Treatment-Naïve and Treatment-Experienced

Glecaprevir-Pibrentasvir in Non-Cirrhotic Genotype 4, 5, or 6
ENDURANCE-4

Source: Asselah T, et al. Clin Gastroenterol Hepatol. 2018;16:417-26.

Glecaprevir-Pibrentasvir in Non-Cirrhotic Genotype 4, 5, or 6 *ENDURANCE-4: Study Features

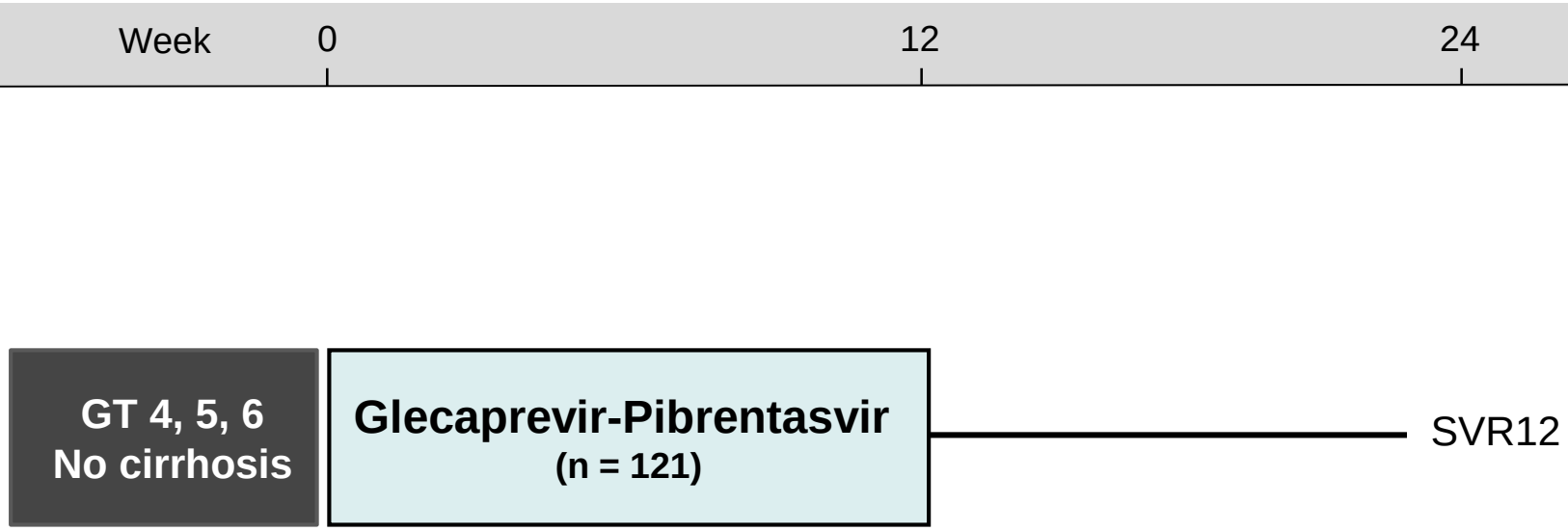
ENDURANCE-4 Trial

- **Design**: Open-label single-arm phase 3 trial to evaluate the safety and efficacy of the fixed-dose combination of glecaprevir-pibrentasvir for 12 weeks in treatment-naïve and treatment-experienced adults with GT 4, 5 or 6 chronic HCV infection without cirrhosis
- **Setting:** Canada, Europe and South Africa
- **Key Eligibility Criteria**
  - Chronic HCV GT 4, 5 or 6
  - HCV RNA ≥1,000 IU/mL at screening
  - Naïve or treated with (1) PEG (or IFN) +/- RBV or (2) SOF + RBV +/- PEG
  - No cirrhosis
  - HIV or chronic HBV coinfection excluded
- **Primary End-Point**: SVR12

***Note**: ENDURANCE-4 was published in conjunction with ENDURANCE-2 and SURVEYOR-II (Part 4)

Source: Asselah T, et al. Clin Gastroenterol Hepatol. 2018;16:417-26.

Glecaprevir-Pibrentasvir in Non-Cirrhotic Genotype 4, 5, or 6
ENDURANCE-4: Study Design



Drug Dosing
Glecaprevir-pibrentasvir (100/40 mg) fixed dose combination; three pills once daily

Source: Asselah T, et al. Clin Gastroenterol Hepatol. 2018;16:417-26.

Glecaprevir-Pibrentasvir in Non-Cirrhotic Genotype 4, 5, or 6
ENDURANCE-4: Baseline Characteristics

Baseline Characteristic	Glecaprevir-Pibrentasvir (n=121)
Age, mean ± SD, years	53 ± 11.0
Male, n (%)	77 (64)
Race, n (%) White Black Asian	 84 (71) 10 (8) 24 (20)
BMI, mean, ± SD kg/m^2	25.7 ± 4.8
IL28B genotype non-CC, n (%)	91 (75)
HCV Genotype, n (%) 4 5 6	 76 (63) 26 (21) 19 (16)
HCV RNA, median (range), log_{10} IU/mL	6.3 (3.6-7.3)
Former IDU, n (%)	32 (26)

Source: Asselah T, et al. Clin Gastroenterol Hepatol. 2018;16:417-26.

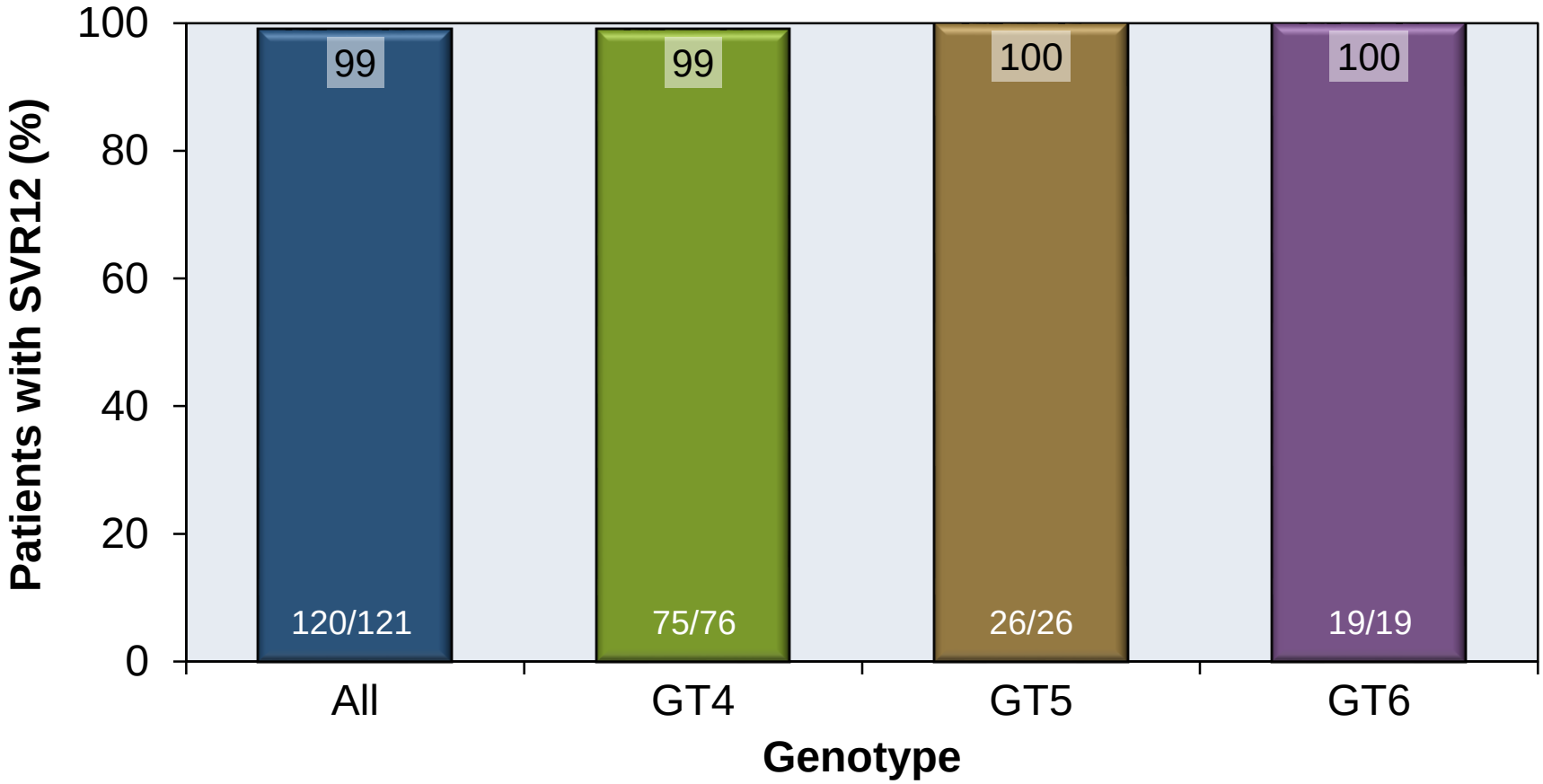
Glecaprevir-Pibrentasvir in Non-Cirrhotic Genotype 4, 5, or 6
ENDURANCE-4: Baseline Characteristics

Baseline Characteristic	Glecaprevir-Pibrentasvir (n=121)
Fibrosis Stage, n (%)	
F0-1	104 (86)
F2	8 (7)
F3	9 (7)
HCV Treatment-Naïve, n (%)	82 (68)
Treatment-Experienced, n (%)	39 (32)
IFN or PEG ± RBV, n (%)	39 (32)
SOF + RBV ± PEG, n (%)	0 (0)
Concomitant PPI use, n (%)	11 (9)

Source: Asselah T, et al. Clin Gastroenterol Hepatol. 2018;16:417-26.

Glecaprevir-Pibrentasvir in Non-Cirrhotic Genotype 4, 5, or 6 ENDURANCE-4: Results

SVR12 (ITT analysis), Overall and by Genotype



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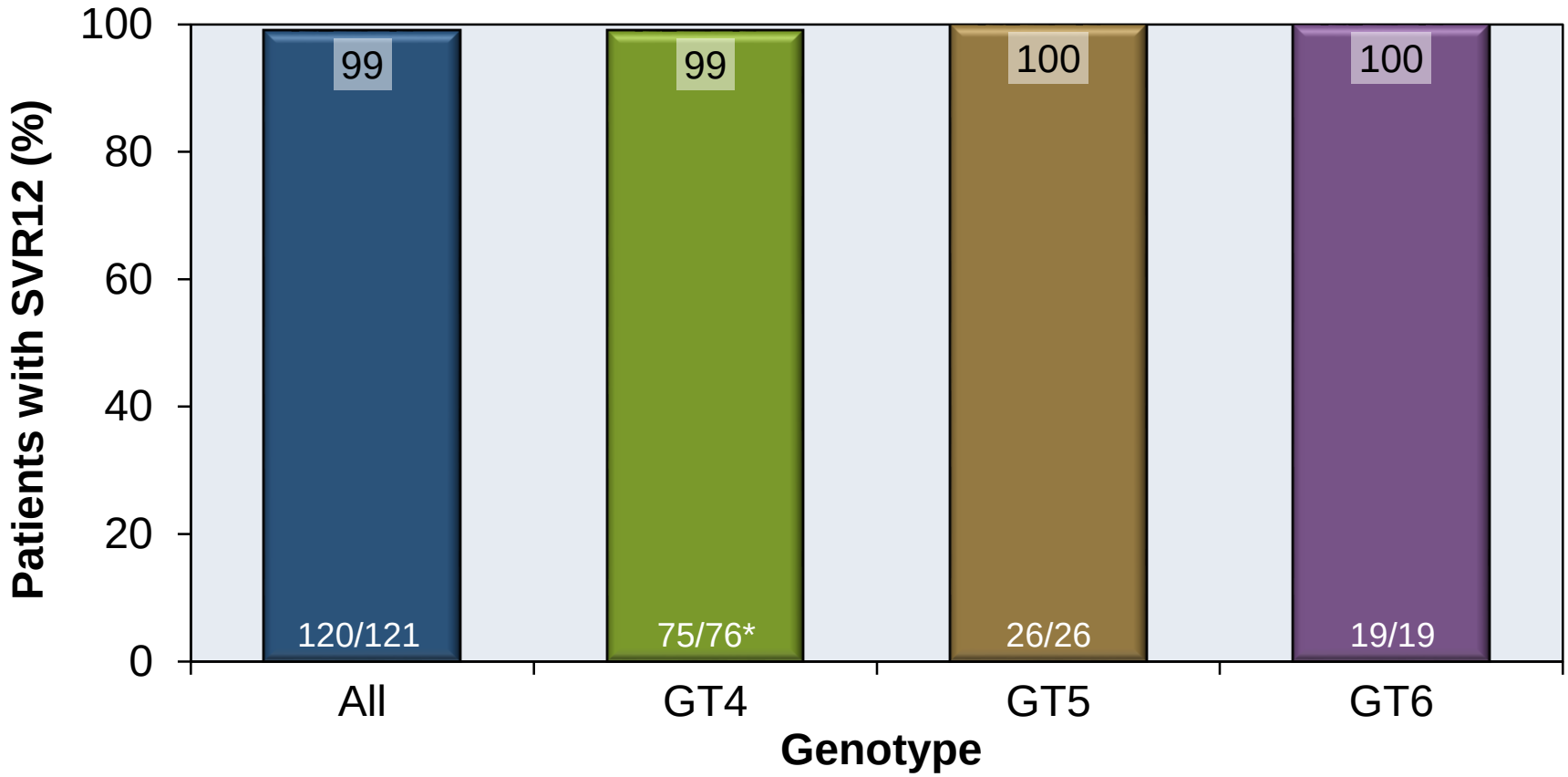
Glecaprevir-Pibrentasvir in Non-Cirrhotic Genotype 4, 5, or 6 ENDURANCE-4: Results

SVR12 (ITT analysis), Overall and by Genotype



*1 patient stopped drug on day 12

Source: Asselah T, et al. Clin Gastroenterol Hepatol. 2018;16:417-26.

Glecaprevir-Pibrentasvir in Non-Cirrhotic Genotype 4, 5, or 6
ENDURANCE-4: Adverse Events

Adverse Events (AEs), n (%)	Glecaprevir-Pibrentasvir (n=121)
AEs leading to drug discontinuation	3 (2.5)*
Serious AEs	1 (0.8)§
AEs occurring in ≥10% of patients	
Fatigue	21 (17)
Headache	25 (21)
Laboratory AEs	
AST grade ≥2 (>3 x ULN)	0
ALT grade ≥2 (>3 x ULN)	0
Total bilirubin grade ≥3 (>3 x ULN)	0

* One patient with anxiety, another with heartburn, third with transient ischemic attack (TIA).
§ Patient with baseline risk factors discontinued drug on day 12 due to TIA.

Source: Asselah T, et al. Clin Gastroenterol Hepatol. 2018;16:417-26.

Glecaprevir-Pibrentasvir in Non-Cirrhotic Genotype 4, 5, or 6 *ENDURANCE-4: Conclusions

Conclusion: “In 3 Phase 3 studies, 8 weeks' treatment with glecaprevir/pibrentasvir produced an SVR12 in at least 93% of patients with chronic HCV genotype 2, 4, 5, or 6 infection without cirrhosis, with virologic failure in less than 1%. The drug combination had a safety profile comparable to 12 week's treatment with glecaprevir/pibrentasvir.”

***Note**: ENDURANCE-4 was published in conjunction with ENDURANCE-2 and SURVEYOR-II (Part 4)

Source: Asselah T, et al. Clin Gastroenterol Hepatol. 2018;16:417-26.

This slide deck is from the University of Washington's *Hepatitis C Online* and *Hepatitis Web Study* projects.

Hepatitis C Online

www.hepatitisc.uw.edu

Hepatitis Web Study

http://depts.washington.edu/hepstudy/

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Hepatitis
web study