HEPATITIS C THERAPEUTIC OPTIONS IN AFRICA

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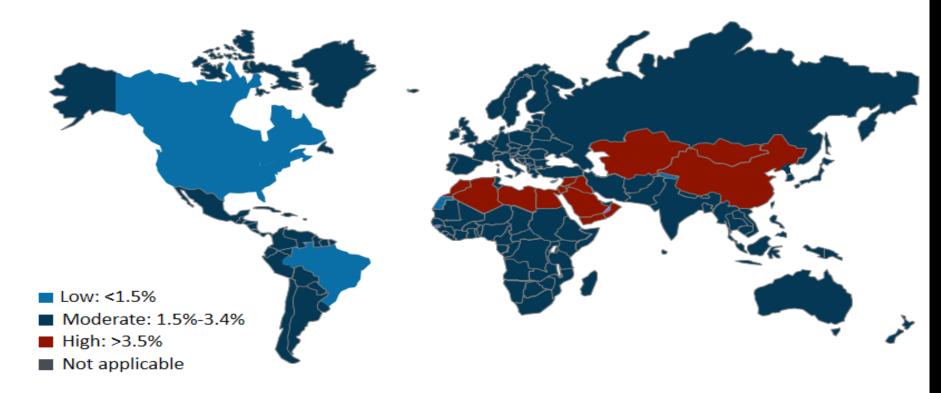
OUTLINE- SOME QUESTIONS

- 1. Is HCV a public health threat in SSA
- 2. Who do we screen?
- 3. HCV treatment in Nigeria
- 4. Challenges to expanding therapeutic access in SSA

INTRODUCTION

- HCV is the most common chronic blood borne infection in the USA affecting at least 3 million Americans
- It is the leading cause of end-stage liver disease and liver transplantation.
- Availability of highly effective, yet highly costly curative therapy for HCV has converted hepatitis care into an emergency requiring urgent responses including an urgency to increase awareness and screening
- WHO elimination goals for 2030

PREVALENCE 170 MILLION PEOPLE

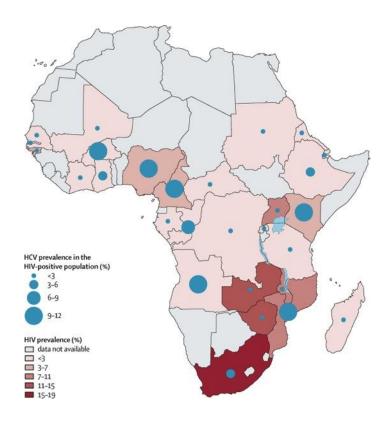


 Estimates are derived from a meta-analysis of data from 232 studies published between 1997 and 2007 and NHANES data up to 2010. Point prevalence estimates are calculated using regional population age weights.

SUB-SAHARAN AFRICA: REGIONAL CHARACTERISTICS & VIRAL HEPATITIS CONTROL

- Population, total 960.1 million 2014
- Low income economies-60%
 - GNI per capita, Atlas method 2013 (\$1,657)
- Urban population 37% (2014)
- Chronic HBV & HCV endemic -Significant public health & economic impact. Estimated 18 million cases of cirrhosis/HCC within next 20 yrs

CHRONIC HCV IN AFRICA- 19 MILLION INFECTIONS- NEARLY 11% OF GLOBAL INFECTIONS



HCV seroprevalence 1.0-14%

 differential historical exposure to iatrogenic transmission

SSA: Highest prevalence

- Cameroon 4.9%,
- Burkina Faso 6.1%
- Gabon 4.9%

SSA: Largest absolute number

- Nigeria- 3.1% (2.5 million persons)
- Ethiopia- 2.7% (1.26 million persons)

CHRONIC HCV IN AFRICA- 19 MILLION INFECTIONS-NEARLY 11% OF GLOBAL INFECTIONS

In Egypt

 Estimated prevalence is 14.7% affecting an estimated 6.8 million

In Africa

Highest prevalence

- Middle Africa (Cameroon)
- West Africa (Burkina Faso, Gabon, Benin)

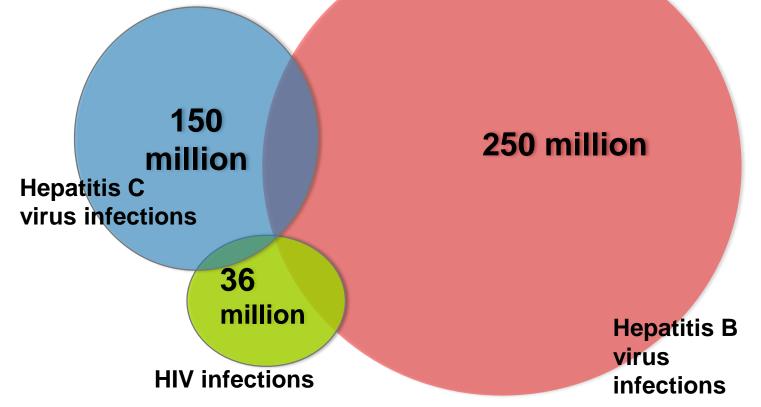
Central /West Africa Country	Estimated adult seroprevalenc e (%)	Estimated number of adult carriers (thousands)
Angola	3.9	370
Burundi	3.1	150
Cameroon	4.9	525
Burkina faso	6.1	475
DRC	2,1	647
Ghana	3.2	426
Ethiopia	2.7	1206
Congo republic	2.9	224
Rwanda	3,1	175
Nigeria	3.1	2575

POOLED HCV SEROPREVALENCE -SSA

Risk	Prevalence	CI		Numb studie		Prevalence (95% CI)
Low risk	2.7%		HCV in HIV Overall	101		5·73 (4·90–6·56)
cohorts					+	
			Central	16		5.87 (3.71-8.04)
Antenatal	3%	2.2-3.8	Southeast	43	-	4.56 (3.70–5.43)
clinics			West	42	→	6.67 (4.83-8.50)
			HIV in HCV			
Blood	2%	1.9-2.1	Overall	40	│ • • • • • • • • • • • • • •	- 15.86 (12.47-19.24)
donors			Central	8	│ <u> </u>	5.89 (2.49-9.31)
HIV	5.7%	4.9-6.6	Southeast	14		→ 33.42 (17.48-49.35)
positive	West	18	│ • ─ ─ ─ ─ ─ ─ ─ ─ ─ ─ ─ ─ ─ ─ ─ ─	10.05 (6.13-13.98)		
P 001110						
					0 2 4 6 8 10 12 14 16 18	8 20
					Prevalence (%)	

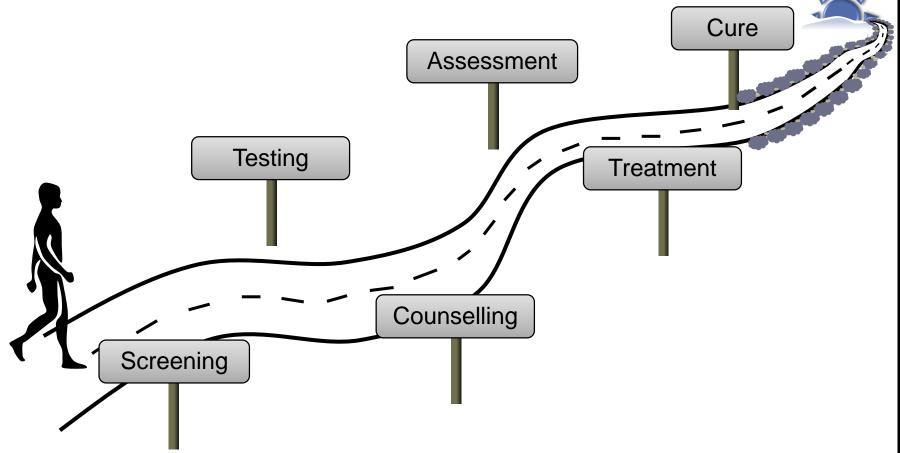
Roa VB, et al. Lancet infectious diseases 2015; 15; 819 to 824

Overlapping Epidemics of HBV, HCV and HIV

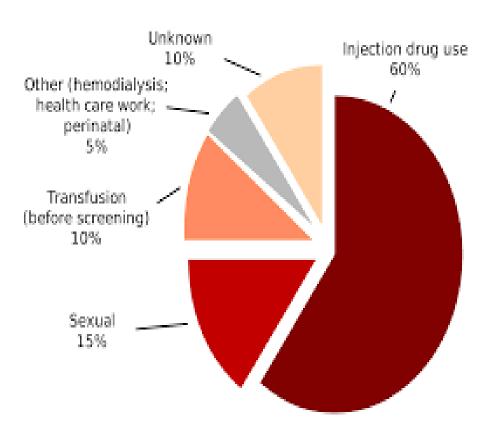


Viral hepatitis has become one of the most clinically important co-morbidity among people living with HIV. (co infection with HBV and HCV occur in 15% and 7%

THE ROAD TO THERAPEUTIC OPTIONS AND CURE FOR HCV INFECTION BEGINS WITH SCREENING



RISK FACTORS FOR VIRAL HEPATITIS-HCV



- latrogenic transmission
- Egypt: parenteral antischistosomal mass treatment in the 1960s and 1970s
- Mass treatment campaigns against yaws, malaria, syphilis in Cameroon, Gabon, CAR and DRC

Pepin j et al, clin Infectious diseases 2010 Njouom R et al, J gen viro. 2009 Ndong-Atome Gr et al, J med virol. 2008

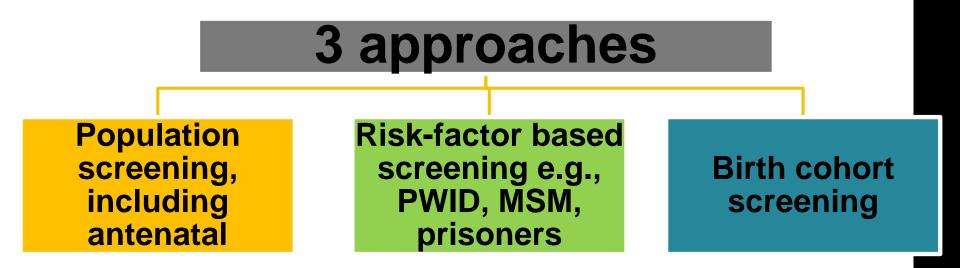
UNSAFE INFECTIONS

Unsafe medical practices and injection safety- as in Egypt and Cameroon-Implications for aggressive & targetted screening



Traditional practices including scarifications, tattoos

WHO TO SCREEN: WESTERN COUNTRIES



- No current universal screening program
- Risk based screening: population dependent
- Birth cohort in USA and Japan, ??? Sub-Saharan Africa

WHO TO SCREEN ?: SUB-SAHARAN AFRICA

Remains unclear

- Persons with persistently abnormal ALT levels.
- Recipients of transfusions (prior) to ???
- Persons with recognized occupational exposures e.g. HCWs
- Exposure to unsafe injection or medical practices
- Children born to HCV-positive women
- HIV positive persons
- ? Traditional practices
- Persons who ever injected illegal drugs

Therapeutic Options: Awareness, Screening, Testing

Barrier	What we need to do to access therapy and promote treatment scale-up
Lack of awareness	 Increase awareness of population /Health care worker /political Information, education, research, advocacy, civil society mobilization
Screening with anti- HCV serological tests NAT for HCV RNA needed for confirmation of viraemia	 Access to testing rural and urban population Integration into existing facilities Health system strengthening Prequalified screening tests Laboratory facilities for viral load and genotype Newer testing strategies –eg HCV Ag test GeneXpert HCV

BACKGROUND: NIGERIA, WEST AFRICA

- **Population :180 million**
- GDP per capita \$1,657
- 60% live below poverty line
- average HBsAg prevalence of 13.7% and anti-HCV prevalence of <2%
- 20 million persons currently infected with hepatitis B.
- 2.5 million persons with HCV



SOGHIN SOCIETY FOR GASTROENTEROLOGY AND HEPATOLOGY IN NIGERIA

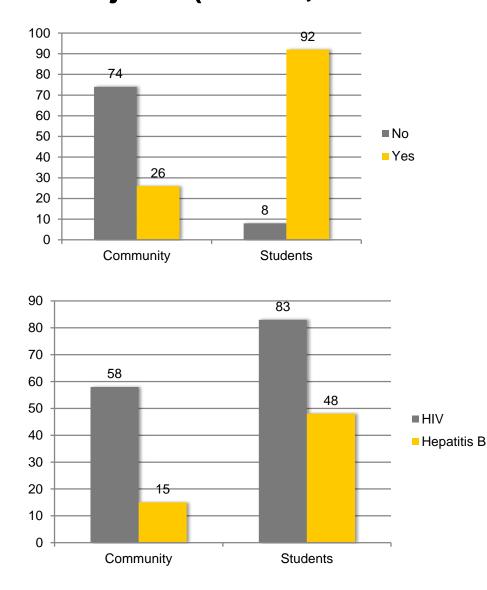


Best of EASL, Lagos, Nigeria, July 2016



Best of EASL, Lagos, Nigeria, July 2016

WHD 2016: community awareness and screening in 750 subjects (community 500 and Health Sciences students-150



Have you ever heard of viral hepatitis B or C?

Do you know your HIV status? Do you know your Hepatitis B o C status?

HCV REGISTRY PILOT-REAL WORLD DATA- NIGERIA 2016

- Commenced July 2015 with availability of generic Sofosbuvir in the Nigerian market
- Over 300 persons treated with sofosbuvir containing regimen to date
- Gastroenterologists/ hepatologists/ National GI professional society-
- 8 recruiting collaborators/sites to date

THE NIGERIAN HCV REGISTRY-PRELIMINARY REAL WORLD DATA- 2016

	N=80
Mean Age, y (range)	59 (40-60)
Male, n (%)	43 (52%)
Risk factor Prior surgery/instrumentation Prior Blood transfusion HIV infection/IVDU/MSM	19 (15%) 12 (9.4%) 0
Genotype 1 Genotype 2 Genotype 3 Genotype 4	62 (78%) 10 (12.5%) 2 8 (10%)
No cirrhosis Cirrhosis compensated Cirrhosis decompensated (child Pugh) score	48 17 10
Regimen type Sof/Peg +/-RIBA Sof/Led or Sof/Daclatasvir	65 10

LESSONS LEARNT FROM REGISTRY PILOT

- 1. "African Birth Cohort' effect
- 2. Significant history of exposure to blood transfusion, health care or hospital admission/surgical intervention etc
 - 1. latrogenic transmission, unsafe injection, blood safety,
- 3. HCV screening driven by clinical features of liver disease or abnormal liver function tests
- 4. Predominant HCV genotype 1
- 5. High cost of care-exceeding 4000 USD

consultation, HCV viral load ± genotype, liver enzyme, fibroscan etc) high end private health care, specialist liver care model. Mostly outof-pocket expenses

- 6. Predominant use of Sof/Peg/Riba (availability)-generic sofosbuvir and Increasing availability/ use of Ledipasvir & Daclatasvir combinations (generic)
- 7. Limitations

GLOBALLY APPROVED HCV DRUGS IN

2016

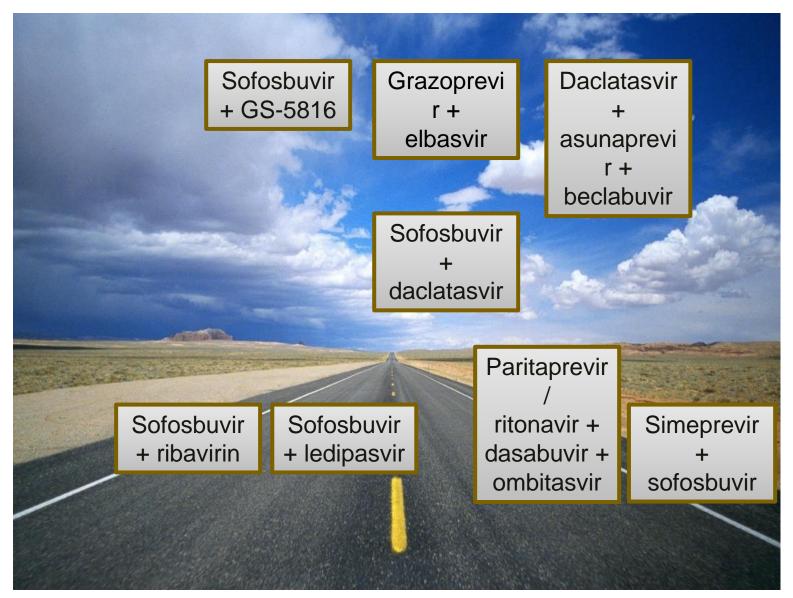
Product	Presentation	Posology
PegIFN-α2a	Solution for injection containing 180, 135 or 90 μg of PegIFN- $\alpha 2a$	Once weekly subcutaneous injection of 180µg (or less if dose reduction needed)
PegIFN-α2b	Solution for injection containing 50 μ g per 0.5 ml of PegIFN- α 2b	Once weekly subcutaneous injection of 1.5 µg/kg (or less if dose reduction needed)
Ribavirin	Capsules containing 200 mg of ribavirin	Two capsules in the morning and 3 in the evening if body weight <75 kg Or Three capsules in the morning and 3 in the evening if body weight ≥75 kg
Sofosbuvir	Tablets containing 400 mg of sofosbuvir	One tablet once daily (morning)
Simeprevir	Capsules containing 150 mg of simeprevir	One capsule once daily (morning)
Daclatasvir	Tablets containing 30 or 60 mg of daclatasvir	One tablet once daily (morning)
Ledipasvir/Sofosbuvir	Tablets containing 400 mg of sofosbuvir and 90 mg of ledipasvir	One tablet once daily (morning)
Paritaprevir/ombitasvir/ ritonavir	Tablets containing 75 mg of paritaprevir, 12.5 mg of ombitasvir and 50 mg of ritonavir	Two tablets once daily (morning)
Dasabuvir	Tablets containing 250 mg of dasabuvir	One tablet twice daily (morning and evening)

PEGASYS (PegIFN-α2a) Roche Products Limited, SmPC, November 2014; VIRAFERON-PEG (PegIFN-α2b), Merck Sharp & Dohme Limited. SmPC June 2014; COPEGUS (ribavirin) Roche Products Limited. SmPC, February 2015; REBETOL (ribavirin) Merck Sharp & Dohme Limited. SmPC May 2014; OLYSIO▼ (simeprevir), Janssen Products LP. SmPC, March 2015; Limited. DAKLINZA▼ (daclatasvir), Bristol-Myers Squibb Pharmaceutical. SmPC October 2014) HARVONI▼ (ledipasvir/sofosbuvir), Gilead Sciences Europe Ltd. SmPC, November 2014 VIEKIRAX▼ (ombitasvir/paritaprevir/ritonavir), AbbVie Ltd. SmPC, January 2015 EXVIERA▼ (dasabuvir), AbbVie Ltd. SmPC, January 2015; SOVALDI▼ (sofosbuvir), Gilead Sciences Europe Ltd. SmPC, March 2015; HARVONII▼ (ledipasvir/sofosbuvir), Gilead Sciences Europe Ltd. SmPC, March 2015; HARVONII▼ (ledipasvir/sofosbuvir), Gilead Sciences Europe Ltd. SmPC, March 2015; HARVONII▼ (ledipasvir/sofosbuvir), Gilead Sciences Europe Ltd. SmPC, November 2015; EASL Guidelines 2015; http://www.hep-druginteractions.org/ (Accessed May 2015)

TREATMENT GUIDELINES

INTERNATIONAL GUIDELINES VS LOCAL PRACTICE

HCV DAA Treatment Landscape



Therapeutic options: Financing access to care and drug therapy

DIRECT COSTS OF TESTS & DRUGS

	Cost (US \$)	Government subsidy/insur ance re- imbursement	The availability of oral DAA have revolutionized the the treatment of HCV with cure rates over 90% even in HIV infected subjects
HBV/HCV serology (5 markers)	30	Nil	Current costs of sofosbuvir,
Biochemistry/ liver enzymes (panel)	40-50	Nil	Per person (12 weeks) 90000 \$84,000
Ultrasound scan CT scan	30-50 200	NIL NIL	80000 · 70000 ·
HBV DNA Viral load HCV RNA Viral load	240-450 240-450	NIL NIL	€ 60000 · \$55,000 553,000 · \$53,000 553,000 · \$53,000
Pegylated interferon (monthly)	800-950	NIL	ون 40000 ·
Sof/ledipasvir (Generic)	25-50	NIL	30000 · \$27,000 20000 ·
Sof/Daclatasvir		NIL	10000 · \$8,000 \$900 \$450 \$100

USA*

USA

UK

Spain

Generic Mini

LMICs

Brazil

KEY CHALLENGES CONFRONTING SSA COUNTRIES THE FIVE "LACKS"



Information gaps on disease and economic burden, treatment need, and potential impact of treatment scale-up (to build investment case)

Not enough advocacy, leadership and commitment for hepatitis response at higher levels

Continued limited global and country funding

Lack of national strategies and plans and dedicated hepatitis department/focal person within MOH in many countries

Public health sector approach to hepatitis treatment still in its infancy and Lack of case finding (testing approaches)

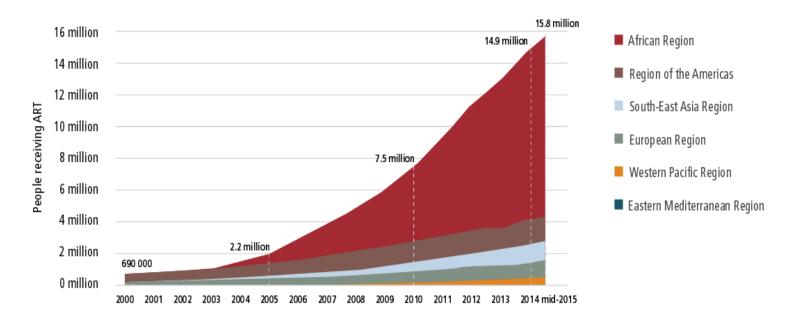
High cost of DAA drugs for HCV

ENHANCING THERAPEUTIC OPTIONS

Appropriate model of care to improve access

- Individualized or public health approach
- Vertical system (like HIV) vs integrative into existing program
- Lessons learnt Public health approach successful in the control of HIV in Africa and likely to succeed in Hepatitis, Lessons from Egypt

Estimated numbers of people receiving antiretroviral therapy globally and by WHO Region and percentage coverage globally, 2000–2015



ENHANCING THERAPEUTIC OPTIONS

Health care financing

- Increased government budget
- Shared costs with other strategies
- Innovations and increased efficiencies

Drug access and funding in RLS

- Generic versions of DAA
- Tiered Pricing
- Government funding/Donor funding
- Treatment subsidization as available for HIV/AIDS, malaria and TB through international donors (Global Funds, PEPFAR, AU, others)

SUMMARY

- Significant disease burden
- Availability of curative therapy for HCV has converted hepatitis care into an emergency requiring urgent responses.
- Despite the therapeutic options, challenge of data hampers disease awareness and recognition.
 Provision of screening and testing and care needed within an integrated public health system
- There is need for a 360 degree response including government commitment and collaboration, strengthening African networks and collaborations, international collaborations (donors, pharmaceuticals) health insurance and HMOs, civil society, academia.





Acknowledgements: Hepatitis B Registry team, SOGHIN

THANK YOU FOR YOUR ATTENTION