

## Stanford Medication Usage Guide Ribavirin (oral)

### Usage

- Ribavirin is FDA approved for the treatment of hepatitis C in combination with IFN- $\alpha$ .
- Off-label: treatment of severe respiratory infections with community-acquired respiratory viruses in immunocompromised patients. E.g.
  - Respiratory syncytial virus (RSV)
  - Metapneumovirus (MPV)
  - Parainfluenza virus (PIV)

### Dosing

CrCl	Oral Ribavirin Dosing Regimen for RSV, MPV, PIV (typical duration 5-10 days)
>50	15 – 20 mg/kg/day in 2 to 3 divided doses; round to nearest 200 mg dose (max dose: 1800 mg/day)
30-50	200 mg TID – 400 mg BID <sup>a</sup>
<29, ESRD, IHD	No recommendations exist. Some experts recommend 200 mg daily <sup>a</sup>
CRRT, PD	No recommendations exist. Consult ASP/ID pharmacist

a. The prescribing information for Rebetol® (capsule) states that use is contraindicated if CrCl <50 mL/min. The prescribing information for Copegus® (tablet) states that the dose should be reduced in patients with CrCl  $\leq$  50 mL/min.

### Pharmacokinetics/Pharmacodynamics

Ribavirin pharmacokinetics	
<b>Bioavailability</b>	64%, increased by high fat meal
<b>Distribution</b>	V <sub>d</sub> 2,825 L (single dose): distribution significantly prolonged in the erythrocyte (16 to 40 days), which can be used as a marker for intracellular metabolism
<b>Metabolism</b>	Hepatically and intracellularly (forms active metabolites)
<b>Half-life elimination</b>	Capsule: 24h (single dose), 298h (steady state) Tablet: 120-170h (single dose), 288h (multiple doses)
<b>Time to peak, serum</b>	3h (capsule) 2h (tablet)
<b>Excretion</b>	61% urine, 12% feces Note: Plasma ribavirin is removed by hemodialysis with an extraction ratio of approximately 50%; however, due to its large V <sub>d</sub> , plasma exposure is not expected to change with hemodialysis

### Monitoring Parameters

- Re-evaluate therapy if adverse reactions occur e.g. clinically significant anemia necessitating transfusion may warrant discontinuation or dose decrease.
- Risk of may be increased in those with renal insufficiency, dialysis, low baseline Hgb levels, or concomitant therapy associated with causing bone marrow suppression.

### Special Circumstances

- Hazardous drug: do not crush/open capsules/tablets. Product is available as oral solution
- Pregnancy FDA risk category X

## References:

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3. Smolders, Elise J., Clara T. M. M. de Kanter, Bart van Hoek, Joop E. Arends, Joost P. H. Drenth, and David M. Burger. "Pharmacokinetics, Efficacy, and Safety of Hepatitis C Virus Drugs in Patients with Liver And/Or Renal Impairment." *Drug Safety* 39, no. 7 (July 2016): 589–611. doi:10.1007/s40264-016-0420-2.
4. Hirsch, Hans H., Rodrigo Martino, Katherine N. Ward, Michael Boeckh, Hermann Einsele, and Per Ljungman. "Fourth European Conference on Infections in Leukaemia (ECIL-4): Guidelines for Diagnosis and Treatment of Human Respiratory Syncytial Virus, Parainfluenza Virus, Metapneumovirus, Rhinovirus, and Coronavirus." *Clinical Infectious Diseases: An Official Publication of the Infectious Diseases Society of America* 56, no. 2 (January 15, 2013): 258–66. doi:10.1093/cid/cis844.
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6. Lexi-comp

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- B.** Gatekeeper: Antimicrobial Stewardship Program
- C.** Review and Renewal Requirement  
This document will be reviewed every three years and as required by change of law or practice
- D.** Revision/Review History:
- E.** Approvals
  1. Antimicrobial Subcommittee: 8/17/2017
  2. P&T: 9/15/2017

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