

Regorafenib PK Fact Sheet

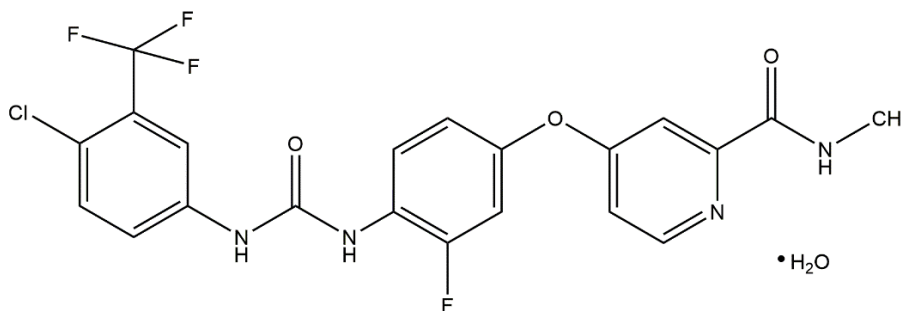
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Details

Generic Name	Regorafenib
Trade Name	Stivarga®
Class	Oncolytics (Protein kinase inhibitors)
Molecular Weight	500.83
Structure	



Summary of Key Pharmacokinetic Parameters

<i>Linearity/non-linearity</i>	Exposure increases dose proportionally up to 60 mg and less than proportionally doses greater than 60 mg.
<i>Steady state</i>	Time to steady state not reported.
<i>Plasma half-life</i>	Regorafenib and M-2: 20-30 h M-5: ~60 h
<i>C_{max}</i>	3.9 µg/mL after oral administration of 160 mg, at steady state.
<i>C₂₄</i>	Not reported
<i>AUC</i>	58.3 µg·h/mL after oral administration of 160 mg, at steady state.
<i>Bioavailability</i>	The mean relative bioavailability of 60 or 100 mg tablets compared to an oral solution is 69% and 83%, respectively.
<i>Absorption</i>	The concentrations of regorafenib and its major pharmacologically active metabolites (M-2 and M-5) are highest when given after a low-fat breakfast, compared to either a high-fat breakfast or fasting condition.
<i>Protein Binding</i>	99.5%.
<i>Volume of Distribution</i>	Not reported.
<i>CSF:Plasma ratio</i>	Not reported.
<i>Semen:Plasma ratio</i>	Not reported.
<i>Renal Clearance</i>	<10% under steady state conditions.
<i>Renal Impairment</i>	No dose adjustment is recommended for patients with renal impairment.
<i>Hepatic Impairment</i>	No dose adjustment is recommended in patients with mild or moderate impairment. Closely monitor patients with hepatic impairment for adverse reactions Regorafenib is not recommended in patients with severe hepatic impairment as this population has not been studied.

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Metabolism and Distribution

<i>Metabolised by</i>	CYP3A4, UGT1A9.
<i>Inducer of</i>	None known.
<i>Inhibitor of</i>	BCRP.
<i>Transported by</i>	BCRP, P-gp.

References

Unless otherwise stated (see below), information is from:

Stivarga® Summary of Product Characteristics, Bayer plc, August 2018.

Stivarga® Prescribing Information, Bayer HealthCare Pharmaceuticals Inc., June 2019.