

Fecal microbiota transplant is associated with lower risk of mortality, hepatic encephalopathy, ascites, and infection patients with severe alcohol-associated hepatitis: A systematic review and meta-analysis

Zohaib Ahmed¹, Andrew Kelly¹; Joyce Badal², Wade M. Lee-Smith³, Manesh Gangwani¹, Yaseen Alastal¹, Mona Hassan¹.

¹Division of Gastroenterology, Department of Medicine, The University of Toledo, Toledo, OH 43614

²College of Medicine and Life Sciences, The University of Toledo, Toledo, OH 43614

³College of University Libraries, The University of Toledo, Toledo, OH 43614

*Corresponding author: zohaib.ahmed@utoledo.edu

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Introduction: Severe alcohol-associated hepatitis (SAH) is an acute, inflammatory liver disease that results in disruptions of the gut microbiome leading to bacterial translocation which drives systemic inflammation and end-organ damage. Despite high risk of mortality, treatment options are limited. Fecal microbiota transplant (FMT) may help restore the balance of healthy bacteria in patients with disrupted gut microbiomes due to SAH, which may decrease systemic inflammation, infection, and mortality. As the role of FMT in the treatment of SAH is not yet established, we conducted a systematic review and meta-analysis to evaluate the currently available literature regarding the impact of FMT on outcomes in patients with SAH.

Methods: A comprehensive search strategy was used to identify studies that reported outcomes of patients with SAH receiving FMT compared to no FMT in Embase, MEDLINE (PubMed), Cochrane Library, Web of Science Core Collection, and Korean Journal Index, and Global Index Medicus. Outcomes of interest included 1-, 3-, and 6- month mortality, overall mortality, and risk of HE, ascites, upper GI bleeding, and infection. RevMan software was used for statistical analysis.

Results: 7 studies with a total of 384 patients were included in the final meta-analysis. Patients who received FMT had significantly lower risk of 1-month mortality (RR: 0.51, 95% CI: 0.29-0.91, p=0.02), 3-month mortality (RR: 0.61, 95% CI: 0.38-0.98, p=0.04), and overall mortality (RR: 0.58, 95% CI: 0.38-0.87, p=0.009) compared to those who did not receive FMT, although the difference in 6-month mortality did not reach statistical significance (RR: 0.73, 95% CI: 0.18-2.89, p=0.65). Patients who

received FMT also had significantly lower risk of hepatic encephalopathy (RR: 0.27, 95% CI: 0.16-0.46, $p < 0.00001$), ascites (RR: 0.47, 95% CI: 0.33-0.67, $p < 0.0001$), noncritical infections (RR: 0.36, 95% CI: 0.21-0.6, $p = 0.0001$), and critical infections (RR: 0.28, 95% CI: 0.17-0.48, $p < 0.00001$). There was no significant difference in risk of upper gastrointestinal bleeding (RR: 0.77, 95% CI: 0.48- 1.24, $p = 0.28$).

Discussion: FMT for SAH is associated with significantly lower risk of 1-month, 3-month, and overall mortality, as well as lower risk of hepatic encephalopathy, ascites, and both critical and non-critical infections. Further studies, particularly large randomized controlled trials, are needed to establish the role of FMT in the treatment of patients with SAH.