

#### Hepatitis B

# Hepatocellular Carcinoma Screening

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Last Updated: June 16, 2023

#### Disclosures

H. Nina Kim, MD MSc

Gilead Sciences provides program funding to my institution

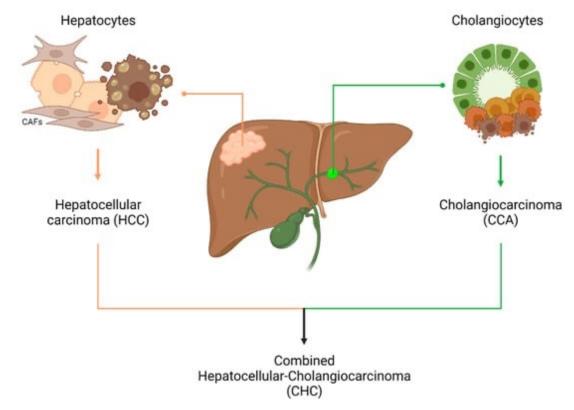


# Hepatocellular Carcinoma (HCC) due to Chronic Hepatitis B

- Terminology
- HCC epidemiology global and US
- Distinct oncogenic risk related to hepatitis B (HBV)
- Evidence to support HCC screening
- Who to screen
- How to screen



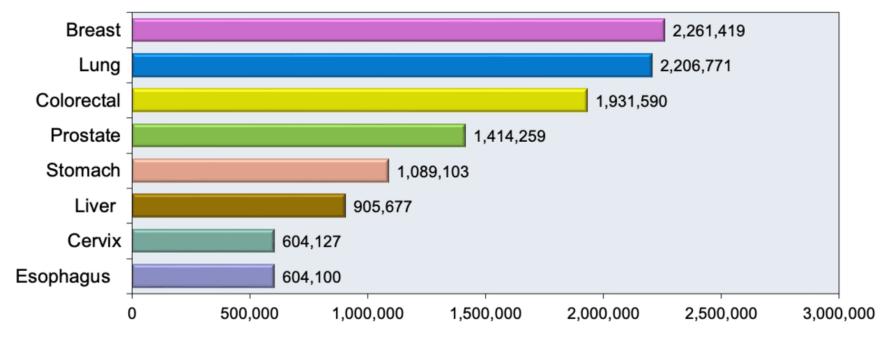
# Terminology of primary liver cancer





# Liver Cancer – Among the leading cancers worldwide

World Estimated Number of New Cancers in 2020, Both Sexes, All Ages

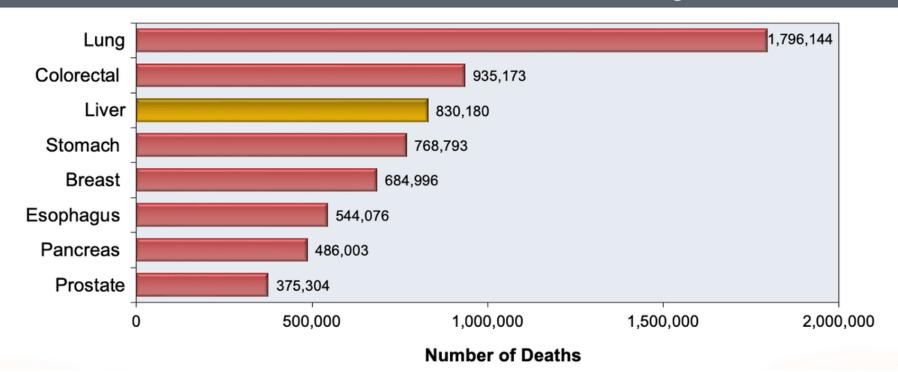


**Number of People with Cancer** 

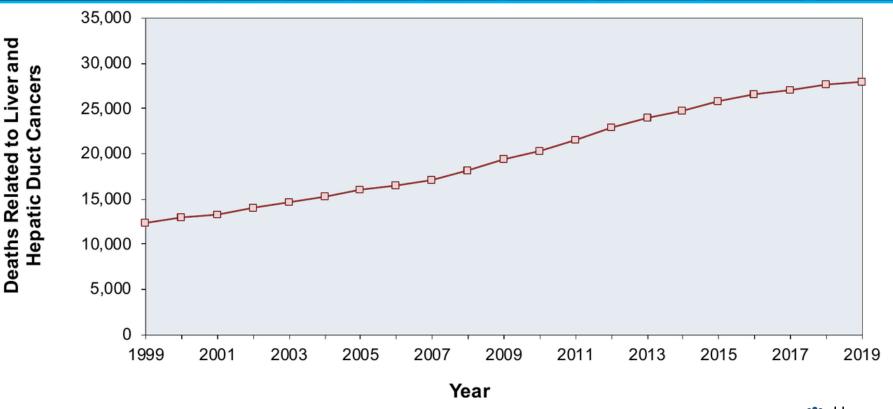


# Liver Cancer – 3<sup>rd</sup> leading cause of cancer death

World Estimated Number of Cancer Deaths in 2020, Both Sexes, All Ages



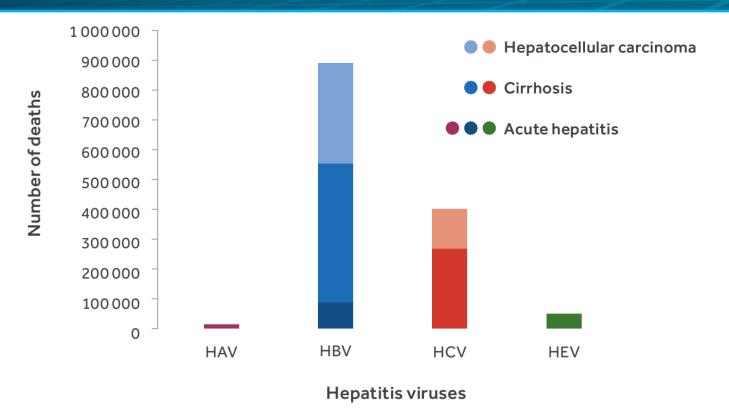
# HCC-related deaths, 1999-2019 in US



Source: U.S. Cancer Statistics Working Group. U.S. Cancer Statistics Data Visualizations Tool, based on 2021 submission data (1999-2019): U.S. Department of Health and Human Services, Centers for Disease Control and Prevention and National Cancer Institute. Released June 2022.

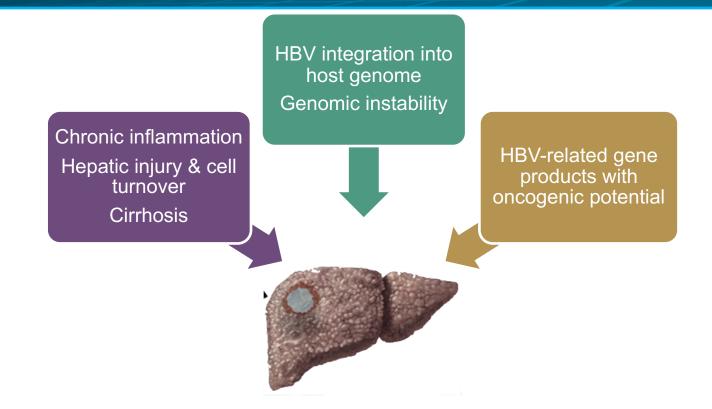


#### Deaths from Viral Hepatitis: Role of HBV-related HCC



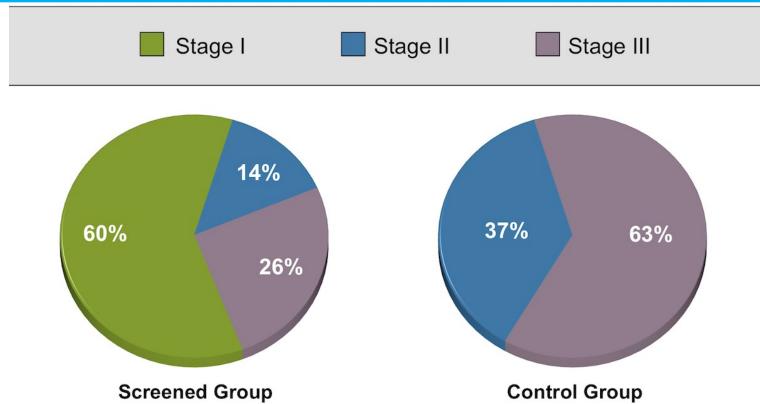


### Mechanisms for HBV Oncogenesis





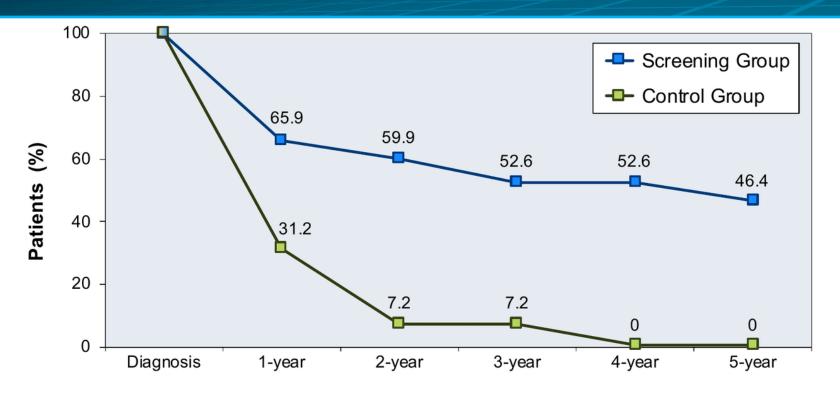
# Cluster randomized trial of HCC screening





Source: Zhang et al, J Cancer Res Clin Oncol. 2004;130:417-22.

### Cluster randomized trial of HCC screening



Survival after Diagnosis of HCC



### Who to Screen for HCC in Persons with chronic HBV (AASLD)

- All persons with cirrhosis
- Black males older than 40 years of age
- Asian males older than 40 years of age
- Asian females older than 50 years of age
- Persons with 1<sup>st</sup> degree family member with history of HCC
- Persons with hepatitis delta coinfection



### Risk Factors for HCC

Host	Viral/Disease	Environmental/Exposure
<ul><li>Age &gt;40 years</li><li>Male sex</li><li>Immune compromised</li></ul>	<ul> <li>Presence of cirrhosis</li> <li>High HBV DNA level (&gt;2000 IU/ml)</li> <li>Genotype C</li> <li>Longer duration of infection</li> <li>Concurrent viral infections (HIV, HCV, HDV)</li> </ul>	<ul> <li>Alcohol</li> <li>Metabolic syndrome (NAFLD, diabetes, obesity)</li> <li>Aflatoxin</li> <li>Smoking</li> </ul>



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# How to Screen: Abdominal Ultrasound every 6 months

#### Advantages

- Inexpensive
- No radiation or IV
- Readily available
- Well tolerated

#### Limitations

- Operator dependent
- Body habitus, hold breath
- Liver factors (nodularity, steatosis)
- Sensitivity ranges from 40-80%



Photo: https://ethnomed.org/resource/abdominal-ultrasound



### Biomarker for HCC Screening

- Serum alpha fetoprotein (AFP)
  - Advantages: Readily available, relatively inexpensive, easy to perform
  - Limitations: Suboptimal sensitivity & specificity
    - (1) Lack of uniform secretion of AFP by HCC tumors, esp. tumors <3 cm<sup>2</sup>
    - (2) Fluctuating and elevated AFP levels in patients with viral hepatitis or advanced fibrotic liver disease without HCC
  - Debate over benefit of adding AFP to ultrasound

	EASL	AASLD	APASL
Biomarker	Not recommended	At discretion of clinician	AFP (+ US)



### Summary

- HCC is a leading cause of cancer-related deaths worldwide, and much of it is HBV-related.
- Overall 5-year survival of HCC in the US is ~20%. Survival depends on the stage of HCC at the time of diagnosis.
- The primary goal of HCC surveillance is to detect disease in an early stage and therefore increase the likelihood of potentially curative therapy.
- Cirrhosis remains a consensus indication for HBV screening. Other indications per AASLD include factors determined by age/race, family hx, delta coinfection.
- The recommended HCC screening method is hepatic ultrasound every 6 months, with or without serum alpha-fetoprotein.









Hepatitis B Primary Care Guidance



HBV Medications >



HBV Vaccines >



Clinical Challenges



Tools & Calculators



#### **Hepatitis B Online**

A free educational website from the University of Washington Infectious Diseases Education & Assessment (IDEA) program



Funded by

Centers for Disease Control and Prevention (CDC)

#### **Course Modules**

Quick Access to Course Information Self Study with Progress Tracker Free CME and CNE/CE

Start the Self-Study Modules »



#### **Hepatitis B Virus**

#### **HBV Epidemiology**

Reviews United States and global HBV incidence and prevalence, populations at risk for HBV acquisition, and the clinical and laboratory criteria for HBV case definitions.

#### Quick Reference >

Rapidly access info about Epidemiology

#### Self-Study CNE/CME

Track progress and receive CE credit

#### **HBV Screening and Diagnosis**

Details the groups considered at priority for HBV testing, the recommended screening and diagnostic tests, and how to interpret HBV diagnostic test results.

#### Quick Reference >

Rapidly access info about Screening and Diagnosis

#### Self-Study CNE/CME

Track progress and receive CE credit

#### **HBV** Immunizations

Identifies indications for HBV vaccine, describes dosing schedules and administration of

#### Quick Reference >

Rapidly access info about Immunization

#### Self-Study CNE/CME

Track progress and receive CE credit

### Acknowledgments

**Hepatitis B Online** is funded by a cooperative agreement from the Centers for Disease Control and Prevention (CDC-RFA- PS21-2105). This project is led by the University of Washington Infectious Diseases Education and Assessment (IDEA) Program.





