Phase 2

Treatment Naïve and Treatment Experienced

Liver Transplantation

Ombitasvir-Paritaprevir-Ritonavir and Dasabuvir + RBV in Liver Transplant Recipients with Recurrent HCV GT1

CORAL-I

Kwo PY, et al. N Engl J Med. 2014;371:2375-82.

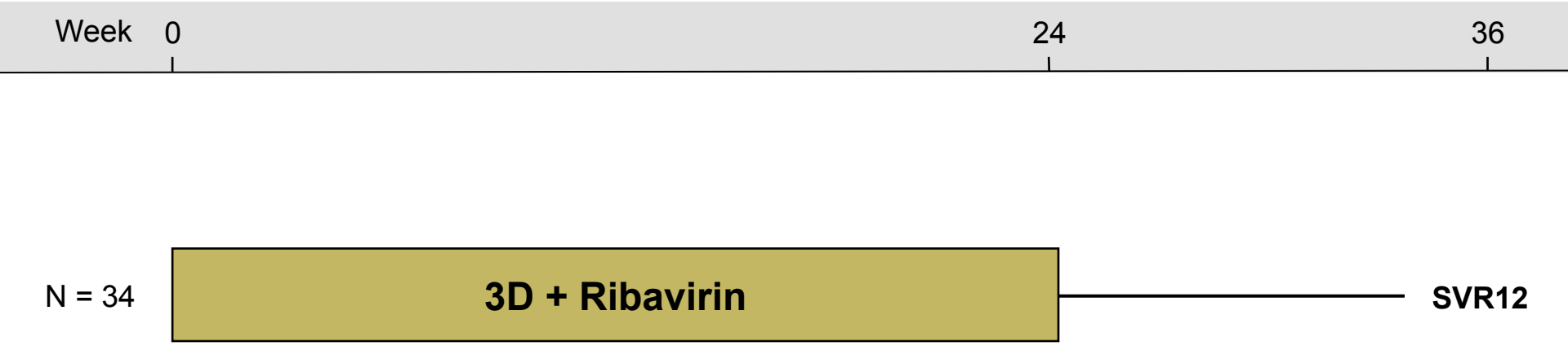
3D + RBV in Liver Transplant Recipients with Recurrent HCV GT1
CORAL-I Trial: Study Design

CORAL-I: Features

- **Design**: Phase 2, open-label, single-arm trial evaluating safety and efficacy of 3D (ombitasvir-paritaprevir-ritonavir + dasabuvir) + ribavirin x 24 weeks in liver transplant recipients with recurrent HCV GT 1
- **Setting**: International
- **Entry Criteria**
  - Chronic HCV infection with genotype 1
  - Liver transplantation due to HCV at least 12 months prior
  - Treatment-naïve after transplantation
  - Pre-transplant treatment with peginterferon + ribavirin allowed
  - Age 18-70
  - Metavir score ≤F2 confirmed by liver biopsy
- **Use of Immunosuppressants**
  - Receiving stable immunosuppressant regimen (tacrolimus or cyclosporin)
  - Tacrolimus or cyclosporin dose based on phase I pharmacokinetic study
  - Prednisone at dose ≤ 5 mg/day permitted but not use of mTOR inhibitors
- **Primary End-Point**: SVR12

Source: Kwo PY, et al. N Engl J Med. 2014;371:2375-82.

3D + RBV in Liver Transplant Recipients with Recurrent HCV GT1
CORAL-I Trial: Regimen



3D = Ombitasvir-Paritaprevir-Ritonavir + Dasabuvir

Drug Dosing

Ombitasvir-Paritaprevir-Ritonavir- (25/150/100 mg once daily) + Dasabuvir: 250 mg twice daily

Ribavirin (RBV): dosing managed per investigator discretion; most patients received 600-800 mg/day

Source: Kwo PY, et al. N Engl J Med. 2014;371:2375-82.

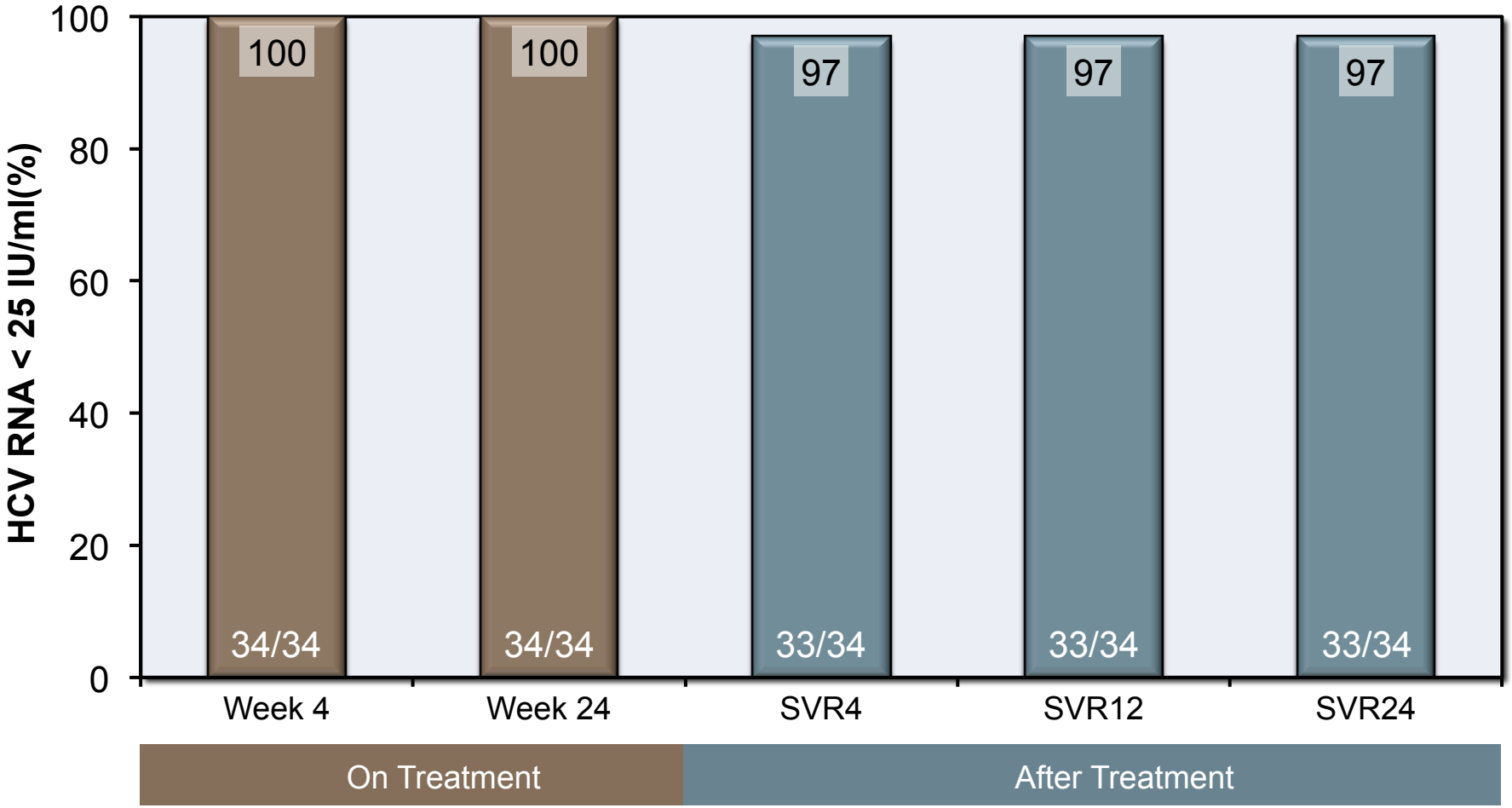
3D + RBV in Liver Transplant Recipients with Recurrent HCV GT1
CORAL-I Trial: Baseline Characteristics

Baseline Characteristic	**3D + Ribavirin** (n=34)
Age (years), Mean	59.6
Male sex–no. (%)	27 (79)
Race–no. (%) White Black Multiple	 29 (85) 4 (12) 1 (3)
Body Mass Index (kg/m^2) Mean	29.7
HCV genotype–no. (%) 1a 1b	 29 (85) 5 (15)
IL28B, non-CC genotype–no. (%)	26 (76)
HCV RNA, $\log_{10}$ IU/ml	6.6
Fibrosis stage (%) F0 F1 F2	 6 (18) 13 (38) 15 (44)

3D = Ombitasvir-Paritaprevir-Ritonavir and Dasabuvir; **RBV** = ribavirin

Source: Kwo PY, et al. N Engl J Med. 2014;371:2375-82.

3D + RBV in Liver Transplant Recipients with Recurrent HCV GT1
CORAL-I Trial: Results



Source: Kwo PY, et al. N Engl J Med. 2014;371:2375-82.

Hepatitis web study

3D + RBV in Liver Transplant Recipients with Recurrent HCV GT1
CORAL-I Trial: Adverse Events

Adverse Event Occurring in > 15% of the 34 Patients Receiving 3D + RBV	
Event	**N (%)**
Any adverse event	33 (97)
Fatigue	17 (50)
Headache	15 (44)
Cough	11 (32)
Anemia	10 (29)
Diarrhea	9 (26)
Insomnia	9 (26)
Asthenia	8 (24)
Nausea	8 (24)
Muscle spasms	7 (21)
Rash	7 (21)
Back pain	6 (18)
Dizziness	6 (18)
Peripheral edema	6 (18
Rhinorrhea	6 (18)

3D = Ombitasvir-Paritaprevir-Ritonavir and Dasabuvir; **RBV** = ribavirin

Source: Kwo PY, et al. N Engl J Med. 2014;371:2375-82.

3D + RBV in Liver Transplant Recipients with Recurrent HCV GT1
CORAL-I Trial: Conclusions

Conclusions: "Treatment with the multitargeted regimen of ombitasvir-ABT-450/r and dasabuvir with ribavirin was associated with a low rate of serious adverse events and a high rate of sustained virologic response among liver-transplant recipients with recurrent HCV genotype 1 infection, a historically difficult-to-treat population."

Note: ABT-450/r = Paritaprevir-Ritonavir

Source: Kwo PY, et al. N Engl J Med. 2014;371:2375-82.

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
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