Hepatitis B is a Treatable Disease!

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Prevalence of Hepatitis B Role of Immigration in Europe



The Netherlands



Annelot

David

HBV Disease: Silent Killer



Hepatitis B under-diagnosed and under-treated



Kim, Hepatology 2004

Hepatitis B: 1990

A preventable disease!

Treatment of Hepatitis B 1990-2010



Adapted from: ClinicalCareOptions.com

* Specific countries only

Goals for treatment

Long-term clinical goals

- Keep patient healthy for as long as possible
- Prevent hepatic decompensation
- Prevent progression to cirrhosis and liver cancer
- Prolong survival

Fung S, Lok ASF. Clinical Gastroenterol Hepatol. 2004;2(10):839–848

Phases Of Infection



What end points are used?

- HBeAg seroconversion
- Suppression in HBV DNA levels
- Histological improvement
- ALT normalization
- HBsAg seroconversion
- Studies have linked high HBV DNA levels with increased risk of:
 - Development of HCC and cirrhosis
 - Disease progression

Keeffe EB, et al. Clin Gastroenterol Hepatol 2006, 4: 936-62. The EASL Jury. J Hepatol. 2003; 39:S3–S25 Chen CJ, et al. JAMA 2006; 295:65–73. Iloeje UH, et al. Gastroenterology 2006; 130:678–686

Treatment Options



Concepts of Treatment Goals in Hepatitis B

Sustained response

No need for antiviral drugs

IMMUNE CONTROL

Pegylated Interferon

Maintained response

Continued need for antiviral drugs

VIRAL CONTROL

Nucleos(t)ide Analogues

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Response to PEG-IFN 6 months post treatment



Treatment duration 48 weeks

Lau, NEJM 2005; Janssen, Lancet 2005; Marcellin, NEJM 2004.

Follow-up of PEG-IFN α-2b in HBeAg (+) CHB: 3 years post-treatment among HBeAg responders



IFNα-2b Treatment is Associated with Prolonged Survival

Proportion of patients surviving



v Zonneveld et al. Hepatology 2004

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Nucleos(t)ide Analogues

Improving the long-term health of HBV patients with NA rests on these pillars



Resistance rates through 6 years among nucleos(t)ide-naïve patients

Entecavir 5-year efficacy data

HBeAg(+) ETV long-term cohort (ETV-022 \rightarrow ETV-901)

Tenofovir 3-year efficacy data

Adapted from 1. Marcellin P et al. N Engl J Med 2008;359:2442–55. 2. Heathcote EJ et al. 60th AASLD Oct 30–Nov 3, 2009; Boston, USA. Poster 483. Available at http://www.natap.org/2009/AASLD/AASLD_35.htm. (Accessed April 2010). 3. Marcellin P et al. 60th AASLD Oct 30–Nov 3, 2009, Boston, USA. Poster 481. Available at http://www.natap.org/2009/AASLD/AASLD_36.htm (Accessed April 2010).

Antiviral treatment delays disease progression

Patients with CHB and cirrhosis or advanced fibrosis

Time to disease progression (months)

Who Should be Treated and With What? PEG-IFN or Nucleoside Analogues?

Balancing the facts

Conclusions

- Major improvement in HBV therapy in last decades!
- Choice of therapy depends on individual patient characteristics: PEG-IFN in selected proportion of patients
- Choose most potent NA with highest genetic barrier
- Therapy with NA may be indefinite in many patients
- More than 95% of HBV in remission with current drugs

Issues for the Future

- Safety & efficacy of longterm therapy
- Response prediction: individualized therapy
- Role of combination therapy
- How to induce long-term cure (HBsAg loss)
- New agents with different sites of action

Hepatitis B: 2010

A treatable disease!