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Hepatic pharmacokinetics of TNP-470 in the rat. A polyglutamated analog of heptestatin, TNP-470 (NSC 679060) is a macromolecular antineoplastic agent that has been shown to inhibit angiogenesis in several models in vivo. However, limited information is available on its pharmacokinetics. To determine the hepatic disposition of this drug, a method for the determination of its concentration in small tissue samples was developed. TNP-470 was extracted from tissue samples by precipitation with TCA. The analyte was assayed by fluorometric

measurement of the inhibition of sulfatase activity at 340 nm, using p-nitrobenzene sulfate as substrate. Limit of quantitation was 0.2 microgram/mL in plasma and tissue samples. Distribution of TNP-470 was best fit by a monoexponential function for tissue concentrations below 1 microgram/g tissue. Following a dose of 1 mg/kg i.v. to rats, hepatic concentrations were highest at 0.5 h. Plasma and urine concentrations increased to maximum levels after 0.5 h and remained above 5 micrograms/mL for up to 12 h. In conclusion, TNP-470 is metabolized to a greater extent in the liver than in any other organ, and the extent of metabolism appeared to be related to the dose administered. The Paris-based fund, which is also backed by French billionaire Alain Robert,

is also understood to be in talks with Saudi Arabia-based Petra Investment Partners to raise about \$1 billion to invest in privately held companies in the Middle East, Southeast Asia and Latin America. The deal would reportedly be split equally between the two funds with the majority of the investments in the fields of software, technology and Fintech, according to The Financial Times. The FT also reported the WS Fund "would initially have four partners, including Saudi billionaire Prince Alwaleed bin Talal, who founded Kingdom Holding Co, and his new venture capital fund". WS Fund has earlier this year reported its shareholders included Kingdom Holding and Roboratio. According to its website, the WS Fund controls a total of €4.1 billion

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