

## How to control HBV in high endemic regions: Asia

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- Consultation: BMS, Gilead, GSK, Novartis, MSD and Roche
- Speaker: BMS, Gilead and GSK

## **Outlines**

- Changing epidemiology of HBV infection in Asia
- Universal HBV vaccination in infants in Asia
- Prevention of HBV MTCT in Asia
- Treatment of CHB in Asia
- Towards elimination of HBV by 2030





## GLOBAL HEALTH SECTOR STRATEGY ON VIRAL HEPATITIS 2016–2021

#### TOWARDS ENDING VIRAL HEPATITIS





Approved at the Sixty-ninth World Health Assembly 23–28 May 2016



## Regional Action Plan for Viral Hepatitis in the Western Pacific 2016–2020

- HBsAg <1% in children under 5-year old, further drop in MTCT</li>
- HepB vaccination in health care workers and medical/health students
- Safe injection
- HBV/HCV, 30% diagnosed, 50% treated, 90% virus suppressed











#### HBsAg Endemicity and HBV genotypes distribution in Asia (1990–2013)



Tian Q, et al. Hepatol Int 2016

Edmunds WJ et al., Epidemiol Infect 1996

# Incremental approach to prevention of HBV infection at birth and in the first years of life





The interventions at the base of the pyramid benefit the largest number and are necessary for those at the top of the pyramid to be effective

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### Infants receiving 3 doses of hepatitis B vaccine by WHO region, 1989–2014



#### WHO, World Health Statistics 2016

### Chronic HBV infection rates in children before and after HBV vaccine introduction, WHO's Western Pacific Region



Hennessey K, et al. Vaccine 2013; 31 (Suppl 9)

### Strategies and achievements for HBV control in Taiwan

- **1965** Australia antigen discovered
- **1984 Universal HBV vaccination for newborns**
- **1997** The incidence of HCC in children decreased
- **2003** Reimbursement of anti-HBV therapy
- 2006 REVEAL-HBV cohort study
- 2011 REACH-B study (community-based natural history study)
- **2012** HBsAg positive rates in the children decreased to 0.9%
- 2013 ERADICATE-B-cohort study (hospital based natural history)
- 2013 Upgraded REVEAL-HBV risk calculator for HCC
- 2014 C-TEAM cohort study Population-based study
- 20?? HBV eradication

Lin CL, et al. J Formos Med Assoc 2015; 14:901-9

#### HCC for Birth Cohorts Born Before and After HBV vaccination Program in 1984 in Taiwan



Chiang CJ, et al. JAMA. 2013; 310:974-6.

# CLD&HCC mortality declined since the launch of viral hepatitis therapy program in 2003 in Taiwan



Chiang CJ, et al. Hepatology 2015;61:1154-62.

### **Changing HBsAg Endemicity in China mainland**

#### 1957-1989

#### 1990-2013

![](_page_14_Figure_3.jpeg)

Schweitzer A, et al. . LANCET 2015. WHO. Global Immunization Data , July 2015 http://www.who.int/immunization/monitoring\_surveillance/Global\_Immunization\_Data.pdf?ua=1

### Prevalence of HBsAg and HepB vaccine coverage for each birth cohort in China mainland

![](_page_15_Figure_1.jpeg)

Cui F, et al. Emerg Inect Dis 2017; 23:765-772

# Coverage of HepBOT in the Eastern, Central and Western China 2002–2009, 2010 survey

![](_page_16_Figure_1.jpeg)

## **Declining HBsAg Prevalence in China mainland**

![](_page_17_Figure_1.jpeg)

HBsAg in general population: 9.75%(120 m)→7.18%(93 m)→ 6%? (70 m ?)

Cui FQ, et al. EID 2017 Cui Y, et al. JGH 2013

#### HBV vaccination reduces the HCC incidence in eastern China

0.77(0.72-0.83) 1.00(0.92-1.09)

1.4(1.2-1.6)

1.1(.93-1.3)

45-49

50+

98(.84-1.14)

40-44

3.4(1.9-6.0)

20-24

10-19

1.9(1.4-2.5)

25-29

1.14(94-1.4) .99(83-1.2)

Age Group (years)

35-39

30-34

Cumulative probability of HCC in the vaccinated and control cohorts.

![](_page_18_Figure_2.jpeg)

Sun Z, et al. Carcinogenesis 2013; 34:1800-05.

### Achievement of WHO/WPRO target of hepatitis control

#### **WPRO target:** HBsAg+ under 5

- 2012: below 2%
- 2017: below 1%

### China achieved:

- 2006: 0.96%
- 2014: 0.32%

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![](_page_19_Picture_8.jpeg)

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![](_page_21_Figure_1.jpeg)

### Prevention of MTCT of HIV, Syphilis and Hepatitis B in China

![](_page_22_Figure_1.jpeg)

He T, et al. Liver Int. 2016;36(Suppl 1):105-8.

### Increased rate of baby deliveries in health facility

![](_page_23_Figure_1.jpeg)

#### 2012 Neonatal deliveries in china

2013 Reported HBV vaccination coverage

![](_page_23_Figure_4.jpeg)

WHO WPRO: Hepatitis B Control Country Profiles 2013

#### NUs used in 3rd trimester of pregnancy: a multi-centre, RCT Study on TDF

![](_page_24_Figure_1.jpeg)

Zhang H, et al. Hepatology 2014;60:468-76

25

### **The Shield Project-Vision and Goals**

Vision :

为了没有乙肝的未来

## Towards A Hepatitis B-Free Future

**Goals** :

- Hundred: engage 100 project hospitals
- Thousand: involve 1000 doctors
- Ten thousands: serve 10000 HBsAg+ pregnant women

![](_page_25_Figure_8.jpeg)

![](_page_25_Picture_9.jpeg)

**Shield Project** 

A hepatitis B-free generation in China

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### rends in the number and reporting quality of the publications:

![](_page_27_Figure_1.jpeg)

Zeng N, et al. Int J Infect Dis 2018; 67:58–64

Cleveland Clinic

![](_page_28_Figure_0.jpeg)

## Antiviral therapy improves the clinical outcomes in HBV-compensated cirrhosis

![](_page_29_Figure_1.jpeg)

Wong GL, et al. Hepatology, 2013, 58:1537-47.

Wu S, et al. Liver Int. 2017

#### Effective viral suppression is necessary to reduce HCC development in 120 cirrhotic patients with CHB: Results of a 10-year follow up

![](_page_30_Figure_1.jpeg)

Zhang W, et al. Medicine 2017; 96:44(e8454) Jia J, et al. JGH 2017

# Guidelines on the prevention and treatment of CHB (2005, 2010, 2015)

![](_page_31_Picture_1.jpeg)

➤Training courses ( 2015~2018) for doctors in secondary hospitals

- ▶31 provinces
  - ~ 60 cities
  - ~2000 hospitals,
  - ~9000 doctors

![](_page_31_Picture_7.jpeg)

![](_page_31_Figure_8.jpeg)

## 国家卫生计生委药物政策与基本药物制度司

![](_page_32_Figure_1.jpeg)

#### 国家卫生计生委办公厅关于公布国家药品价格谈判结果的通知

中华人民共和国国家卫生和计划生育委员会 2016-05-20

#### TDF 1500 RMB/Mon → 490 RMB/Mon 国卫办药政函〔2016〕515号

市卫生计生委,新疆生产建设兵团卫生局:

药品价格谈判结果印发你们,请按照国家卫生计生委等7部门印发的《关于做好 构的通知》(国卫药政发〔2016〕19号)有关精神抓紧落实。

底前完成谈判药品集中挂网。在2016—2017年的采购周期内,不再另行组织谈 构和使用谈判药品。

宣侣验合规费用范围的地区,要进一步巩固完善医保制度和支付

2015年国家药品价格谈判结果

	通用名	商品名	生产企业	包装规格	谈判价格
4	富马酸替诺福韦二吡呋酯	韦瑞德	葛兰素史克(中国) 投资有限公司	300mg×30片/瓶	490元
	盐酸埃克替尼	凯美纳	贝达药业股份 有限公司	125mg×21片/盒	1399元
	盐酸吉非替尼	易瑞沙	阿斯利康制药 有限公司	250mg×10片/盒	2358元

![](_page_32_Picture_11.jpeg)

### Medicine is not luxury item. 药品不是奢侈品

![](_page_32_Picture_13.jpeg)

### Preliminary Data from China-Registry of Hepatitis B (CR-HepB) Increasing use of the first-line NAs for CHB (2003-2016)

![](_page_33_Picture_1.jpeg)

![](_page_33_Figure_2.jpeg)

Shan S, et al. APASL 2017

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## **Estimation of HBV-CLD in China mainland**

![](_page_35_Figure_1.jpeg)

### Scale-up therapy is need to reduce deaths due to HBV-HCC in China

![](_page_36_Figure_1.jpeg)

Source: WHO China (preliminary results)

# Summary

- 1. MTCT and early childhood transmission play a key role in HBV high endemicity in Asia
- 2. Universal hepB vaccination could dramatically reduce HBsAg in the young age group in Asia
- 3. Timely birth dose of hepB vaccine & HBIG are essential to prevent MTCT in HBeAg+ mothers
- 4. NA therapy at late pregnancy could further reduce MTCT in mothers with high viral load
- 5. Scale up of effective antiviral therapy could reduce cirrhosis, decompensation and HCC

![](_page_38_Picture_0.jpeg)