

## Supplementary Materials

### Local and systemic thrombotic complications in cirrhotic patients with hepatocellular carcinoma

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### Supplementary Table 1. Risk Factors associated with PVT occurrence in patients with cirrhotic patients

| Author, Years                               | Type of study                   | Patients with Liver Disease (n)  | PVT   | Risk Factor  |
|---|---------------------------------|--|---|--|
| Nonami <i>et al.</i> , 1992 <sup>[48]</sup> | Retrospective study (1989-1990) | 849 patients who underwent LT (87 HCC and cirrhosis, 47 HCC without cirrhosis) | HCC (+) incidence 34.8% vs. HCC without cirrhosis incidence 8.5%, Post necrotic cirrhosis | Encephalopathy $P < 0.02$<br>Ascites $P < 0.005$<br>Gastrointestinal bleeding $< 0.001$<br>Previous splenectomy $< 0.01$ |

|   |  |  |   |  |
|---|--|--|---|--|
|   |  | cirrhosis  | incidence   |  |
|   |  |  | 15.7%   |  |
| Davidson <i>et al.</i> , 1994 <sup>[49]</sup> | Prospective study (1988-1992)                  | 132 patients who underwent OLTx (12 cryptogenic cirrhosis, 22 cirrhosis and HCC, 5 autoimmune hepatitis) | Cirrhosis + HCC incidence (6/22 (27.3%) vs. Cirrhosis incidence (10/110 (9.1%)) | Autoimmune chronic active hepatitis $\chi^2:13.3, P < 0.001$<br>Cryptogenic cirrhosis $\chi^2:7.2, P < 0.01$<br>HCC $\chi^2:5.7, P < 0.05$ |
| Ravaioli <i>et al.</i> , 2011 <sup>[50]</sup> | Retrospective study (1998-2008)                | 889 patients LT candidates (282 with HCC)  | HCC (+) incidence 37/282 (13%)  | HCC significantly associated with PVT risk at multivariate analysis (HR = 1.81, $P < 0.05$ )   |
| Zanetto <i>et al.</i> , 2017 <sup>[20]</sup>  | Prospective study (2012-2013) follow-up 1 year | 41 patients with cirrhosis and HCC (CHILD A/B/C 20/12/9) 35 patients with non HCC cirrhosis (CHILD       | HCC (+) incidence 10/41 (24.4%) vs HCC (-) incidence 4/35 (11.4%)               | HCC(+) HR: 10.34, $P = 0.03$<br>Thromboelastogram: Maximum Clot Firmness (MCF* > 25 mm), HR = 6, $P = 0.001$                               |

|   |   |   |  |   |
|---|---|---|--|---|
|   |   | A/B/C   |  |   |
|   |   | 9/17/9)   |  |   |
| Cagin <i>et al.</i> ,<br>2016 <sup>[51]</sup> | Retrospective<br>study (2009-<br>2014)        | 461 patients<br>with<br>cirrhosis:<br>HCC + 69<br>(15%),<br>HCC- 392<br>(85%)   | HCC (+)<br>prevalence<br>(13/69<br>18.8%),<br>HCC (-)<br>(32/392<br>8.2%)            | HCC significantly<br>associated with PVT $P <$<br>0.001   |
| Serag <i>et al.</i> ,<br>2020 <sup>[52]</sup> | Prospective<br>study<br>(follow-up 1<br>year) | 44 patients<br>with<br>cirrhosis +<br>HCC<br>(CHILD<br>A/B/C<br>12/20/12)<br>47 patients<br>with<br>cirrhosis<br>(CHILD<br>A/B/C<br>14/18/14) | HCC (+)<br>incidence<br>(10/44<br>22.7%);<br>HCC(-)<br>incidence<br>(6/47<br>12.7%)  | Differences between PVT<br>(+) and PVT (-) in all<br>cirrhosis with and without<br>HCC:<br>In cirrhosis with HCC<br>Annexin A5/PS + MP ratio<br>$P < 0.001$<br>PS + MPs $P < 0.001$<br>Portal flow velocity $P <$<br>0.001 ** |
| Faccia <i>et al.</i> , 2022 <sup>[38]</sup>   | Retrospective<br>study (1982-<br>2017)        | 7,445<br>hospitalized<br>cirrhotic<br>patients<br>(HCC+<br>1,524)   | HCC (+)<br>prevalence<br>(162/1524<br>10.6%);<br>HCC (-)<br>prevalence<br>( 220/5921 | (multivariate logistic<br>regression analyses)<br>Endoscopic signs of portal<br>hypertension OR = 1.33, $P$<br>= 0.02<br>Hepatic encephalopathy<br>OR = 13.98, $P < 0.0001$   |

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|  |                     |                            |                         |   |
|--|---------------------|----------------------------|-------------------------|---|
|  |                     |                            | 3.7%)                   | HCC OR 4.59, $P < 0.0001$<br>Diabetes OR = 1.68, $P = 0.0001$<br>Abdominal<br>surgery/invasive procedure<br>OR = 2.03, $P < 0.0001$   |
| Senzolo <i>et al.</i> , 2023 <sup>[44]</sup> | Retrospective study | 750 cirrhotic HCC patients | 88/750 PVT at diagnosis | (multivariate analyses of the occurrence of PVT at HCC diagnosis)<br>Pre-treatment total tumor volume ( $P < 0.001$ )<br>Clinically significant portal hypertension ( $P = 0.005$ ) |

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\* Maximum Clot Firmness (MCF) is the maximum amplitude in millimeters reached in the thromboelastogram. MCF > 25 mm was associated with a 5-fold increased PVT risk [RR: 4.8 (2–11.3),  $P = 0.0001$ ]; \*\* Cut off HCC (+): Annexin A5/MP ratio < 0,0277; PS + MPs > 38.7 nm/L, Portal flow velocity < 15 cm/sec; Cut off HCC (-): Annexin A5/MP ratio < 0,0028; PS + MPs > 35.3 nm/L, Portal flow velocity < 15 cm/sec. PVT: portal vein thrombosis; LT: liver transplant; MCF: maximum clot firmness; HCC: hepatocellular carcinoma; HR: hazard ratio.