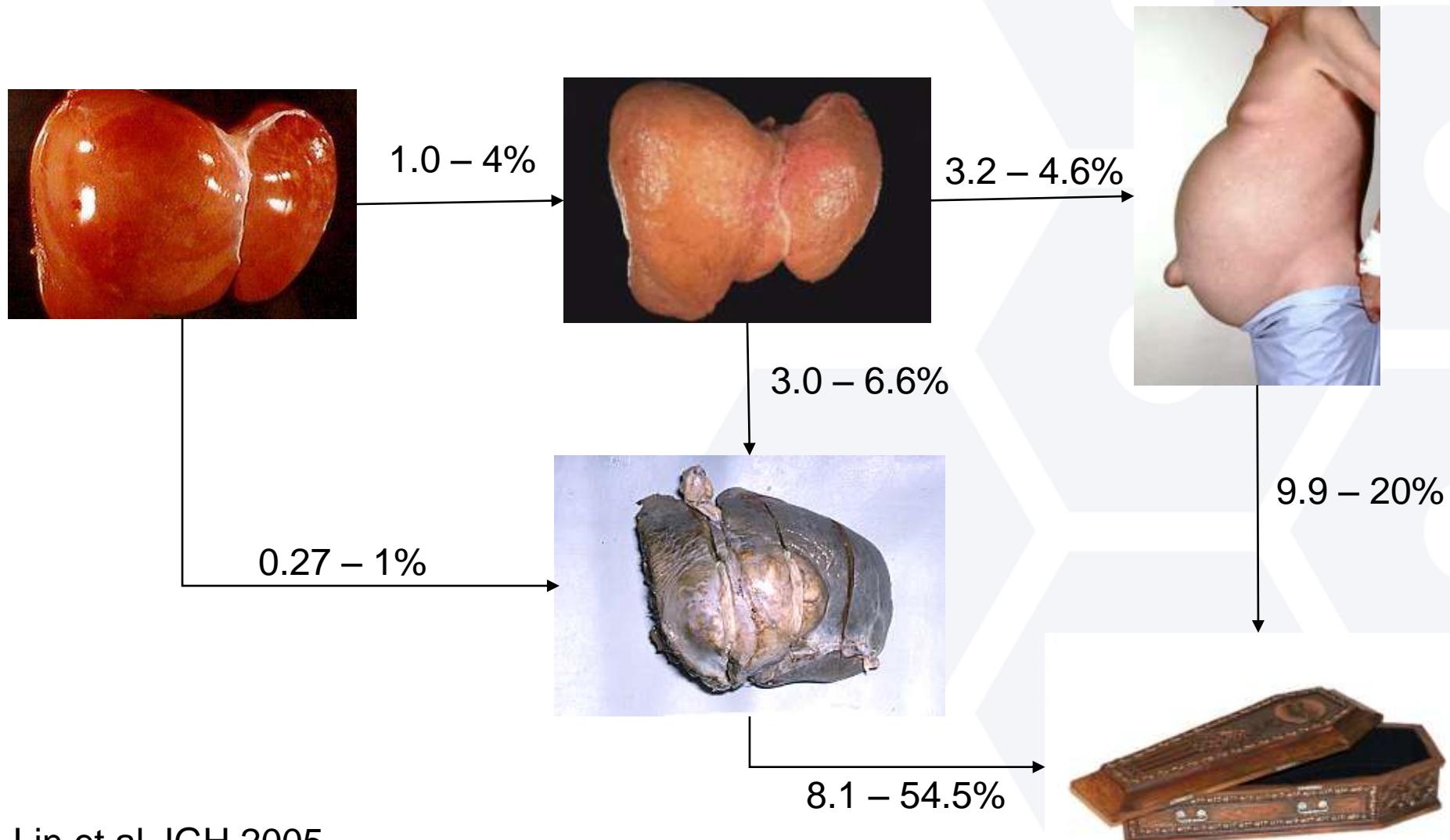


Summit Conference Hepatitis B & C

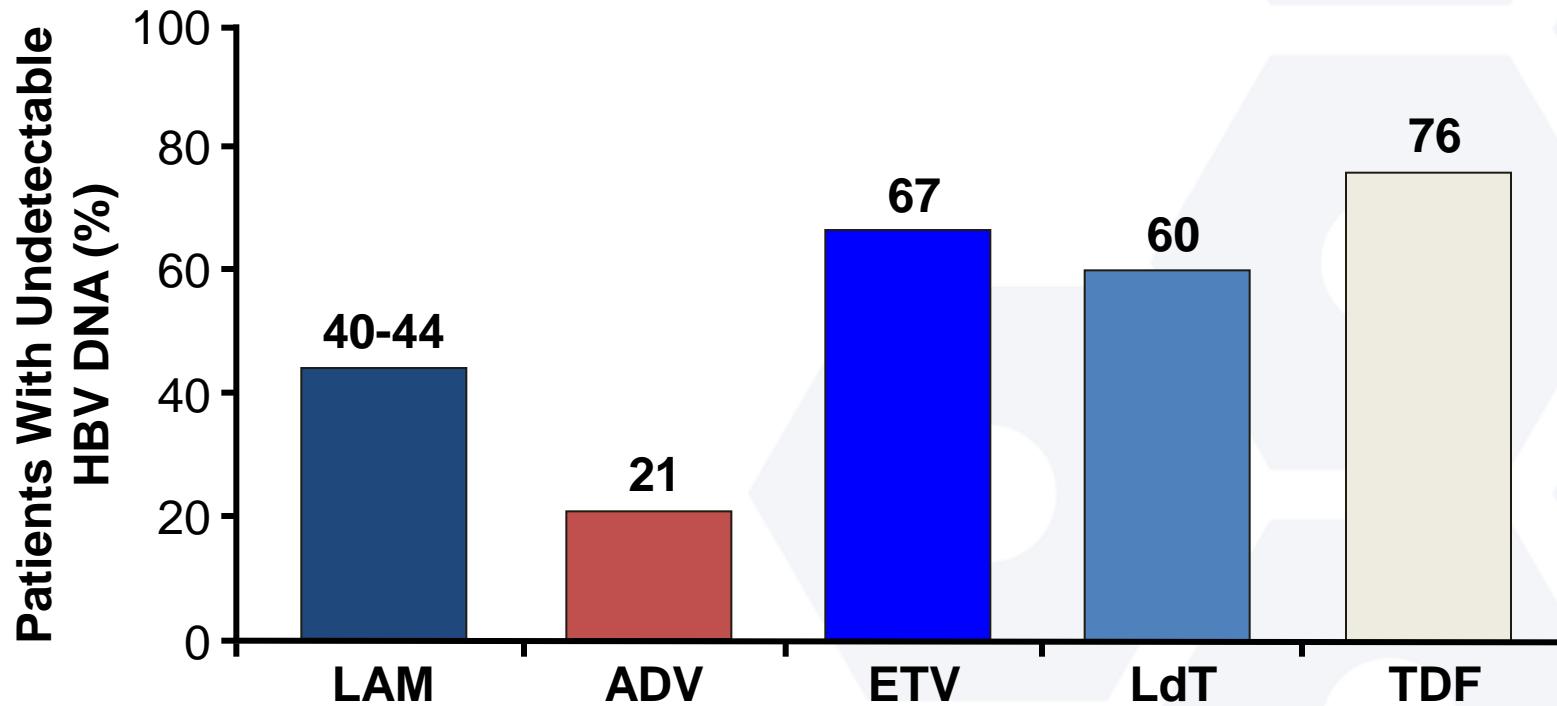
Advanced Liver Diseases Are Treatable

Mark Thursz
Vice-Secretary EASL

Progression of Disease in HBV

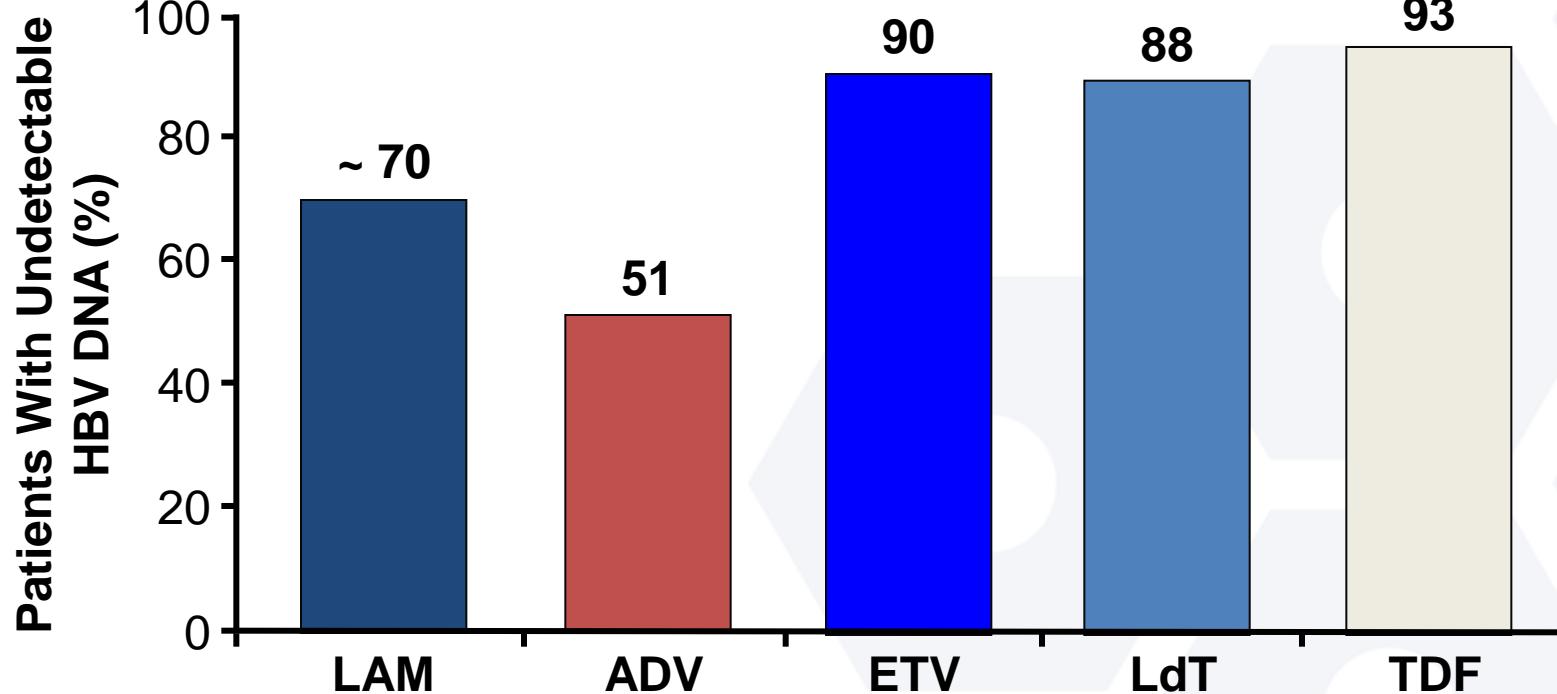


Virologic Response in HBeAg+ Patients (Undetectable HBV DNA at Wk 48-52)



Adapted from Lok AS, et al. Hepatology. 2007;45:507-539.
Heathcote EJ, et al. AASLD 2007. Abstract LB6.

Virologic Response in HBeAg- Patients (Undetectable* HBV DNA at Wk 48-52)



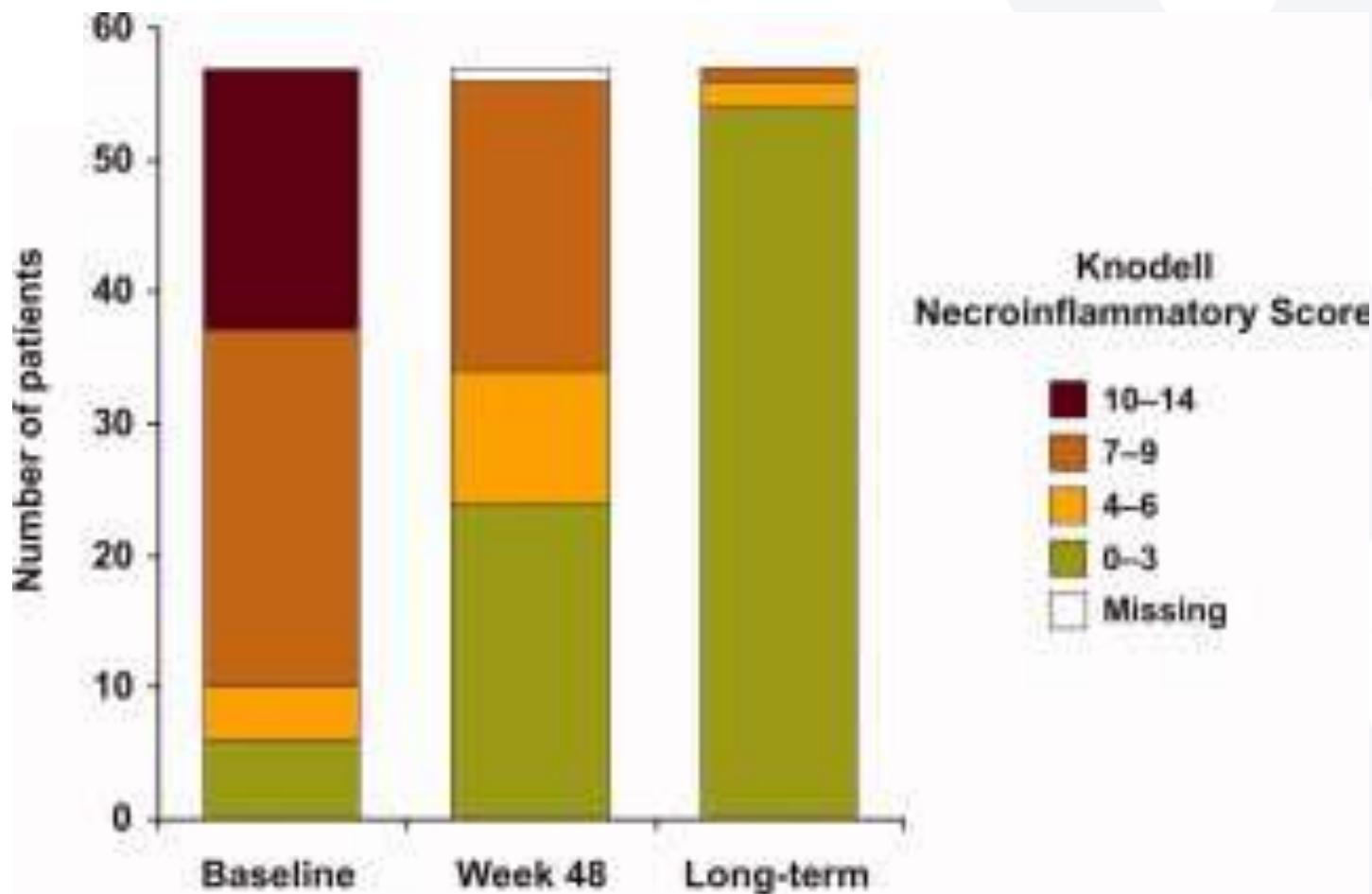
Adapted from Lok AS, et al. Hepatology 2007;45:507-539.
Marcellin P, et al. AASLD 2007. Abstract LB2.

Key questions

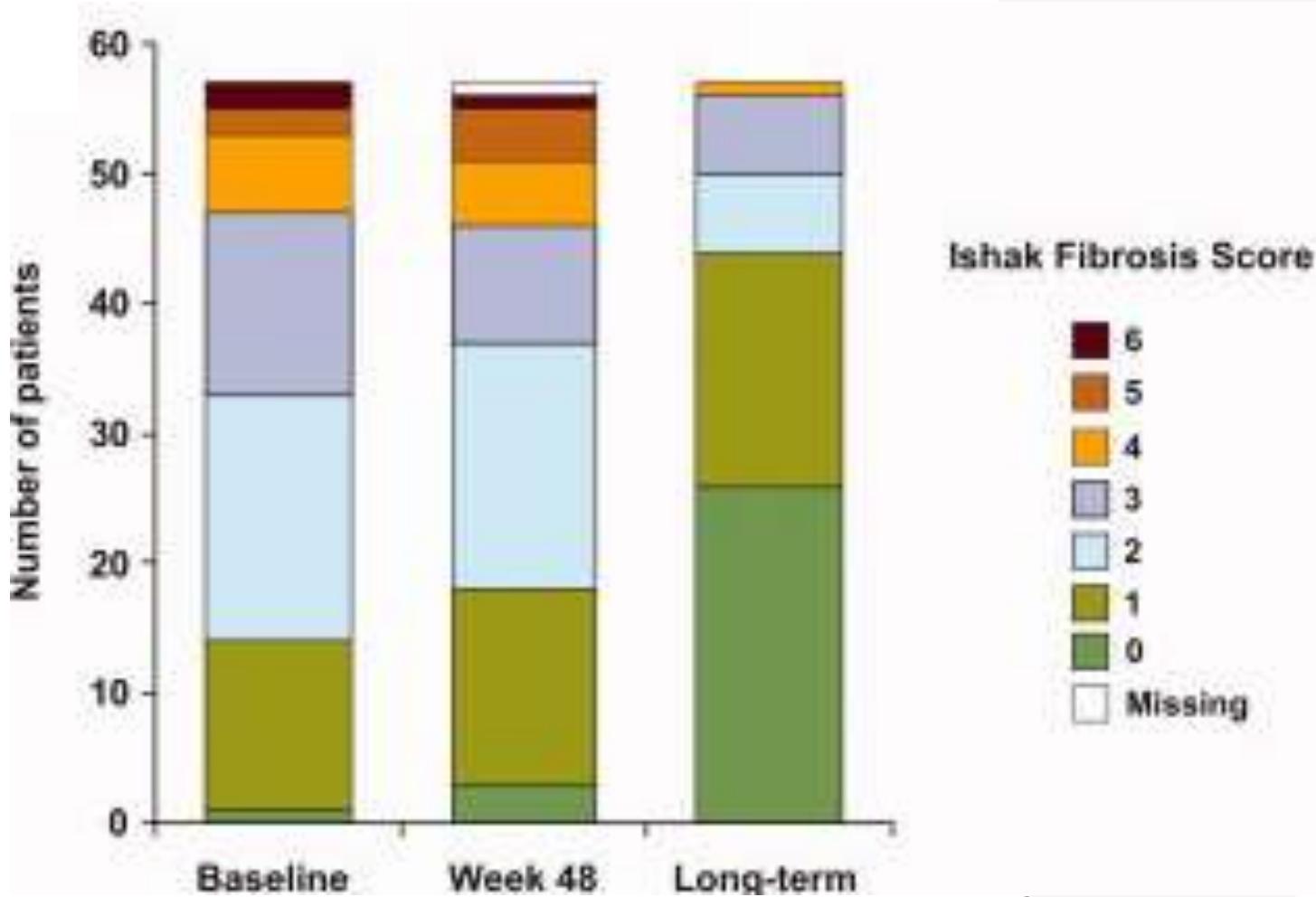
- Does treatment improve the liver histologically?
- Does treatment reverse the progression of liver disease?
- Does treatment prevent progression of cirrhosis to decompensation?
- Does treatment prevent progression of cirrhosis to cancer?
- Does treatment prolong life?

Impact of Treatment on Histology

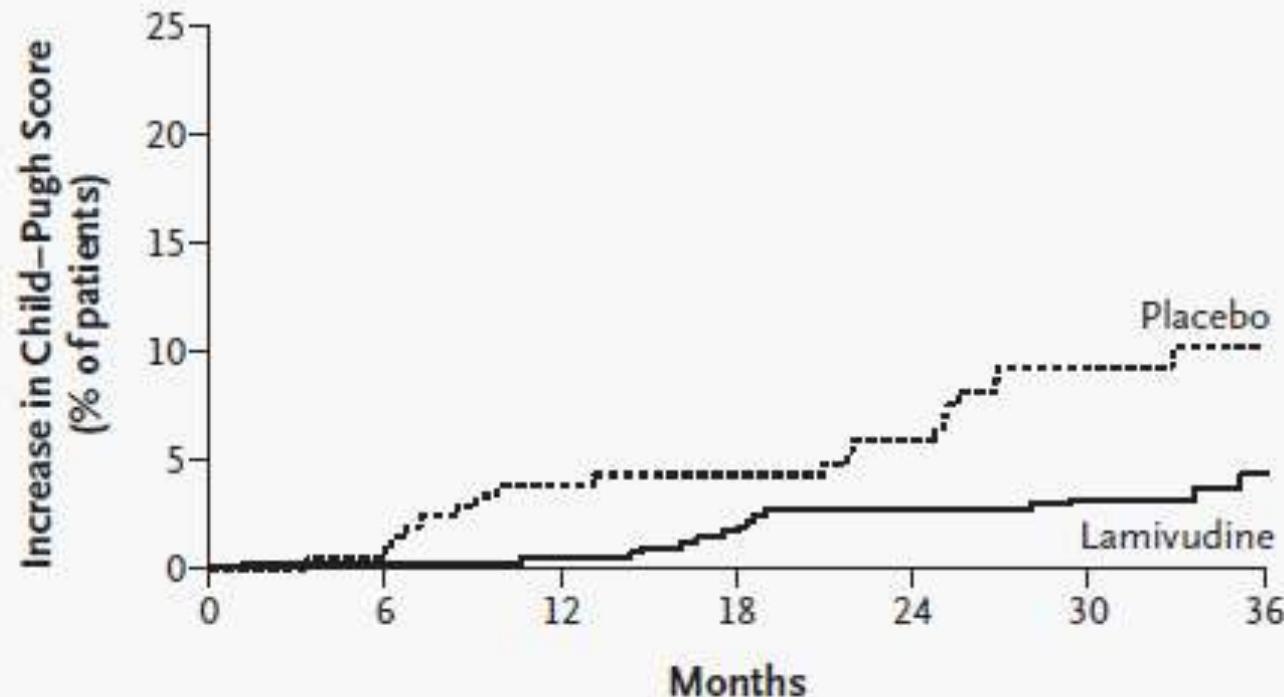
Inflammation



Impact of treatment on Histology Fibrosis



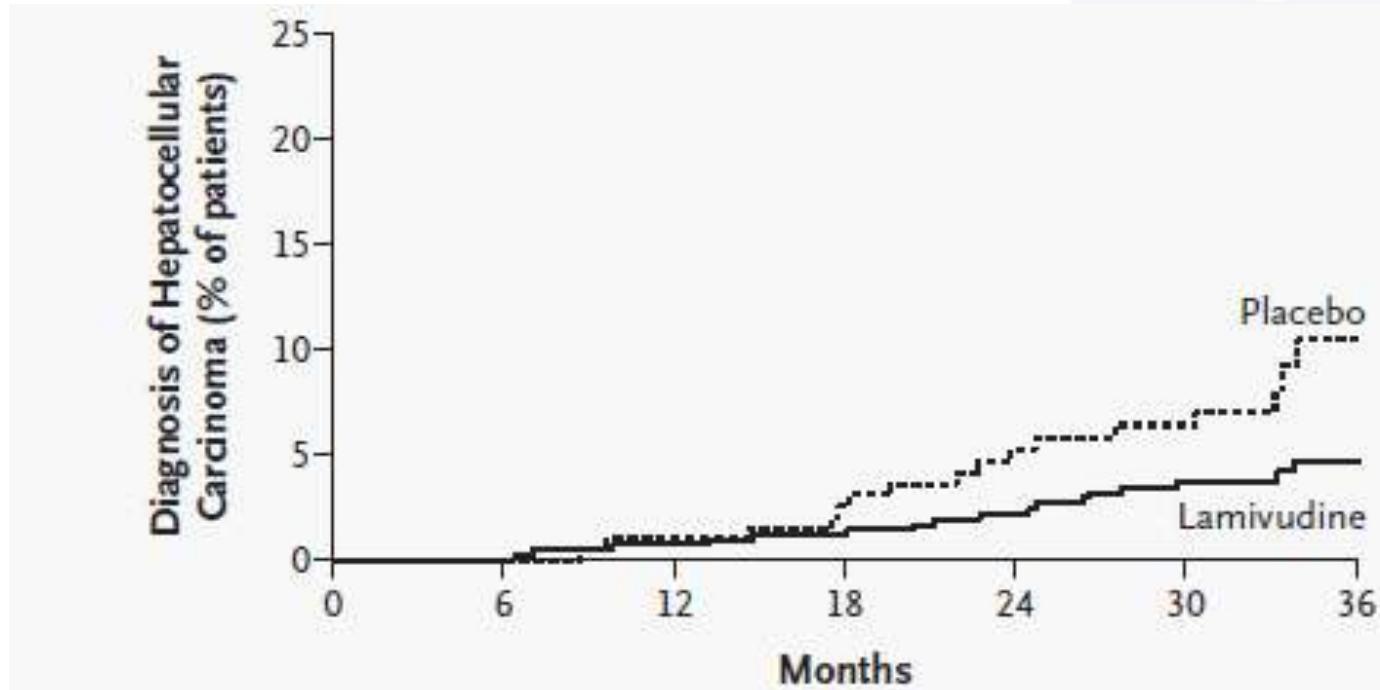
Impact of Treatment on Progression Cirrhosis - Decompensation



No. at Risk

Placebo	215	209	198	184	173	153	43
Lamivudine	436	429	417	400	385	347	122

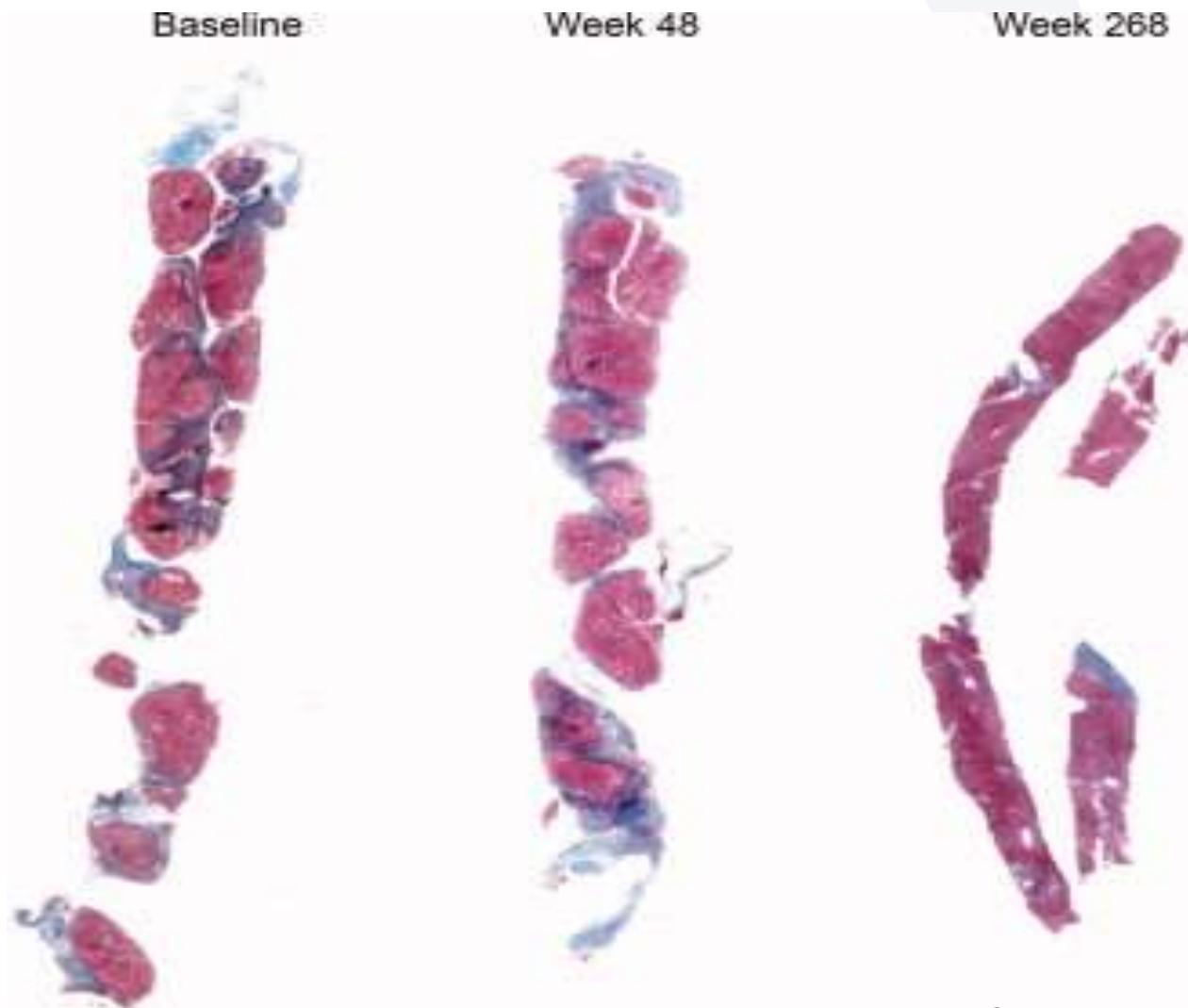
Impact of Treatment on Progression Cirrhosis - Cancer



No. at Risk

	6	12	18	24	30	36
Placebo	215	209	198	184	173	153
Lamivudine	436	429	417	400	385	347

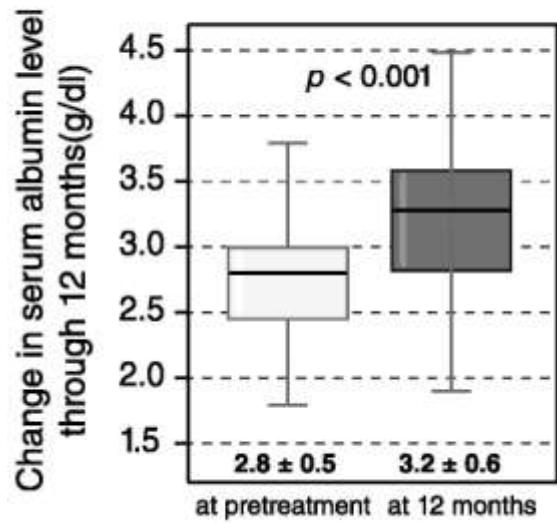
Reversal of Cirrhosis



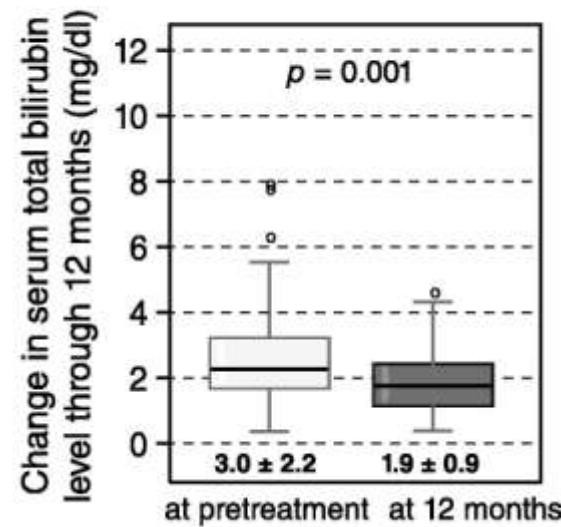
Chang. Hepatology 2010

Treatment of Decompensated HBV

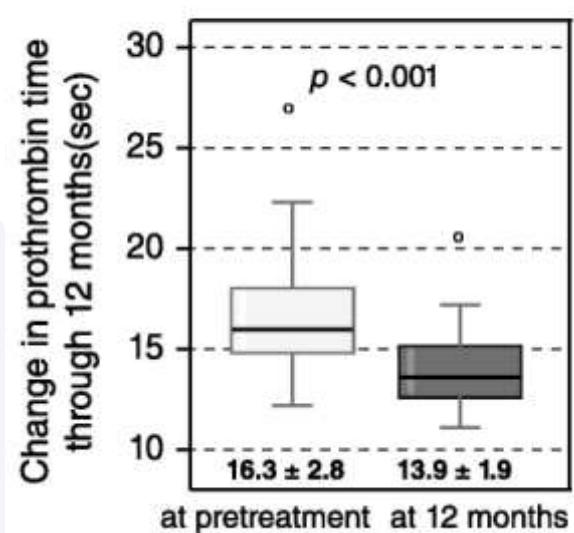
A



B

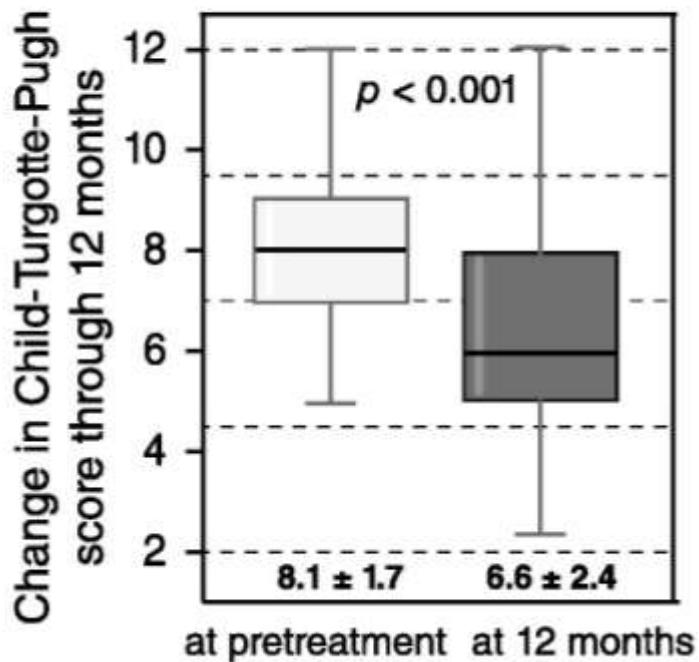


C

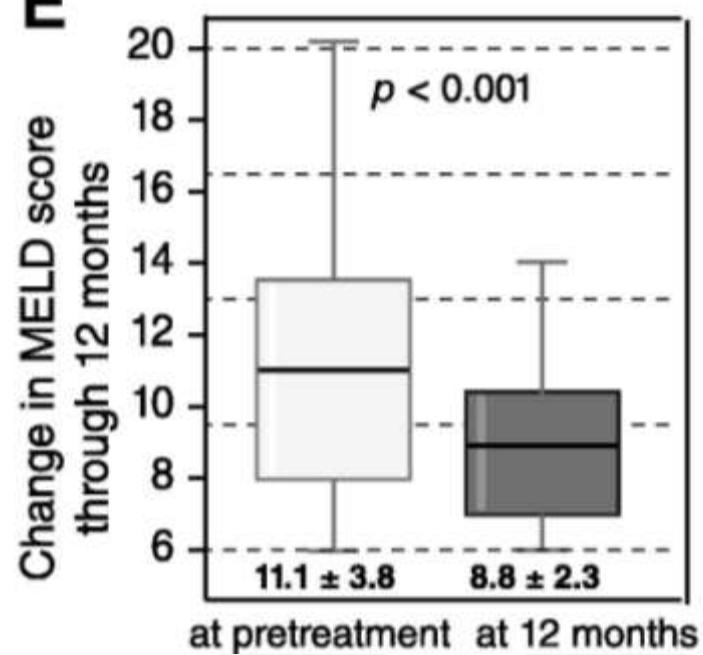


Treatment of Decompensated HBV

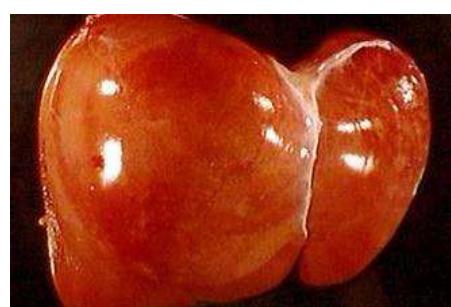
D



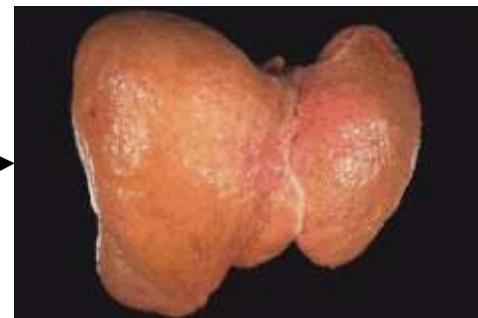
E



Progression of Disease in HCV



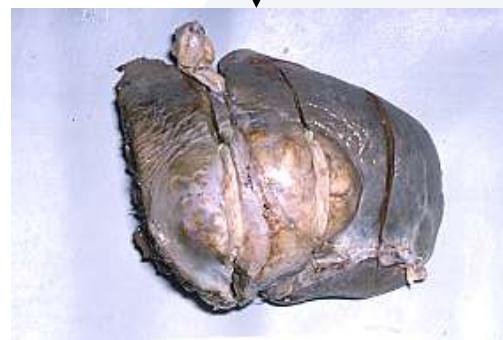
7.7%



6.3%



3.7%



4.6%

9.9 – 20%



8.1 – 54.5%

Progression to HCC in Patients with HCV Infection

Clinical Setting	Geographic Area	No Studies	No Patients	Mean follow up (years)	HCC incidence (%)	95% CI
Chronic hepatitis	Europe	1	329	4.2	0	
	Japan	6	1451	6.2	1.8	1.5 – 2.05
Cirrhosis	Europe & US	13	1284	4.5	3.7	3.2-4.17
	Japan	7	626	5.8	7.1	6.19 – 7.96

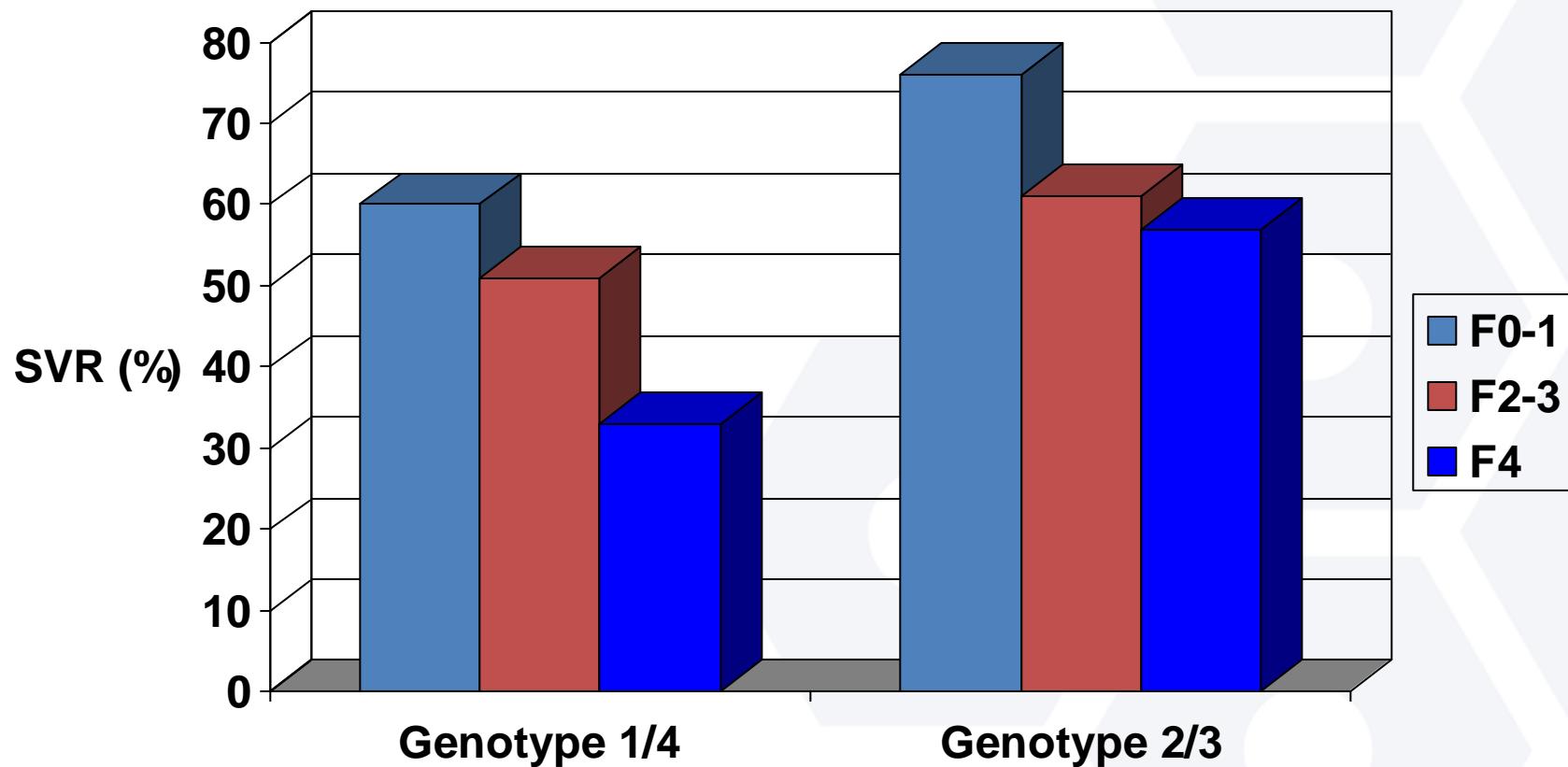
Key questions

- Does treatment improve the liver histologically?
- Does treatment reverse the progression of liver disease?
- Does treatment prevent progression of cirrhosis to decompensation?
- Does treatment prevent progression of cirrhosis to cancer?
- Does treatment prolong life?

Cirrhosis – can it be treated?

- Risks
 - Decompensation
 - Neutropaenia
 - Thrombocytopenia

Treatment Outcome According to Histological Severity



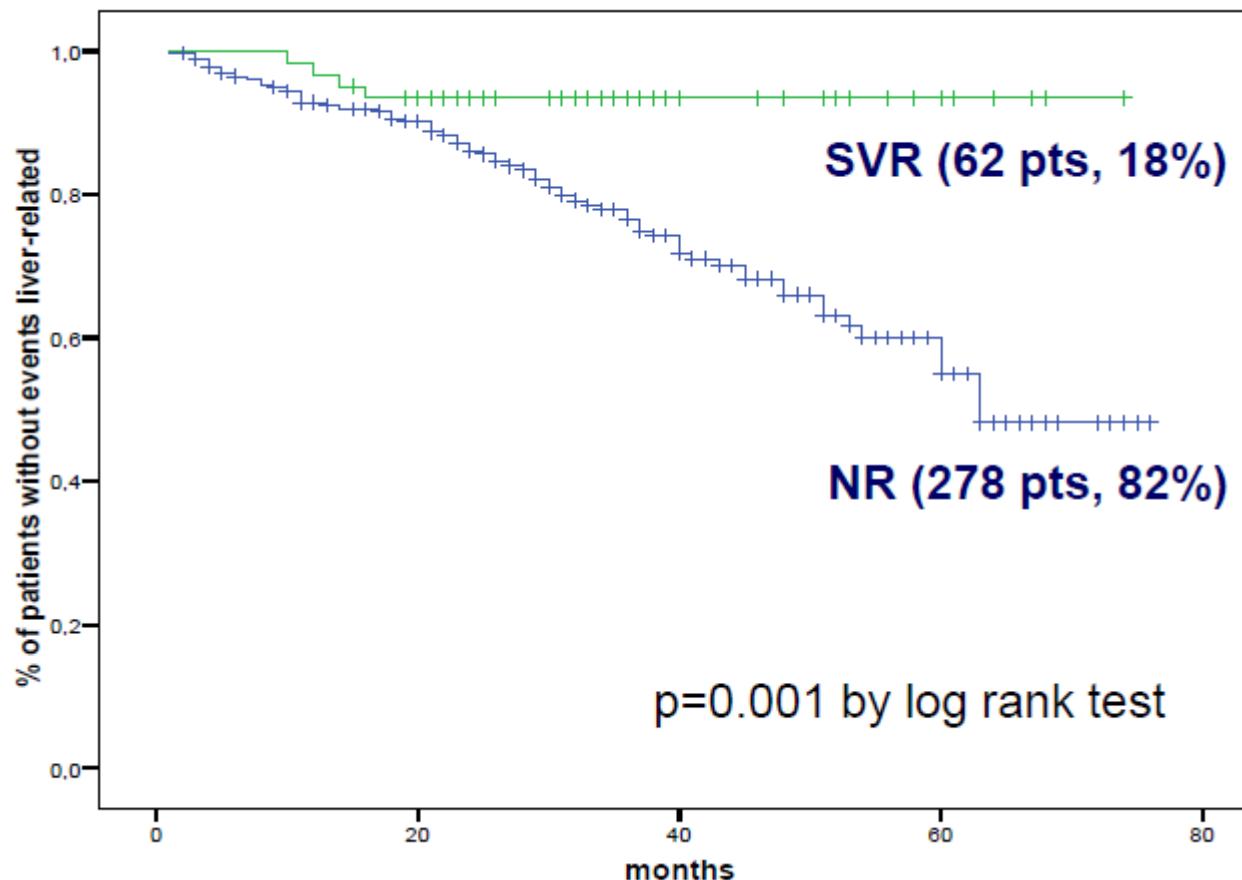
Progression to Cirrhosis According to Treatment Outcome

Author	Year	Follow up (Years)	Rate of cirrhosis in SVR	Rate of cirrhosis in non-SVR
Sata	1997	0.6 – 3.8	0 / 25	9 / 38
Lau	1998	10 – 13	0 / 5	2 / 5
Camma	1998	0.7 – 9	0 / 62	5 / 360
Ajello	1999	10	1 / 10	0 / 21
Morisco	1999	4	0 / 39	12 / 115
Gianni	2001	1 – 6	0 / 15	3 / 21
Shindo	2001	8 – 11	0 / 67	62 / 183
Gallego	2000	4	0 / 11	33 / 87
Swain	2004	8	0 / 989	8 / 997
Vedt	2004	1.6	6 / 110	3 / 15
Cianco	2006	7	0 / 83	3 / 86
Chavaldhamrong	2006	3	0 / 171	27 / 171
			0.43%	7.7%

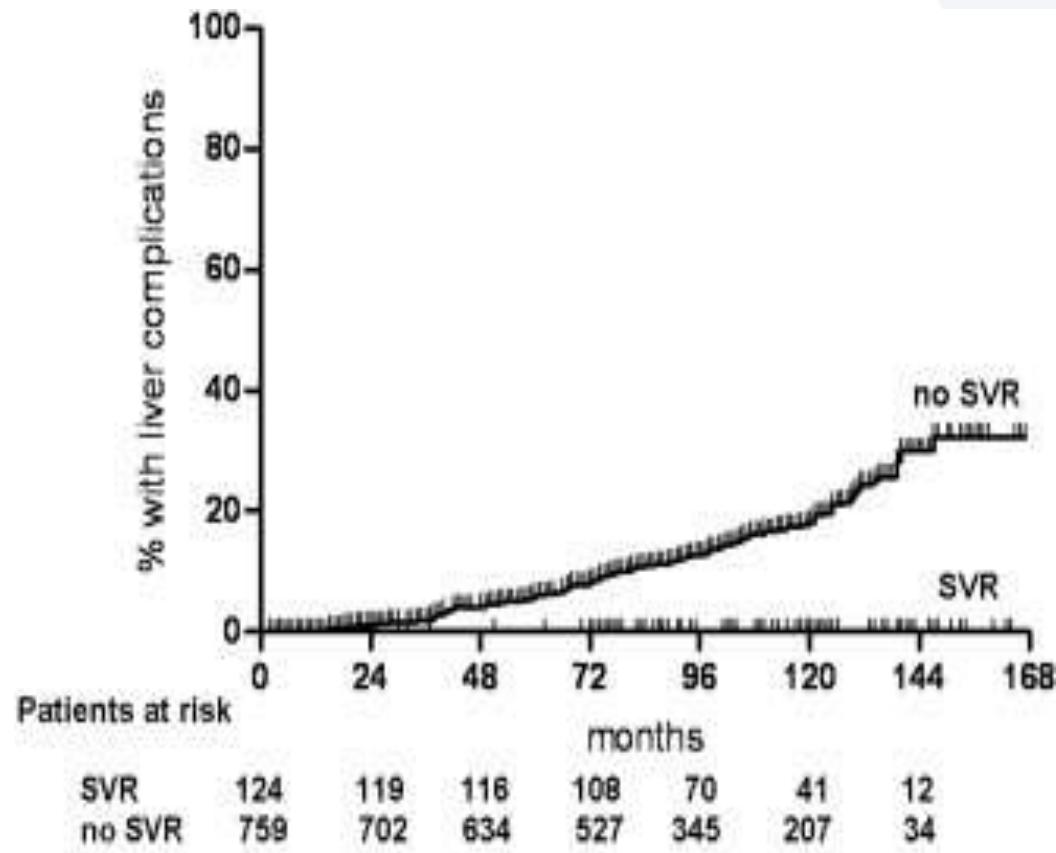
Effect of Treatment on Liver Related Death

Factor	OR	P
Age	1.05	P < 0.005
Treatment	0.54	P = 0.02
SVR vs Untreated	0.05	P < 0.003
No SVR vs Untreated	0.72	P = NS

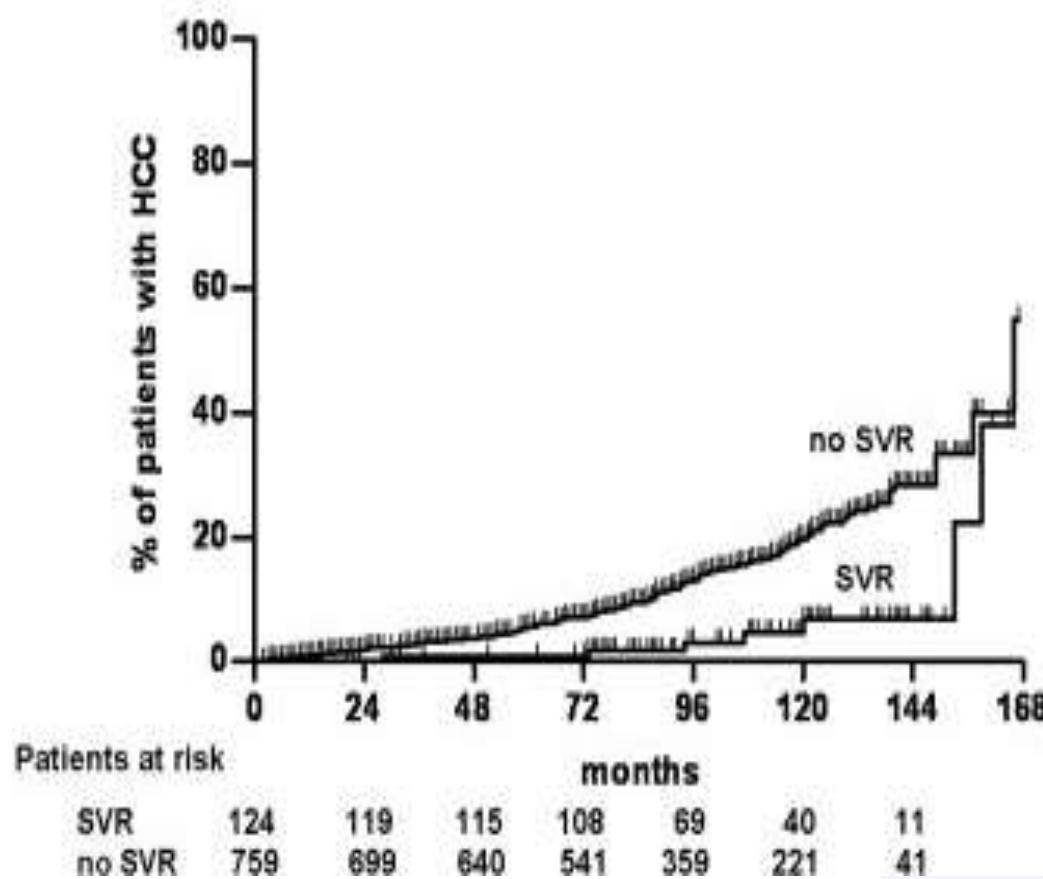
Impact of Viral Elimination on Progression Cirrhosis - Decompensation



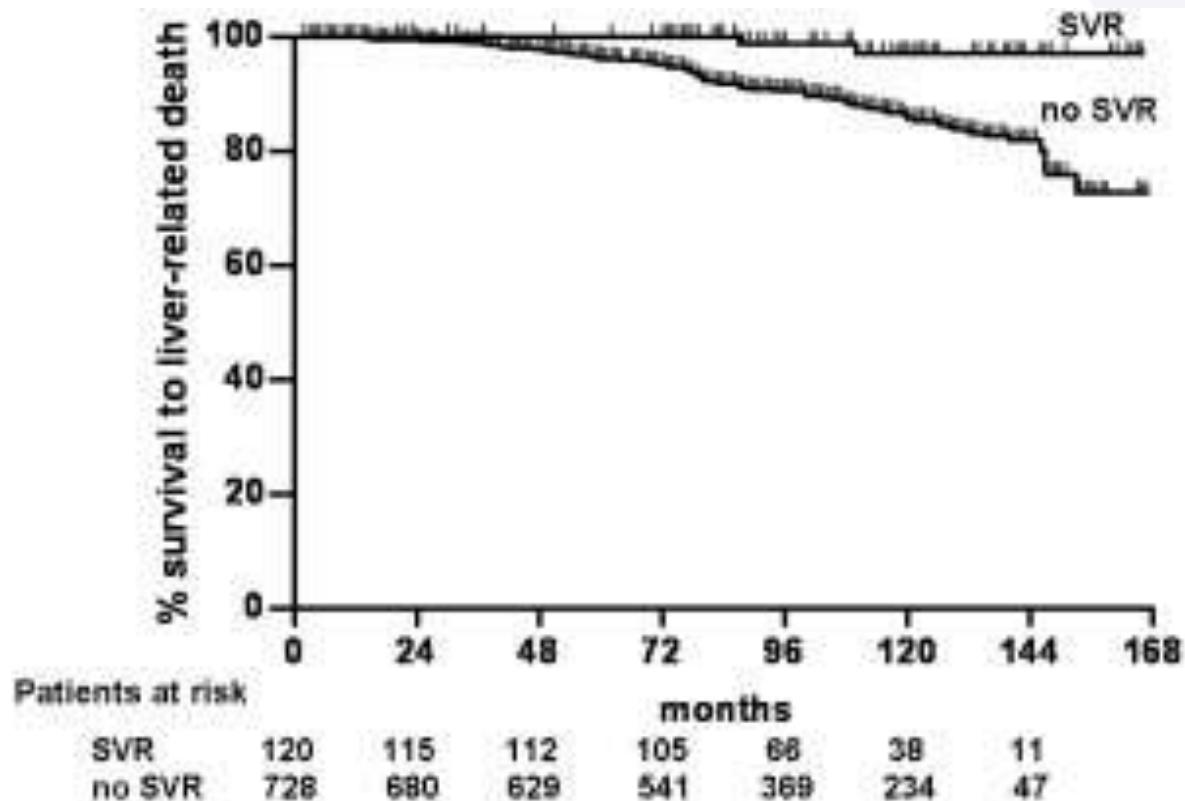
Impact of Viral Elimination on Progression Cirrhosis - Decompensation



Impact of Viral Elimination on Progression Cirrhosis - Cancer



Viral Elimination Reduces Mortality



Summary

- HBV in cirrhotics can be controlled
- Suppression of HBV reduces the risk of disease progression
- Decompensated HBV cirrhosis can be reversed
- HCV in cirrhotics can be cured
- Elimination of HCV reduces the risk of HCC / Liver related death
- Cirrhosis may be reversible