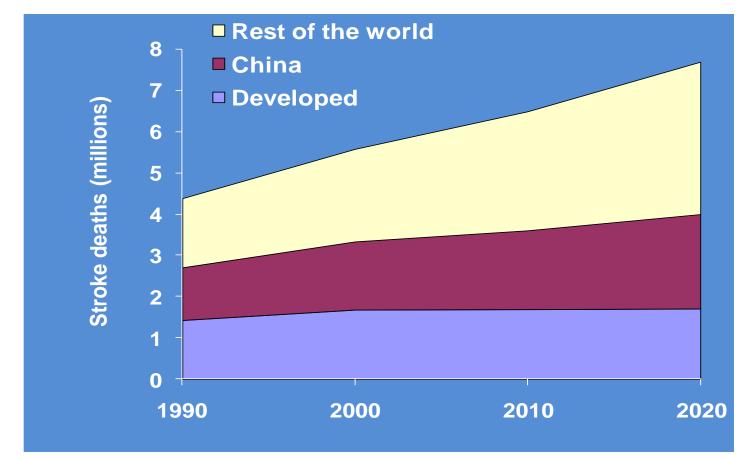


Age standardized Incidence of Hemorrhagic Stroke 1990



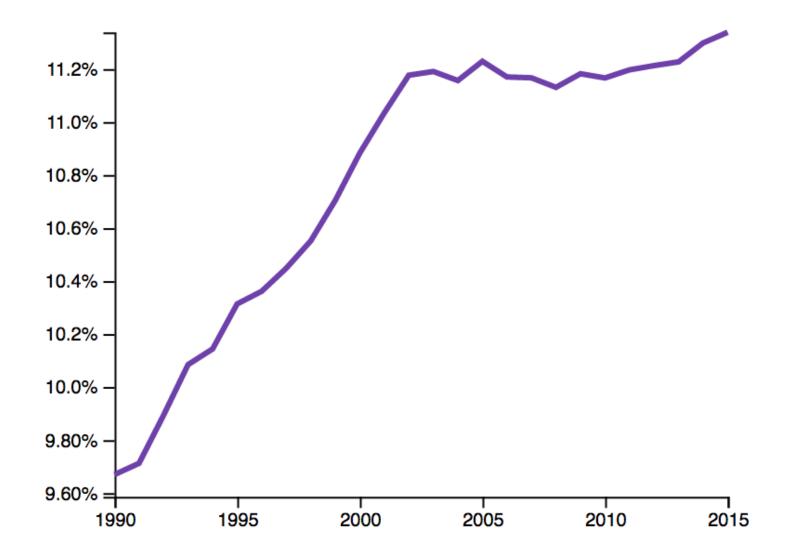
Age standardized Incidence of Hemorrhagic Stroke 2010

Global stroke deaths 1990-2020



Reference: WHO. Atlas of Heart Disease and Stroke. www.WHO.int 2004

Proportional contribution of deaths from stroke to all causes of deaths in the world (1990-2015)



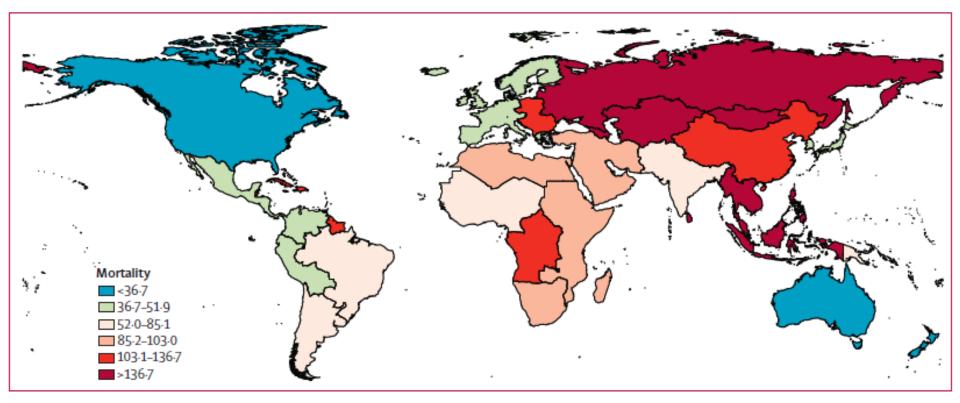
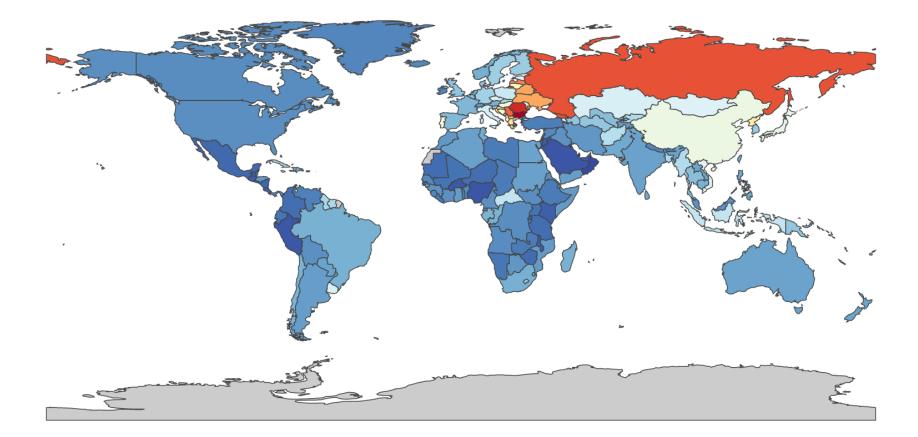


Figure 2: Age-standardised stroke mortality per 100 000 people for 2010

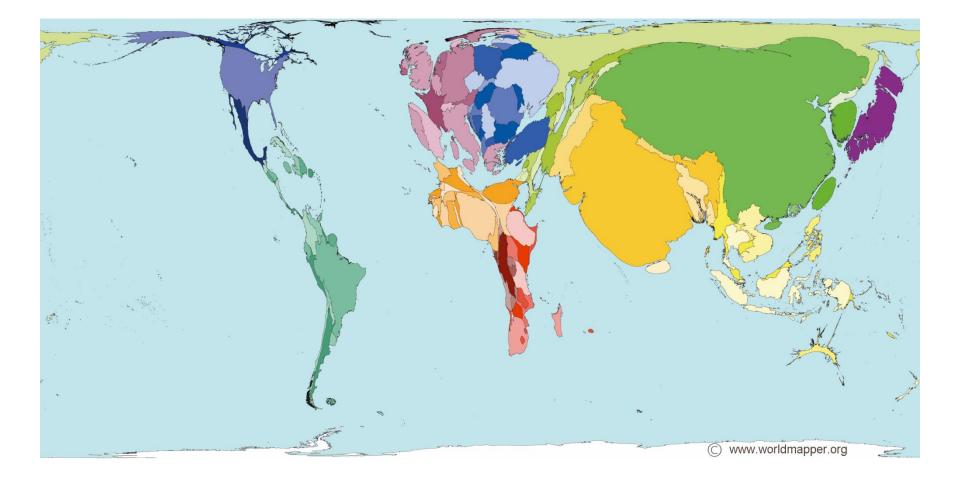
Feigin VL Lancet. 2014 ;383:245-54.

Geographical variations in stroke mortality rates in 2015





Worldwide burden of all stroke deaths

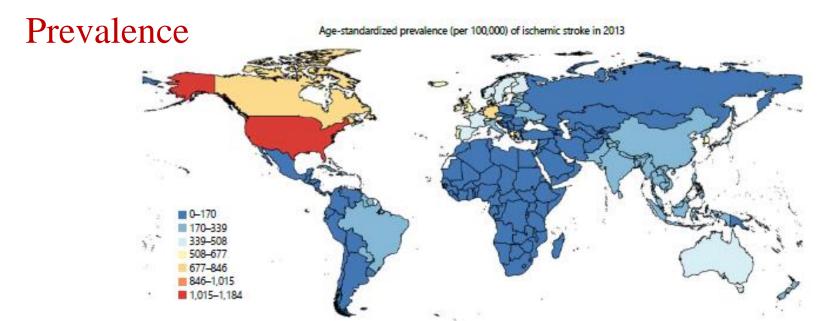


Size of country proportional to the proportion of worldwide stroke deaths in that country

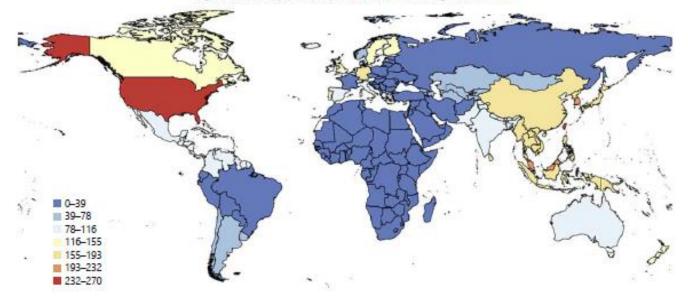
The reality of health care in areas of stroke burden



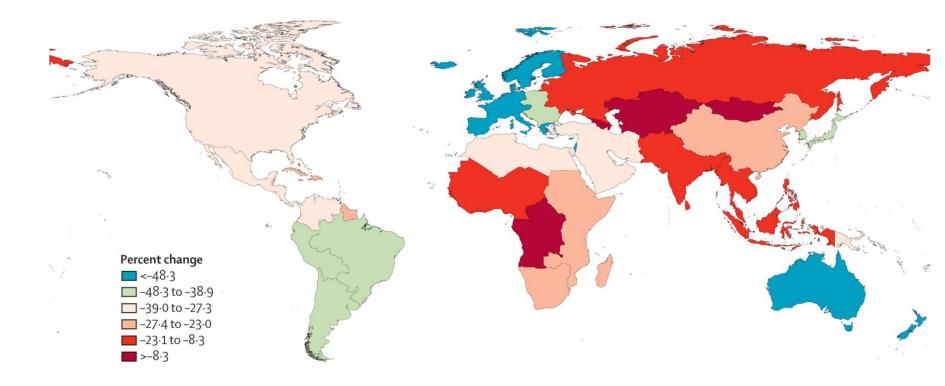




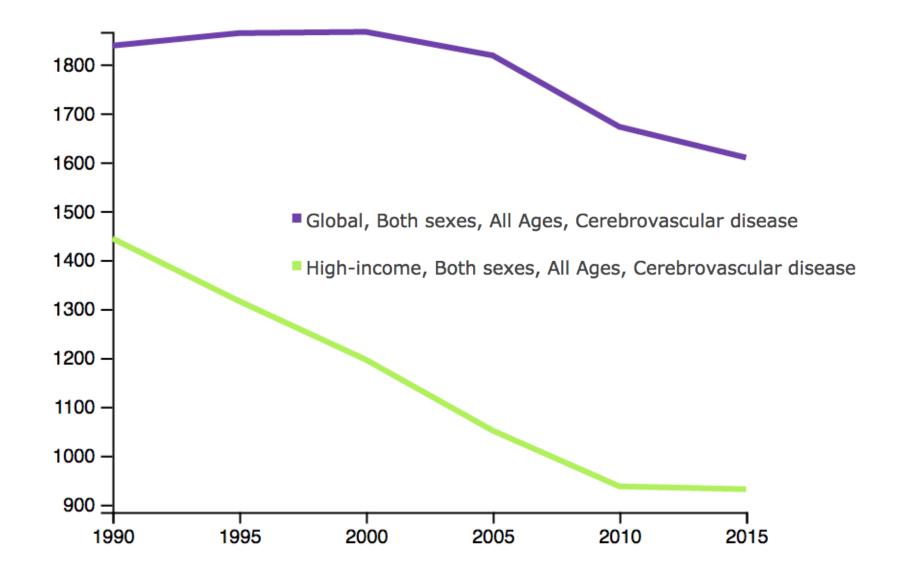
Age-standardized prevalence (per 100,000) of hemorrhagic stroke in 2013



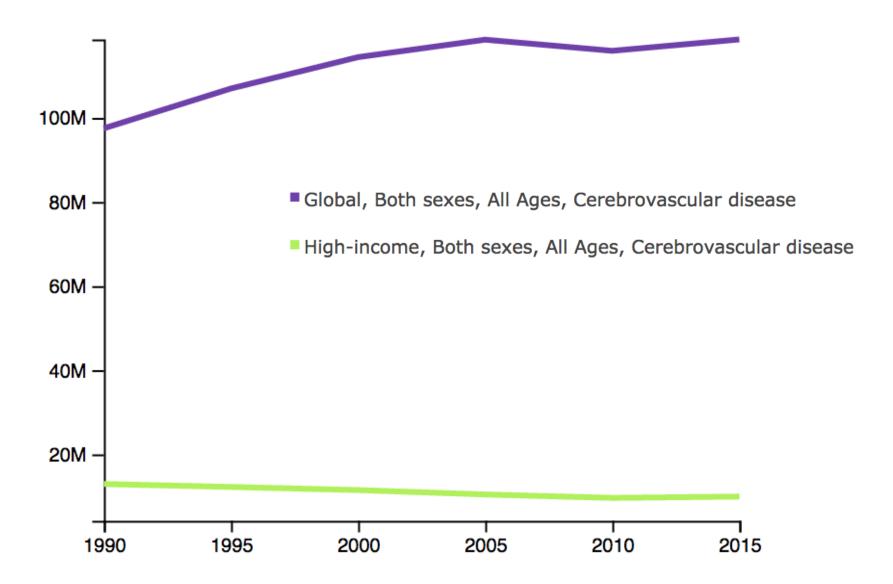
Feigin et al, 2015



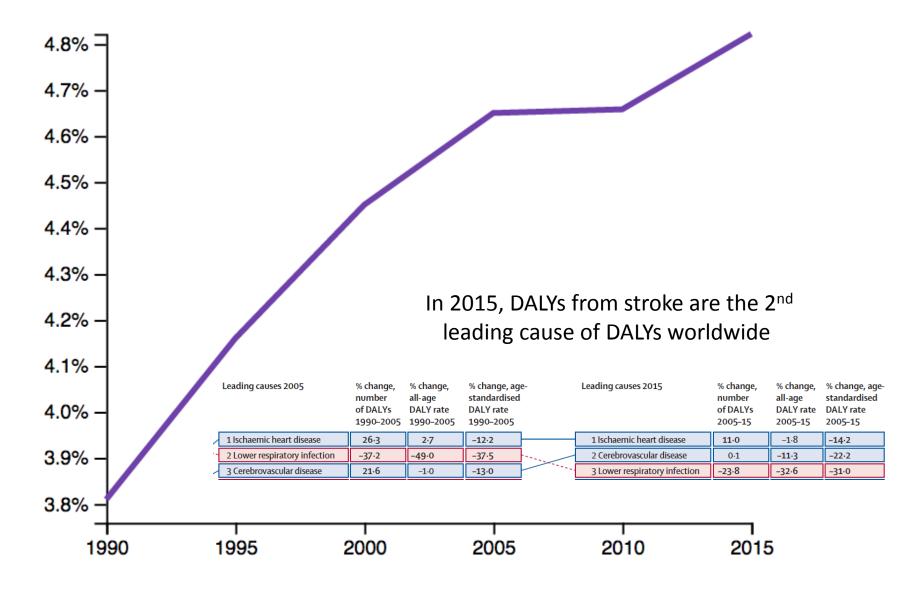
Global trends in stroke DALYs rates per 100,000 person-years (1990-2015)



Absolute number of DALYs from stroke in the world (1990-2015)



Proportional contribution of DALYs from stroke to DALYs from all causes in the world (1990-2015)

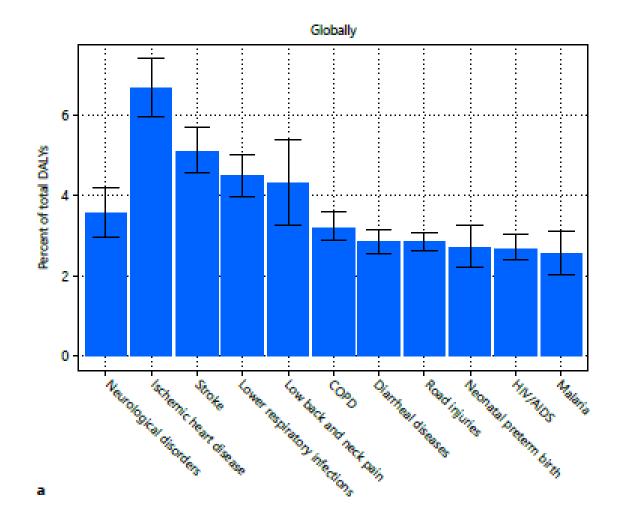


Estimated number of disability-adjusted life-years (DALYs) lost to stroke in 2020

Developed world: 10 million Developing world: 52 million

Total 62 million

Source: Murray and Lopez, Lancet 1997



Epidemiology of Stroke in SSA

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	Journal Code	Manuscript No.		No. of pages: 7	PE: Saranya Shree

CLINICAL INVESTIGATIONS

Contribution of Noncommunicable Diseases to Medical Admissions of Elderly Adults in Africa: A Prospective, Cross-Sectional Study in Nigeria, Sudan, and Tanzania

Rufus O. Akinyemi, PhD, *[†] Isameldin M. H. Izzeldin, MBBS,[‡] Catherine Dotchin, MD,^{†§} William K. Gray, PhD,[§] Olaleye Adeniji, MBBS, * Osheik A. Seidi, MBBS,[‡] Josephine J. Mwakisambwe, MD,[¶] Carl J. Mhina, MD,[¶] Florence Mutesi, MD,[¶] Helen Z. Msechu, MD,[¶] Kien A. Mteta, MD,[¶] Mayada A. M. Ahmed, MBBS,[‡] Shahd H. M. Hamid, MBBS,[‡] Nazik A. A. Abuelgasim, MBBS,[‡] Sumia A. A. Mohamed, MBBS,[‡] Ashraf Y. O. Mohamed, MBBS,[‡] Fidelis Adesina, MBChB,^{*} Mohammed Hamzat, MBBS,^{*} Taiwo Olunuga, MBChB,^{*} Venance P. Maro, MMed,[¶] and Richard Walker, MD^{†§}

RESULTS: In Africa, noncommunicable diseases (NCDs) accounted for 80.4% (n = 703) of admissions (n = 874), and tuberculosis, malaria, and the human immunodeficiency virus and acquired immunodeficiency syndrome accounted for 4.5% (n = 39). Stroke (n = 213, 24.4%) was the most common reason for admission, followed by cardiac or circulatory dysfunction (n = 155, 17.7%). Rates of hypertension were remarkably similar in the United Kingdom (45.8%) and Africa (40.0%).

Stroke Literacy among Africans

European Journal of Neurology 2009

doi:10.1111/j.1468-1331.2009.02666.x

Knowledge and perception of stroke amongst hospital workers in an African community

R. O. Akinyemi^{a,b}, O. S. Ogah^a, R. F. Ogundipe^a, O. A. Oyesola^c, A. A. Oyadoke^c, M. O. Ogunlana^d, F. M. Otubogun^a, T. F. Odeyinka^a, B. S. Alabi^e, J. O. Akinyemi^f, J. K. Osinfade^a and R. N. Kalaria^b

^aDepartment of Medicine, Federal Medical Centre, Abeokuta, Nigeria; ^bInstitute for Ageing and Health, Newcastle General Hospital, Newcastle upon Tyne, UK; ^cDepartment of Community Medicine and Primary Care, Federal Medical Centre, Abeokuta, Nigeria;

About 30% of workers in an hospital (majorly non – clinical) could not identify the brain as the organ affected in stroke
 90% correctly identified hypertension as a risk factor
 14% considered evil spirit/witchcraft as cause of stroke
 spiritual healing was most preferred by 13%

S27

The burden of stroke in Africa: a glance at the present and a glimpse into the future

Mayowa O Owolabi, Sally Akarolo-Anthony, Rufus Akinyemi, Donna Arnett, Mulugeta Gebregziabher, Carolyn Jenkins, Hemant Tiwari, Oyedunni Arulogun, Albert Akpalu, Fred Stephen Sarfo, Reginald Obiako, Lukman Owolabi, Kwamena Sagoe, Sylvia Melikam, Abiodun M Adeoye, Daniel Lackland, Bruce Ovbiagele, as members of the H3Africa Consortium

Abstract

Objective: Information on the current burden of stroke in Africa is limited. The aim of this review was to comprehensively examine the current and projected burden of stroke in Africa. **Methods:** We systematically reviewed the available literature (PubMed and AJOL) from January 1960 and June 2014 on stroke in Africa. Percentage change in age-adjusted incidence velocity, ischaemic stroke proportion, mean age and case fatality compared to high-income countries remain unknown.

Conclusions: While the available study data and evidence are limited, the burden of stroke in Africa appears to be increasing.

Stroke Incidence in Africa

- Hospital-based may show trends but not representative
- Community based, door-to-door, multiple case ascertainment methods, including verbal autopsies, over 3 years, age- standardized preferred. Owolabi MO West Indian Medical Journal. 2011; 60:412-421
- 315.9 per 100 000 (281.6-352.3) in Dar-es-Salaam (urban) Tanzania in 2006 Walker R Lancet Neurol 2010; 9: 786-92
- o 108·6 per 100 000 (95% CI 89·0−130·9) in Hai (rural) Tanzania in 2006
- 58.0 per 100 000 in Lagos, 2007. ?under-reporting (surveillance for 1 year, verbal autopsies not utilized, prehospitalization deaths not captured, patients who presented in hospitals outside the community not captured). Danesi MA J Neurol Sci. 2013;331:43-7.

Table 1. Studies reporting crude incidence of stroke in Africa									
			Crude incidence per 100 000 per year						
Year	Country/location/setting	Author	Overall	Male	Female	Age			
	Hospital-based								
1984	Libya, Benghazi, urban	Ashok ²⁸	63	69	58	15+			
1985	South Africa: Atteridgeville and Mamelodi, suburban areas of Pretoria, urban	Rosman ²⁹	101	108	93	20+			
1991	Zimbabwe, Harare, urban	Matenga ¹⁸	31	30	32	All			
1993	Libya, Benghazi, urban	El Zunni ³⁰	48	52	42	15+			
2006	Mozambique, Maputo, urban	Damasceno ¹⁹	149	174	128	15+			
	Population/community-based								
1975	Nigeria, Ibadan, urban	Osuntokun ²⁰	26	25	13	All			
1993	Egypt, Sohag, mixed*	Kandil ³¹	180	100	85	All			
1993	Egypt, Sohag, urban	Kandil ³¹	150	90	53	All			
1993	Egypt, Sohag, rural	Kandil ³¹	210	97	119	All			
2006	Tanzania, Hai, rural	Walker ³²	95	107	77	All			
2006	Tanzania, Dares Salaam, urban	Walker ³²	108	115	100	All			
2007	Nigeria, Lagos, urban	Danesi ²⁴	25	28	21	All			
2007	Egypt, Al-Kharga, mixed*	Farghaly ²²	250	270	230	All			
2007	Egypt, Al-Kharga, rural	Farghaly ²²	230	250	220	All			
2007	Egypt, Al-Kharga, urban	Farghaly ²²	260	280	240	All			
2012	Egypt,Al Quseir, urban	El Tallawy ²¹	181	212	150	20+			
*Combined rates including both rural and urban communities.									

Stroke mortality in SSA

- In hospital based studies, 1- month case fatality rate of up 50% (ICH – 72%, C I- 27%) were reported by Damasceno et al. Stroke 2010.
- In the INTERSTROKE Study, 1 month CFR was 22% in Africa (compared to 4% in HIC)
- In the Tanzanian Incident Stroke Study, case fatality rate was 28.7% at 1 month and 84.3% at 3 years (Walker et al, 2010)

Stroke Prevalence in Africa

- Southern Africa Stroke Prevention Initiative (SASPI), demographic surveillance in 2001, a stroke prevalence study in Agincourt, rural South Africa, with diagnosis of stroke based on the WHO definition of stroke, yielded an agestandardized (Segi world population) stroke prevalence of 290 per 100 000 people over the age of 15 years.
- In men, the prevalence was 281 per 100 000 and in women, 315 per 100 000 Connor MD, Lancet Neurol 2007; 6: 269-78.
- Crude prevalence rate of stroke in urban Nigeria 2005/2006 was 114/100,000 Danesi MA Neuroepidemiology 2007;28:216-23

Population – based prevalence rates of stroke in Africa

			Crud per			
Year	Country llocation/setting	Author	Overall	Male	Female	Age
1982	Nigeria, Igbo-Ora, rural	Osuntokun ³⁷	58	_	_	All
1985	Tunisia Kelibia, mixed*	Atia- Romdhane⁴	42	-	-	All
1988	Ethiopia, central Ethiopia, rural	Tekle Haimanot ³⁴	15	_	-	20-85
1993	Egypt, Sohag, mixed*	Kandil ³¹	508	520	490	All
1993	Egypt, Sohag, urban	Kandil ³¹	410	460	470	All
1993	Egypt, Sohag, rural	Kandil ³¹	540	510	570	All
1994	Tanzania, Hai, rural	Walker ⁴²	127	155	103	15+
2002	South Africa: Agincourt Health and Population Unit, Limpopo province, rural	Connor ⁴³	243	188	296	15+
2006	Nigeria, Lagos, urban	Danesi ³⁸	114	151	69	All
2009	Benin, Cotonou, urban	Cossi ⁴⁴	460	610	360	15+
2009	Egypt, Al-Kharga, mixed*	Farghaly ²²	560	610	510	All
2009	Egypt, Al-Kharga, urban	Farghaly ²²	580	620	530	All
2009	Egypt, Al-Kharga, rural	Farghaly ²²	520	580	458	All
2010	Tanzania, Hai district, rural	Dewhurst ³⁹	2300	2971	1752	70+
2010	Egypt, Assuit, urban	Khedr ³⁵	963	1174	736	All
2013	Egypt, Qena, mixed*	Khedr ^₄	922	1103	726	All
*Com	bined rates including both r	iral and urbar	comm	mities		

*Combined rates including both rural and urban communities.

Stroke Types in SSA – Data from the INTERSTROKE Study

	Overall (n=13447)	Western Europe, North America, Australia (n=1917)	Eastern and central Europe, Middle East (n=1394)	South America (n=1471)	China (n=3987)	South Asia (n=2850)	Southeast Asia (n=855)	Africa (n=973)
Age, years	62-2	66-7	63·9	65-8	61-9	59-6	56-6	58-7
	(13-6)	(13-4)	(13·4)	(14-3)	(12-5)	(12-9)	(13-0)	(15-2)
Age ≤45 years	1582	141	143	123	364	451	156	204
	(11-8%)	(7·4%)	(10·3%)	(8·4%)	(9·1%)	(15-8%)	(18·3%)	(21·0%)
Women	5434	781	556	652	1606	1017	352	470
	(40·4%)	(40-7%)	(39-9%)	(44·3%)	(40-3%)	(35·7%)	(41·2%)	(48-3%)
Intracerebral haemorrhage	3059	128	117	348	1102	785	285	294
	(22.7%)	(6·7%)	(8·4%)	(23-7%)	(27-6%)	(27-5%)	(33-3%)	(30·2%)
Ischaemic stroke	10388	1789	1277	1123	2885	2065	570	679
	(77-3%)	(93·3%)	(91·6%)	(76-3%)	(72-4%)	(72-5%)	(66-7%)	(69-7%)
OCSP classification*								
Total anterior circulation infarct	673/10388	71/1789	72/1277	176/1123	127/2885	105/2065	33/570	89/679
	(6-5%)	(4·0%)	(5-6%)	(15-7%)	(4·4%)	(5·1%)	(5-8%)	(13·1%)
Partial anterior circulation infarct	4872/10388	809/1789	631/1277	404/1123	1306/2885	945/2065	383/570	394/679
	(46·9%)	(45·2%)	(49·4%)	(36-0%)	(45·3%)	(45-8%)	(67·2%)	(58-0%)
Posterior circulation infarct	1509/10388	353/1789	265/1277	146/1123	372/2885	258/2065	63/570	52/679
	(14·5%)	(19·7%)	(20·8%)	(13-0%)	(12·9%)	(12·5%)	(11·1%)	(7·7%)
Lacunar infarction	2789/10388	536/1789	267/1277	212/1123	1051/2885	549/2065	79/570	95/679
	(26-9%)	(30-0%)	(20·9%)	(18·9%)	(36-4%)	(26-6%)	(13·9%)	(14·0%)
Undetermined	545/10388	20/1789	42/1277	185/1123	29/2885	208/2065	12/570	49/679
	(5·3%)	(1·1%)	(3·3%)	(16·5%)	(1·0%)	(10·1%)	(2·1%)	(7·22%)

O'Donnell et al, 2016.

Why is the burden of stroke increasing in low to middle income countries?

Feigin VL et al. Lancet Neurol 2009; 8:355-69

- The average age of the populations increases
- Exposure to cardiovascular risk factors increases:
 - Smoking
 - Raised blood pressure
 - Raised blood glucose concentrations
 - Westernised diets
 - Low in fruit and vegetables
 - High in fat and salt
 - Physical inactivity

Risk Factors

Risk factors for ischaemic and intracerebral haemorrhagic stroke in 22 countries (the INTERSTROKE study): a case-control study



Martin J O'Donnell, Denis Xavier, Lisheng Liu, Hongye Zhang, Siu Lim Chin, Purnima Rao-Melacini, Sumathy Rangarajan, Shofiqul Islam, Prem Pais, Matthew J McQueen, Charles Mondo, Albertino Damasceno, Patricio Lopez-Jaramillo, Graeme J Hankey, Antonio L Dans, Khalid Yusoff, Thomas Truelsen, Hans-Christoph Diener, Ralph L Sacco, Danuta Ryglewicz, Anna Czlonkowska, Christian Weimar, Xingyu Wang, Salim Yusuf, on behalf of the INTERSTROKE investigators^{*}

- 10 risk factors are associated with 90% population attributable risk (PAR) risk of stroke
- history of hypertension (OR 2.64)
- current smoking (OR 2.09)
- waist-to-hip ratio(OR 1.65 for highest vs lowest tertile)
- *diet risk score (OR 1.35)for highest vs lowest tertile);*
- regular physical activity (OR 0.69)
- diabetes mellitus (OR 1.36)
- alcohol intake (OR 1.51) for more than 30 drinks per month or binge drinking
- psychosocial stress (OR 1.30)
- depression (OR 1.35)
- cardiac causes (OR 2.38)
- ratio of apolipoproteins B to A1 (OR 1.89) highest vs lowest tertile

- Globally, 90% of the burden of stroke are attributable to modifiable risk factors
- Clusters of behavioural, metabolic and environmental risk factors:
- Behavioural (smoking, poor diet, low physical activity)
- Metabolic (\uparrow SBP, \uparrow BMI, \uparrow FPG, \uparrow TC, \downarrow GFR)
- Environmental (household air pollution from from solid fuels, ambient particulate matter pollution, lead exposure)

Global burden of stroke and risk factors in 188 countries, during 1990–2013: a systematic analysis for the Global Burden of Disease Study 2013



Valery L Feigin, Gregory A Roth, Mohsen Naghavi, Priya Parmar, Rita Krishnamurthi, Sumeet Chugh, George A Mensah, Bo Norrving, Ivy Shiue, Marie Ng, Kara Estep, Kelly Cercy, Christopher J L Murray, Mohammad H Forouzanfar, for the Global Burden of Diseases, Injuries, and Risk Factors Study 2013 and Stroke Experts Writing Group*

Summary

Background The contribution of modifiable risk factors to the increasing global and regional burden of stroke is Lancet Neurol 2016

Rank 1-5 11-15 6-10 > 15 Rick factor	Slobal	High-income Asia Pacific	Western Europe	Australasia	High-income North America	CentralEurope	Southern Latin America	Eastern Europe	ast Asia	Fropical Latin America	Central Latin America	South east A sia	Central Asia	Andean Latin America	North Africa and Middle East	Caribbean	5outh A sia	Dceania	Southern sub- Saharan Africa	Eastern sub- Saharan Africa	Cen tral sub- Saharan Africa	Western sub- Saharan Africa	
High systolic blood pressure	1	1	1	1	1	1	1	1	1	1	1	1	1	1	1	1	1	1	1	1	1	1	
Diet low in fruits	2	2	- 4	- 4	3	- 4	- 4	3	2	3	4	2	2	3	3	3	2	3	2	2	2	2	
High body-mass index	3	6	2	2	2	2	2	2	6	2	2	6	3	2	2	2	8	2	3	5	5	4	
Diet high in sodium	4	3	6	10	6	3	8	6	3	- 4	8	8	- 4	7	4	12	5	13	7	9	11	8	
Smoking	5	4	5	5	5	6	5	5	4	6	6	4	6	6	8	5	7	5	5	8	10	12	
Diet low in vegetables	6	5	3	3	4	5	3	4	11	5	3	3	5	4	5	4	4	4	4	4	4	5	
Ambient particulate matter (2.5 µm) pollution	7	8	11	14	12	8	12	10	5	9	9	9	8	9	6	11	6	15	11	12	6	6	
Household air pollution from solid fuels	8						15		7	14	12	5	12	11	14	7	3	6	8	3	3	3	
Diet low in whole grains	9	7	9	8	9	7	6	7	8	7	5	7	7	5	7	6	10	7	10	10	7	9	
High fasting plasma glucose	10	10	10	9	7	11	7	9	9	8	7	10	9	10	9	10	9	8	9	11	8	11	
Low physical activity	11	9	8	6	8	10	11	11	13	11	13	11	13	12	10	8	12	9	13	14	12	10	
Low glomerular filtration rate	12	12	7	7	13	9	10	12	14	10	10	14	10	8	11	9	11	12	6	7	9	7	
Alcohol use	13	11	12	11	10	12	9	8	12	12	11	15	11	14	16	15	15	11	12	13	14	14	
Lead exposure	14	14	14	13	15	14	16	15	10	15	16	12	14	15	13	13	13	14	14	6	13	13	
High total cholesterol		13	13	12	11	13	13	13	16	13	14	13	15	13	12	14	14	10	15	15	15	15	
Second-hand smoke		15	15	15	16	15	14	14	15	16	17	16	16	17	15	17	16	16	16	16	16	16	
Diet high in sugar-sweetened beverages	17	16	16	16	14	16	17	16	17	17	15	17	17	16	17	16	17	17	17	17	17	17	

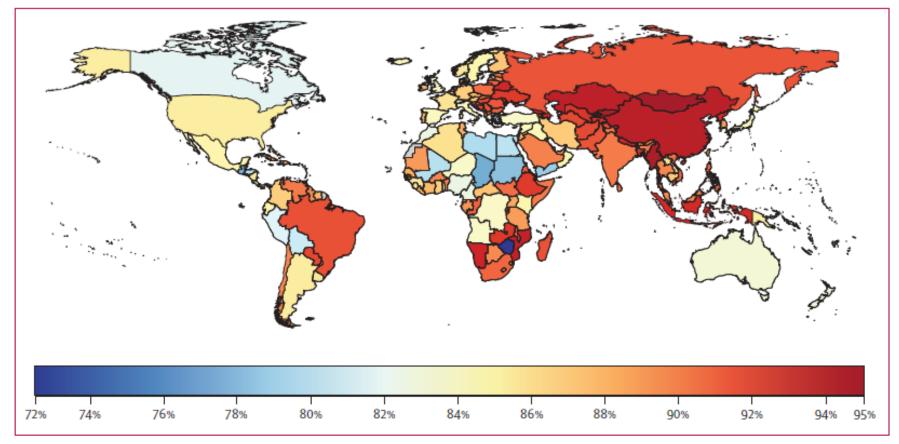
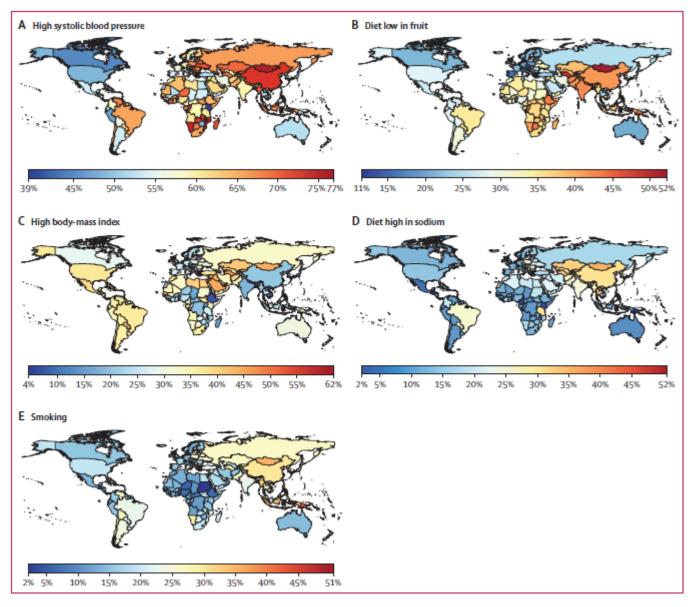
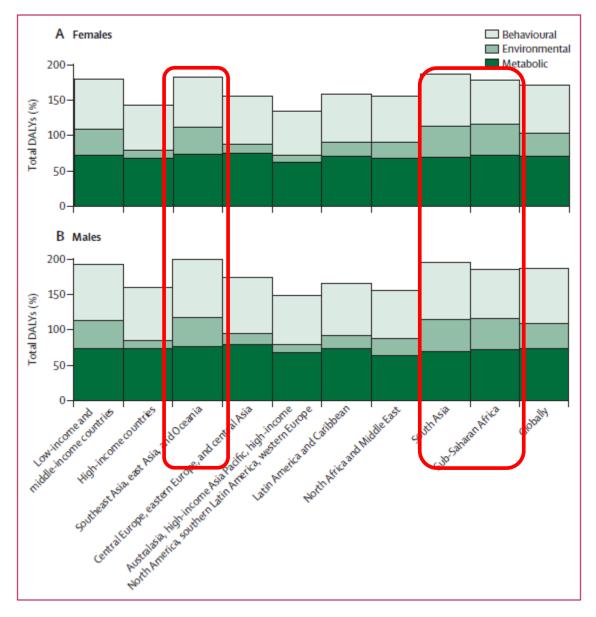


Figure 1: Stroke-related DALYs attributable to all modifiable risk factors combined for both sexes in 2013 DALY–disability-adjusted life-year.

Feigin et al, 2016



Feigin et al, 2016



Feigin et al, 2016

Global and regional effects of potentially modifiable risk factors associated with acute stroke in 32 countries (INTERSTROKE): a case-control study

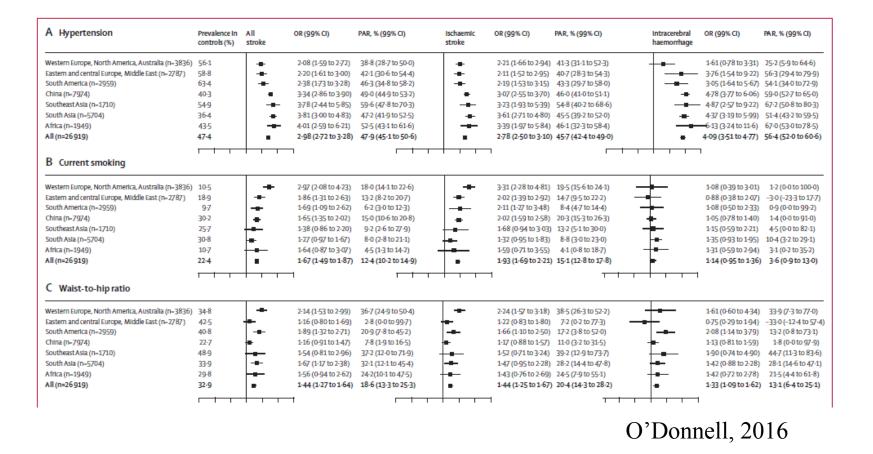


Martin J O'Donnell, Siu Lim Chin, Sumathy Rangarajan, Denis Xavier, Lisheng Liu, Hongye Zhang, Purnima Rao-Melacini, Xiaohe Zhang, Prem Pais, Steven Agapay, Patricio Lopez-Jaramillo, Albertino Damasceno, Peter Langhorne, Matthew J McQueen, Annika Rosengren, Mahshid Dehghan, Graeme J Hankey, Antonio L Dans, Ahmed Elsayed, Alvaro Avezum, Charles Mondo, Hans-Christoph Diener, Danuta Ryglewicz, Anna Czlonkowska, Nana Pogosova, Christian Weimar, Romaina Iqbal, Rafael Diaz, Khalid Yusoff, Afzalhussein Yusufali, Aytekin Oguz, Xingyu Wang, Ernesto Penaherrera, Fernando Lanas, Okechukwu S Ogah, Adesola Ogunniyi, Helle K Iversen, German Malaga, Zvonko Rumboldt, Shahram Oveisgharan, Fawaz Al Hussain, Daliwonga Magazi, Yongchai Nilanont, John Ferguson, Guillaume Pare, Salim Yusuf; on behalf of the INTERSTROKE investigators*

Summary

Background Stroke is a leading cause of death and disability, especially in low-income and middle-income countries. Published Online We sought to quantify the importance of potentially modifiable risk factors for stroke in different regions of the world ^{July 15, 2016}

O'Donnell, 2016



- Data confirm regional/geographic variation of PAR of stroke risk factors
- Implications for region- and country- specific primary and secondary prevention

Drivers of changing stroke epidemiology

- Demographic transition (population growth and ageing)
- Urbanization and changing environments
- Changing risk factor pattern
- Racial (? Genetic factors, gene gene, gene environment interactions)

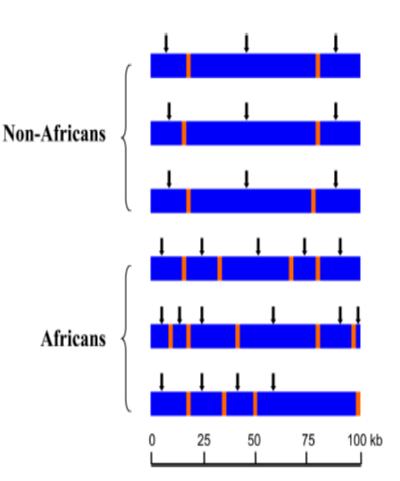
Is Genetics important for Stroke in Africa ?

Stroke Disparities and African ancestry

- Significant racial (ethnic) and geographical (environmental) disparities occur in stroke
- African Americans have a higher predisposition, worse severity and often poorer outcomes compared to Caucasian Americans (US Studies).
- Black stroke survivors had poorer cognitive outcome compared to other racial groups (South London Stroke Register Study)
- Inherited genetic variations offer a possible explanation for the observed peculiarities of stroke in populations of African ancestry

Genetic diversity of African populations

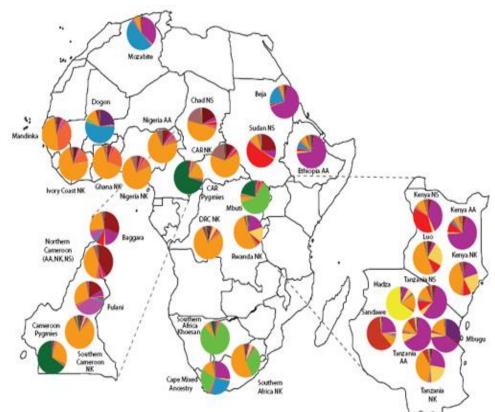
- The African population exhibits great genetic diversity.
- This contributes to variations in disease pattern eg Hbs and malaria .



Campbell and Tishkoff; Annu Rev Genomics Hum Genet. 2008 ; 9: 403–433

Genetic diversity of African populations

- The import of this on the pathobiology, phenomics and pharmacogenomics of cerebrovascular disorders are largely unexplored.
- Exploring genomics of diseases has great potentials for deeper insight, new therapeutics, neuro-protective agents and personalized medicine



Campbell and Tishkoff; Annu Rev Genomics Hum Genet. 2008; 9: 403–433

Genomics and health disparities

Adapted Gene	Beneficial Trait	Negative Outcome
HBB	Confers incomplete resistance to lethal forms of malaria	Sickle Cell Trait Sickle Cell Disease
Regulatory Deficiencies of HBA and HBB	Confers incomplete resistance to lethal forms of malaria	α and β thalassemia
APOL1	Protection against Human African Trypanosomiasis	Kidney disease
ATP1A1, AQP2, CSK	Climate adaptation	Hypertension and Osmoregulation
DMD	Confers protection against Lassa fever	Duchenne Muscular Dystrophy

Rotimi et al, 2016

The burden of stroke in Africa: a glance at the present and a glimpse into the future

Mayowa O Owolabi, Sally Akarolo-Anthony, Rufus Akinyemi, Donna Arnett, Mulugeta Gebregziabher, Carolyn Jenkins, Hemant Tiwari, Oyedunni Arulogun, Albert Akpalu, Fred Stephen Sarfo, Reginald Obiako, Lukman Owolabi, Kwamena Sagoe, Sylvia Melikam, Abiodun M Adeoye, Daniel Lackland, Bruce Ovbiagele, as members of the H3Africa Consortium

CVJAFRICA • Volume 26, No 2, H3Africa Supplement, March/April 2015

Stroke genomics in people of African ancestry: charting new paths

RO Akinyemi, B Ovbiagele, A Akpalu, C Jenkins, K Sagoe, L Owolabi, F Sarfo, R Obiako, M Gebreziabher, E Melikam, S Warth, O Arulogun, D Lackland, A Ogunniyi, H Tiwari, RN Kalaria, D Arnett, MO Owolabi, for the SIREN investigators as members of the H3Africa Consortium



Contents lists available at ScienceDirect

Journal of the Neurological Sciences

journal homepage: www.elsevier.com/locate/jns

Review article

Neurogenomics in Africa: Perspectives, progress, possibilities and priorities



S2

S39

Rufus O. Akinyemi ^{a,b,*}, Mayowa O. Owolabi ^c, Tolulope Oyeniyi ^d, Bruce Ovbiagele ^e, Donna K. Arnett ^f, Hemant K. Tiwari ^f, Richard Walker ^g, Adesola Ogunniyi ^c, Raj N. Kalaria ^{b,**} and SIREN group of H3Africa Consortium





Stroke Investigative Research and Education Network (SIREN)



National Institute of Neurological Disorders and Stroke Reducing the burden of neurological disease...

NIF

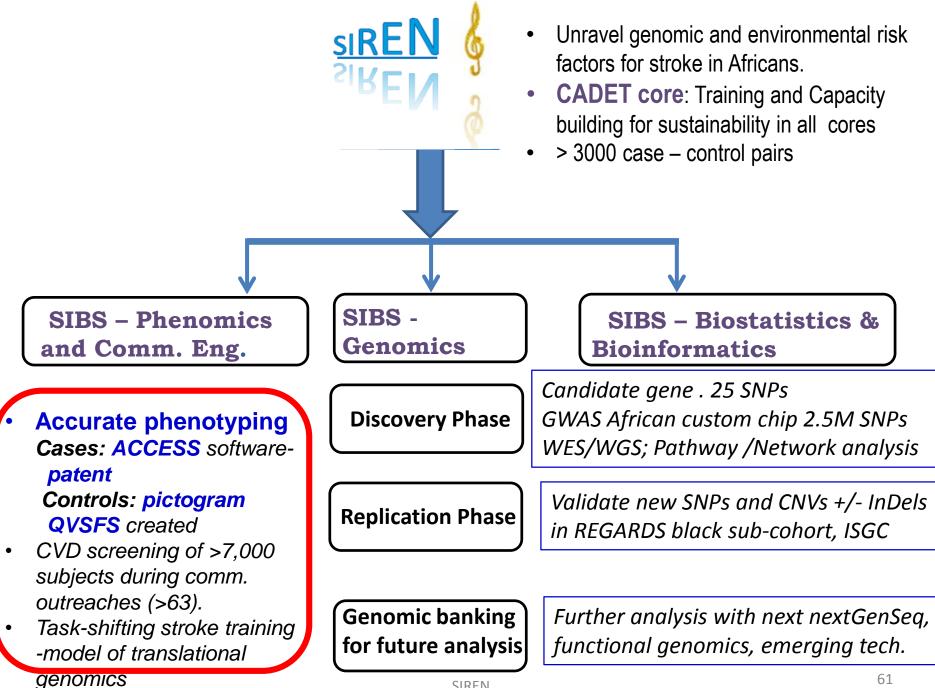


SIREN: Central Hypothesis and Objective

To develop suitable interventions, the stroke epidemic in Africa needs accurate epidemiologic, phenotypic and genomic characterization (first pillar of Stroke Quadrangle).

- ✓ Aim: To evaluate the *genomic and environmental risk factors for stroke in sub-Saharan Africa*, while simultaneously **building** *sustainable capacities* in phenomics, biobanking, genomics, biostatistics and bioinformatics.
- ✓ Hypothesis: Distribution of unique and novel genomic and environmental factors will account for the peculiar and hitherto unreported <u>pattern, incidence, type, subtype and outcome</u> of stroke among individuals of African ancestry.





AIM on ClearCanvas Enriched Stroke phenotyping Software : ACCESS

 For standard/uniform reporting templates across multiple sites in multicenter studies.
 USER-FRIENDLY

Facilitates...

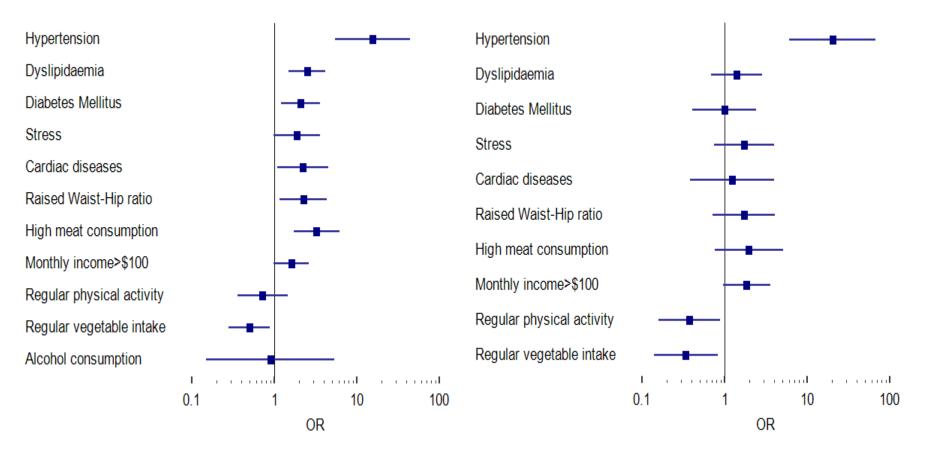
- concordant and reproducible classification of all stroke subtypes by multiple investigators
- Multi-level adjudication/ quality control of image report
- Export of image data to other formats -XML map to Excel and merges easily with REDCap database
- Archiving, backup & security

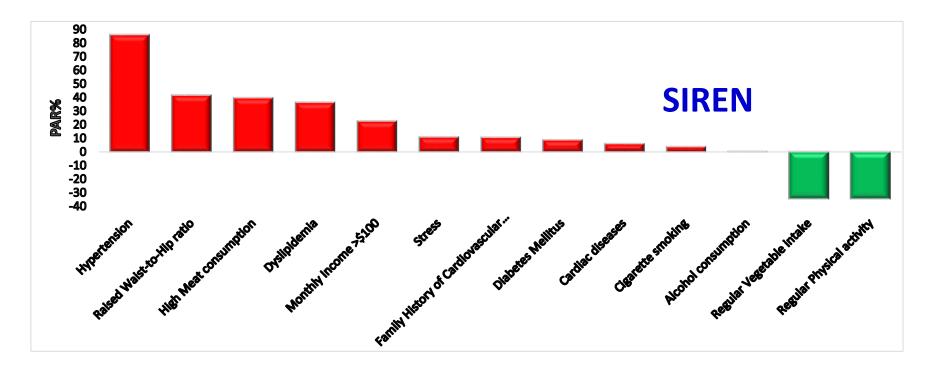


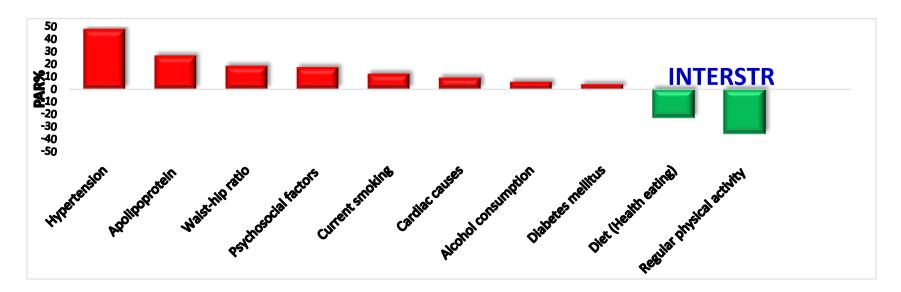
Preliminary Results

Risk factors for Ischemic stroke (64 %)

Risk factors for hemorrhagic stroke (36%)



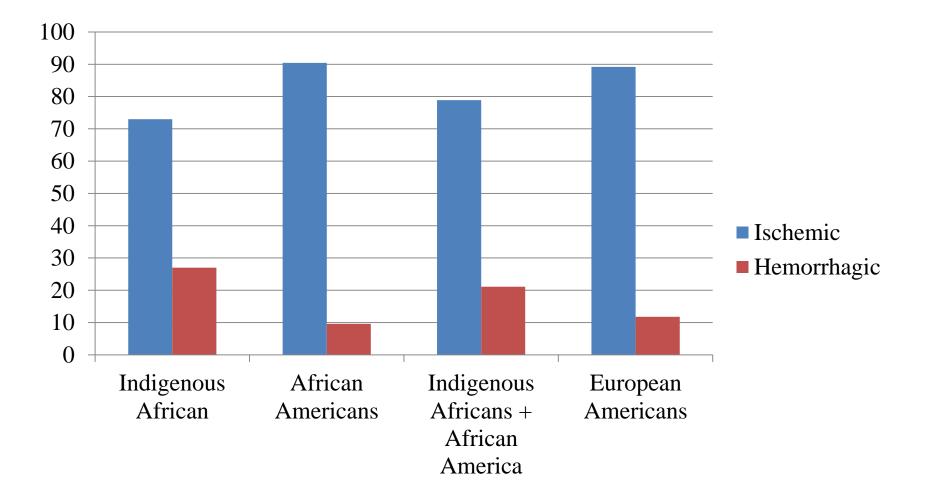


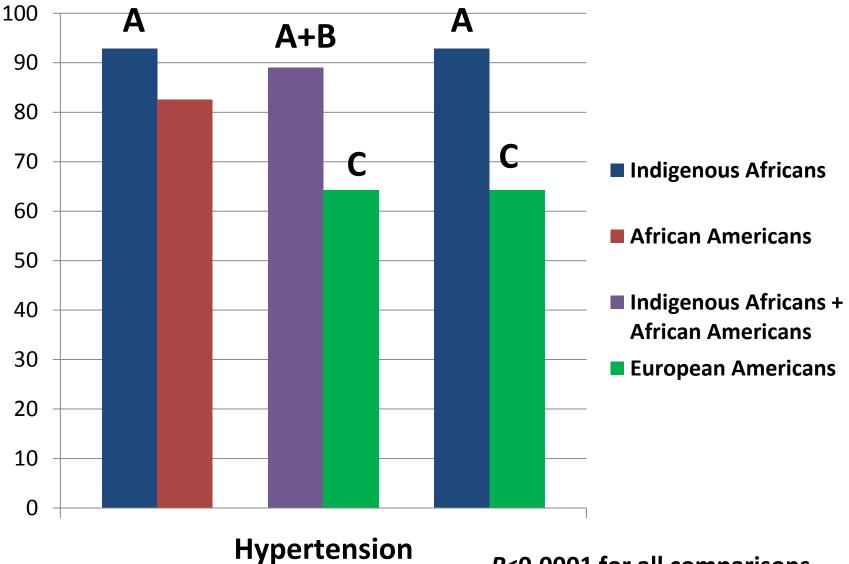


Comparison with REGARDS (African Americans and European Americans)

- We compared stroke type and contributions of vascular risk factors among indigenous Africans (IA), African Americans (AA) and European Americans (EA)
- Using harmonized assessments and standard definitions, we compared data on stroke type and established risk factors for stroke in acute stroke cases age>55 years in SIREN and REGARDS
- There were 811 IA, 452 AA and 665 EA stroke subjects, with mean age of 68.0±9.3, 73.0±8.3 and 76.0±8.3 years respectively (p<0.0001).
- Hemorrhagic stroke was higher in IA (27%) compared to AA (8%) and EA (5.4%; p < 0.001). Lacunar strokes were highest in IA (47.1%), followed by AA (35.1%), and then EA (21.0%; p < 0.0001).

Proportion of primary stroke types





P<0.0001 for all comparisons

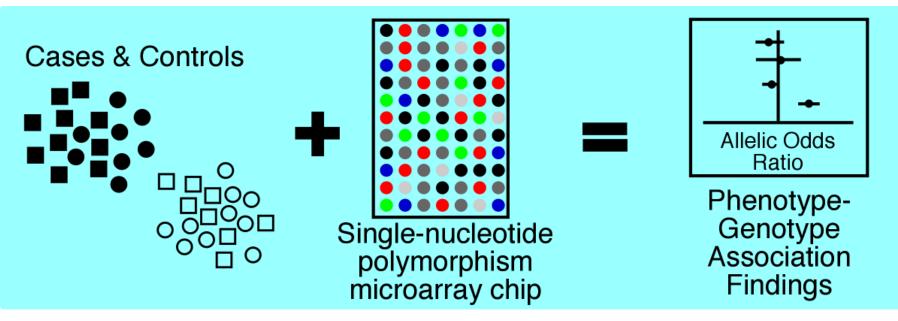
- Hypertension was highest in IA (92.8%), followed by AA (82.5%) and then EA (64.2%; p<0.01).
- Diabetes mellitus (DM) was similar in IA (38.3%) and AA (36.8%) but lower in EA (21%; p<0.0001).
- Pre-morbid sedentary lifestyle was similar in AA (37.7%) and EA (34%) but lower in IA (8%).
- Behavioural factors such as sedentary lifestyle, alcohol and smoking may contribute to the higher proportion of ischemic stroke in AA compared to IA,
- <u>Inherited genetic variations offer a possible explanation for</u> <u>the higher proportion of hypertension and DM among stroke</u> <u>subjects of African ancestry as well as the proportion of risk</u> <u>that remains unexplained by traditional and emerging risk</u> <u>factors alone.</u>

Candidate gene studies

			Location	Minor		
CHR	Gene	SNP	Basepair	Allele	OR	p-value
7	IL6	rs1800796	22766246	С	1.54	0.01329
4	PITX2	rs2200733	111710169	Т	0.8239	0.08378
4	CSN3	rs3775745	71113074	G	1.192	0.09064
19	APOE	rs7412	45412079	Т	1.258	0.1331
9	CDKN2A/CDKN2B	rs2383207	22115959	А	0.6124	0.1644
17	ACE	rs4343	61566031	G	1.162	0.2242
9	CDKN2A/CDKN2B	rs1333040	22083404	С	1.099	0.3574
19	APOE	rs429358	45411941	С	1.108	0.3878
7	HDAC9	rs11984041	19031935	Т	0.8924	0.3938
22	CELSR1	rs6007897	46780521	Т	1.093	0.3958
9	ANRIL	rs10757274	22096055	G	0.8909	0.4125
7	IL6	rs2069832	22767433	А	2.503	0.4322
5	CD14	rs2569190	140012916	А	0.9187	0.442
22	APOL1	rs73885319	36661906	G	0.9478	0.6075
7	HDAC9	rs2107595	19049388	А	0.9331	0.6105
4	PITX2	rs2634073	111665783	Т	0.9512	0.6228
22	CELSR1	rs9615362	46795141	С	1.049	0.6506
9	ANRIL	rs10757278	22124477	G	1.046	0.7664
22	APOL1	rs60910145	36662034	G	0.9743	0.7858
16	ZFHX3	rs879324	73068678	А	0.9791	0.8793
	intergenic region near					
1	TSPAN2	rs12122341	115655690	G	1.027	0.8994
16	ZFHX3	rs16971456	73013036	G	1.013	0.9263
7	HDAC9	rs28688791	19039605	С	0.9932	0.9552
11	Sickle cell (HBB)	rs334	5248232	Т	NA	NA

Next Steps.....

Genome-Wide Association Study using the H3Africa Consortium Chip



•Can probe for associations between a million (+?) genetic variants (SNP) and stroke phenotypes

- •May identify novel genes-i.e., genes not previously suspected of playing a role in stroke
- •Multiple testing can be problematic. Candidate genes identified here must be examined in replication studies

Tackling the growing Stroke burden in Africa



Stroke Turns 40

Cerebrovasc Dis 2010;30:127–147 DOI: 10.1159/000315099 Published online: May 24, 2010

Stroke: Working toward a Prioritized World Agenda

Vladimir Hachinski¹ Geoffrey A. Donnan^{5, a, h} Philip B. Gorelick^{8, b, h} Werner Hacke^{30, c, h} Steven C. Cramer ^{10, d, h}

- The impact of stroke will grow into the future as populations age, and globalization grows with attendant lifestyle changes – especially in developing economies including Nigeria.
- To mark 40 years of the journal 'Stroke' Vladmir Hachinski, the WFN president led a synergium to advance the frontiers of stroke in all facets in all the regions of the world.

Stroke Neurologic Manpower in Africa

- No Stroke specific postgraduate medical education programme in Africa (academic or professional)
- Most African strokologists are primarily neurologists with different levels of further training and research interest in stroke.
- Stroke units are springing up Cape Town, Accra, Benin city, Ilorin, Cairo, with residents in Neurology, Internal Medicine and Geriatrics rotating through them.

"PROBLEMS CAN BECOME OPPORTUNITIES WHEN THE RIGHT PEOPLE COME TOGETHER."

ROBERT SOUTH

© Lifehack Quotes

Opportunities in stroke medicine in Africa

- Paucity of strokologists and allied health workers for effective MDT stroke research and practice.
- Neuroimaging challenges availability and cost
- Lack of stroke rehab facilities
- Inequity of access to health care
- Stroke literacy and cultural beliefs
- Lack of stroke systems of care

Strategic keys to bridging the stroke training gap in Africa

- Global Health Perspective of Stroke Medicine
- African and North South Networking and Collaborations
- Strong Leadership and Team work
- Creativity and Innovation : eg.Deploying IT
- Mentorship
- Tapping into the African Diaspora: Turning 'Brain drain' to 'Brain gain'
- 'Training Service Research' Synergy

Strengthening capacity for health research in Africa

James A GWhitworth, Gilbert Kokwaro, Samson Kinyanjui, Valerie A Snewin, Marcel Tanner, Mark Walport, Nelson Sewankambo

Kaddumukasa et al. Human Resources for Health 2014, 12:75 http://www.human-resources-health.com/content/12/1/75



CASE STUDY

Open Access

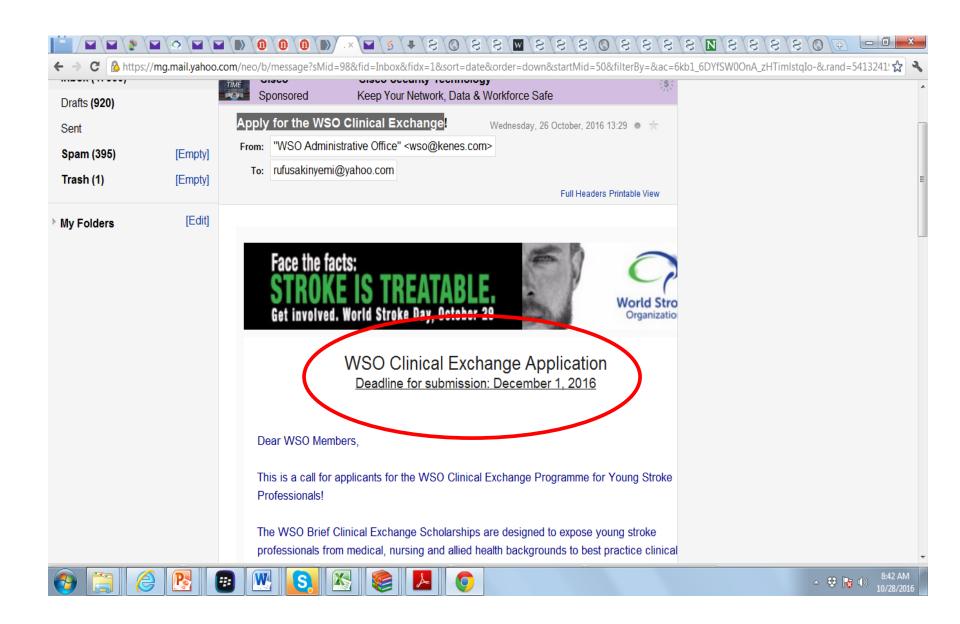
Global medical education partnerships to expand specialty expertise: a case report on building neurology clinical and research capacity

Mark Kaddumukasa^{1*}, Elly Katabira¹, Robert A Salata², Marco A Costa³, Edward Ddumba⁴, Anthony Furlan⁵, Angelina Kakooza-Mwesige⁶, Moses R Kamya¹, James Kayima¹, Chris T Longenecker², Harriet Mayanja-Kizza¹, Charles Mondo¹, Shirley Moore⁷, Svetlana Pundik⁸, Nelson Sewankambo⁹, Daniel I Simon², Kathleen A Smyth¹⁰ and Martha Sajatovic¹¹

World Stroke Academy (WSA)

- The World Stroke Academy is a global online educational resource aimed at major stroke issues for CME of healthcare professionals
- Professional section:
 - Case based adaptive e-Learning
 - Interactive eLearning lectures
 - Webcast/podcast past congresses
 - Article reviews
 - Guidelines
 - CME certification
- Patient/carers section:





Guidelines

World Stroke Organization Global Stroke Services Guidelines and Action Plan

Patrice Lindsay^{1,2,*,†}, Karen L. Furie^{3,4,†}, Stephen M. Davis^{5,6,†}, Geoffrey A. Donnan^{6,7,†}, and Bo Norrving^{8,†}

Int J Stroke 2014;9:4-13

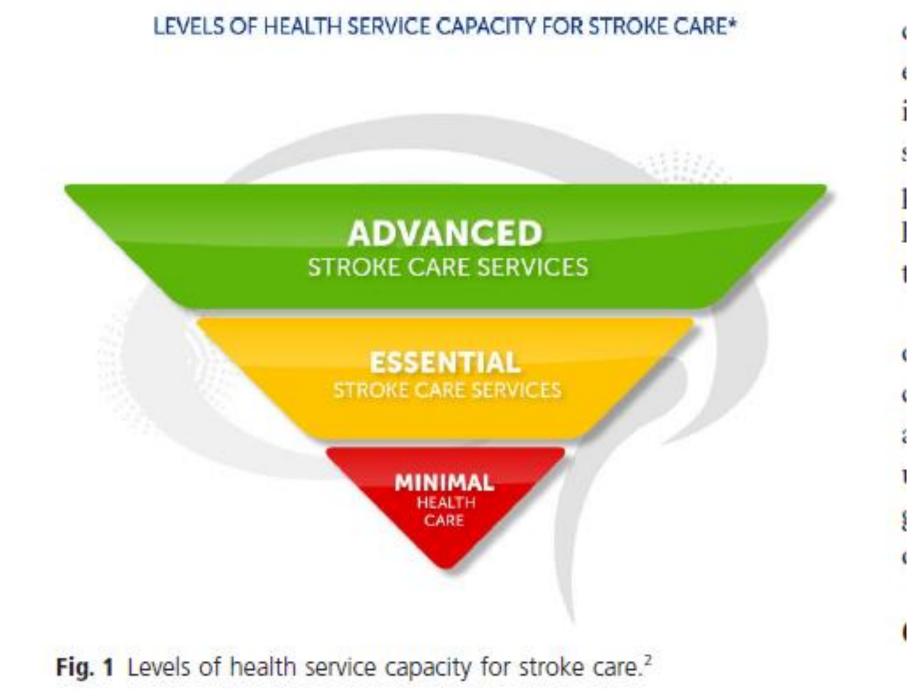


Table 1 Health service capacity for stroke care checklists*		
Advanced stroke services	Essential stroke services	Minimal healthcare services
 Access to advanced diagnostic services Access to physicians with stroke expertise Access to advanced interventions in addition to tPA, such as interventional radiology and neurosurgery Access to specialist rehabilitation therapists Access to community programs for recovery after stroke Fully coordinated stroke care provided across geographically discrete regions 	 Access to basic diagnostic services – laboratory, ECG, CT scan, ultrasound Access to nurses Access to physicians, although may not be stroke specialists Access to acute thrombolysis with tPA Access to elements of stroke unit care, including members of an interdisciplinary stroke team Access to rehabilitation services Access to stroke prevention therapies such as aspirin, lifestyle change recommendations, blood pressure management Limited coordinated stroke care provided across geographically discrete regions 	 Variable access to healthcare workers (nurses or lay workers) Very limited access to physicians No access to diagnostic services or hospital care Limited access to the most basic lifestyle preventative advice Care provided in local communities without coordination across defined geographic regions

*These checklists should be used for self-assessment and for stroke services planning. The goal is to achieve as many checkmarks as possible and continually strive to provide the highest level of stroke services that is realistically and reasonably attainable, given local and regional resources and circumstances.

CT, computed tomography; ECG, electrocardiogram; tPA, tissue plasminogen activator.

Stroke medicine sub-specialty interest among young African neurologists and trainees

- 77 young African neurologists and trainees (IWGYNT – African group database)
- 10 African countries
- Sub specialty interest (multiple choices)
 - Stroke medicine : (35 (45.5%))
 - Epileptology : 28 (36.4%)
 - Movement disorders : 21 (27.3%)
 - Neuroinfections : 17 (22.1%)
 - Cognitive Neurology: 10 (13.0%)

Section Editors

Johan A. Aarli, MD Oded Abramsky, MD, PhD, FRCR

Global Perspectives



James H. Bower, MD Amadou Gallo Diop, MD, PhD Riadh Gouider, MD Erich Schmutzhard, MD Addrean African A total G

the home of no

8th REGIONAL TEACHING COURSE IN SUB-SAHARAN AFRICA MAPUTO, MOZAMBIQUE 10 - 12 NOVEMBER, 2016

Organised by EAN - European Academy of Neurology

In cooperation with:

African Academy of Neurology American Academy of Neurology International Brain Research Organisation International PD and Movement Disorder Society World Federation of Neurology

World Stroke Organisation

and Eduardo Mondlane University of Maputo, Mazambique of Neurological Societies inuing medical education courses have always been as many participants from his is an ideal opportunity the world to cooperate in

JE

VOL. 26 • NO. 1 • FEBRUARY 2011



THE OFFICIAL NEWSLETTER OF THE WORLD FEDERATION OF NEUROLOGY

Nigeria-Florida Alliance Focuses on Training

BY DIANA M. SCHNEIDER, PH.D.

new alliance between the Florida Society of Neurology and the Nigerian Society of Neurological Sciences and Nigerian Stroke Society - the Nigeria-Florida Neuroscience Partnership - aims to develop neurology training and services in the West African country and eventually to expand to include research collaborations, exchange programs, mentoring, patient management, teleconferencing, and other collaborative efforts.

The Nigerian Stroke Society (NSS) is an affiliate of the Nigerian Society of Neurological Sciences (NSNS) and the World Stroke Organisation. All NSNS members interested in stroke are NSS members. It was inaugurated in 2009. Prof. Yomi Ogun is president, Dr. Njide Okubadejo is vice president, and Dr. Bamidele Osalusi is secretarygeneral.

The partnership was forged in response to recent study findings demonstrating that Nigeria's high stroke and associated mortality rates, coupled with a shortage of trained neurologists and a lack of understanding of stroke in patients and health care providers

risk for further straining its already diminished resources. The organizers

also plan to focus on the development of clinical programs such as stroke units, independent subspecialty clinics, neurosurgical subspecialties, as well as the creation of guidelines for how neurological emergencies can

be approached

neurologists

and neurosurgeons, and by practitioners in pediatrics, internal medicine, emergency medicine, and general surgery when neurologists and neurosurgeons are not available.

The first Nigeria-Florida Neuroscience Partnership (NFNP) conference was held Nov. 2-5 in Lagos, Nige-

alike, put the country at Nigeria - Florida Neuroscience Partnership ria. The Joint Steering

Committee was co-chaired by Dr. Michael Finkel, representing the Florida Society of Neurology (FSN) and the World Neurology Foundation (WFN); Prof. Yomi Ogun of the NSNS, with the help of Dr. Niide Okubadejo, Dr. Rufus Akinyemi, and Dr. Bunmi Ogunrin, all members of both the NSNS and NSS; and Dr. Hubert Fernandez, Dr. Ali Malek, and Dr. Susan Naselli, all of the FNS.

Dr. Susan Naselli of the Florida Neurological Society presents a

neurological Tool Kit to a participant at the partnership meeting.

See Nigeria-Florida • page 5

INSIDE

A WFN Association of Parkinsonism and Related Disorders-sponsored teaching course aimed to update neurologists on advances in the field and encourage medical students to consider it as a specialty. PAGE 3

Europe

A recent report says many aspects of epilepsy care in Europe are seriously underresourced. PAGE 6

Travelling Fellowships

The WFN invites applications for the 2011 Junior Travelling Fellowships, and last year's Fellows share their experiences and lessons learned at their selected conferences. PAGES 2 AND 8

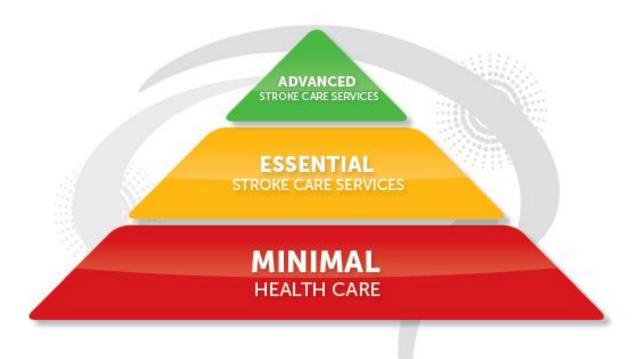


by



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What about allied stroke professionals ?



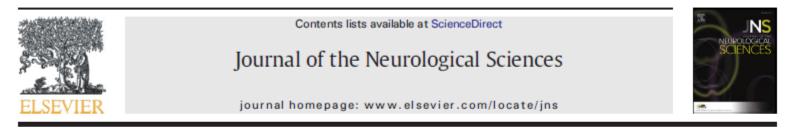
Nurse - centered discharge planning may be effective for home – based rehabilitation

Comprehensive Discharge Planning and Home Follow-up of Hospitalized Elders A Randomized Clinical Trial

Mary D. Naylor, PhD	Context Comprehensive discharge planning by advanced practice nurses has dem- onstrated short-term reductions in readmissions of elderly patients, but the benefits of more intensive follow-up of hospitalized elders at risk for poor outcomes after dis- charge has not been studied.	
Dorothy Brooten, PhD		
Roberta Campbell, MSN		
Barbara S. Jacobsen, MS	Objective To examine the effectiveness of an advanced practice nurse-centered discharge	
Mathy D. Mezey, EdD	planning and home follow-up intervention for elders at risk for hospital readmissions.	
Mark V. Pauly, PhD	Design Randomized clinical trial with follow-up at 2, 6, 12, and 24 weeks after in- dex hospital discharge.	
J. Sanford Schwartz, MD		
-	 Setting Two urban, academically affiliated hospitals in Philadelphia, Pa. 	

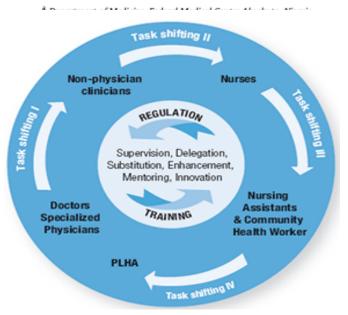
JAMA, February 17, 1999-Vol 281, No. 7

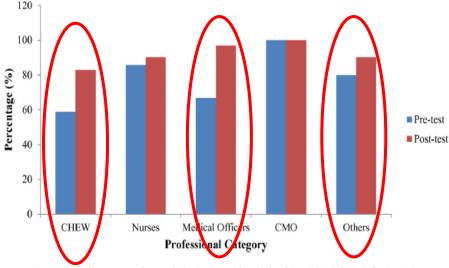
Conclusion: An advanced practice nurse–centered discharge planning and home care intervention for at-risk hospitalized elders reduced readmissions, lengthened the time between discharge and readmission, and decreased the costs of providing health care.



Task-shifting training improves stroke knowledge among Nigerian non-neurologist health workers

Rufus O. Akinyemi ^{a,g,*}, Mayowa O. Owolabi ^b, Philip B. Adebayo ^c, Joshua O. Akinyemi ^d, Folajimi M. Otubogun ^a, Ezinne Uvere ^b, Olaleye Adeniji ^a, Osimhiarherhuo Adeleye ^a, Olumayowa Aridegbe ^a, Funmilola T. Taiwo ^f, Shamsideen A. Ogun ^e, Adesola Ogunniyi ^b





CrossMark

Fig. 3. Association between respondents' professional category and knowledge of thrombolysis before and after the training.









Final words.....

Building neurological services in Africa

International institutions will need to work in partnership with Africa to meet the growing burden of neurological disorders across the continent. Many see the creation of the World Federation of Neurology's Africa Committee as an important step in the right direction. Rebecca Craven reports.

www.thelancet.com/neurology Vol 8 January 2009



Faculty at EAN RTC in Dakar, Senegal, July 2013

future of neurological sciences in Africa"

- Prof. Alfred Njamnshi.

 \checkmark "I like the dreams of the future better than the history of the past"

 Thomas Jefferson.
 ✓ "The future belongs to those who believe in the beauty of their dreams"

-Eleanor Roosevelt

Summary

- An increase in global stroke burden with regional variations, LMIC being worst affected
- Growing burden of Stroke in Africa
- The current epidemiology is driven by certain modifiable and non modifiable factors
- Capacity building, including task shifting approaches and stroke services devp are needed to curtain the growing burden in Africa.

Acknowledgements



• UK Raj Kalaria Peter Sandercock (WSO) Janet Slade Peter Sandercock

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Bruce Ovbiagele Donna Arnett Hemant Tiwari Mulugeta G. Samantha Hurst Philip Gorelick

• Nigeria

Adesola Ogunniyi Mayowa Owolabi Joshua Akinyemi Godwin Ogbole Benjamin Aribisala Philip Adebayo Akin Ajani Bayo Ojo Adeola Akintola Ezinne Uvere Sylvia Melikam Osi Adeleye Mayowa Ogunrombi



- Valery Feigin for a couple of slides
- My Teachers, Friends and Family





for Neurochemistry

