Vatsalya, V., Kong, M., Gobejishvili, L., Chen, W.-Y., Srivastava, S., Barve, S., McClain, C., & Joshi-Barve, S. (2019). Urinary acrolein metabolite levels in severe acute alcoholic hepatitis patients. *American Journal of Physiology-Gastrointestinal and Liver Physiology*, *316*(1), G115-G122. <a href="https://doi.org/10.1152/ajpgi.00209.2018">https://doi.org/10.1152/ajpgi.00209.2018</a>

## **Definitions**

- **Acrolein**: A harmful chemical produced when the body processes alcohol and is also found in the environment (e.g., cigarette smoke).
- **3-Hydroxypropylmercapturic Acid (HPMA)**: A substance measured in urine that indicates the level of acrolein in the body.
- **Severe Acute Alcoholic Hepatitis (AAH)**: A serious liver condition caused by heavy alcohol use, leading to liver inflammation and damage.
- **Proinflammatory Cytokines**: Proteins (e.g., IL-1 $\beta$ , IL-8, TNF $\alpha$ ) that increase inflammation and can lead to liver cell death.

# **Key Findings**

- Patients with severe AAH had higher levels of HPMA in their urine compared to those with non-severe AAH and healthy individuals.
- HPMA levels were linked to disease severity and liver cell death in patients with severe AAH.
- The presence of specific cytokines (IL-1 $\beta$ , IL-8, TNF $\alpha$ ) along with HPMA was strongly associated with liver cell death.

## **Introduction**

This study explores the relationship between acrolein levels in the body and liver damage in patients with severe acute alcoholic hepatitis (AAH). Acrolein is a toxic chemical produced when alcohol is metabolized, and high levels can indicate severe liver damage. The researchers measured urinary HPMA to determine acrolein levels and studied their association with liver disease severity and inflammation markers.

## **Main Content**

## **Background**

Chronic alcohol use can lead to liver disease, ranging from simple fatty liver to severe conditions like hepatitis and cirrhosis. This study investigates the role of acrolein, a harmful byproduct of alcohol metabolism, in liver damage. It aims to see if HPMA in urine can be a marker for disease severity in AAH patients.

## Methods

## • Study Population:

- o 30 patients with AAH (16 severe, 14 non-severe) and 10 healthy individuals were included.
- o Patients were diagnosed at hospitals in Louisville, Kentucky.

#### Measurement of HPMA:

- o Urine samples were collected from all participants.
- HPMA levels were measured using a specific laboratory technique.

## • Assessment of Liver Injury and Inflammation:

- Blood samples were collected to measure proinflammatory cytokines and liver injury markers.
- o Disease severity was assessed using standard clinical scores (e.g., MELD score).

#### Results

#### • HPMA Levels:

- Patients with severe AAH had significantly higher HPMA levels compared to non-severe patients and healthy controls.
- No significant difference in HPMA levels was found between non-severe patients and healthy controls.

#### Association with Disease Severity:

- HPMA levels showed a complex relationship with disease severity, being higher in severe cases.
- o In patients with severe AAH, HPMA levels were associated with liver injury markers and the presence of proinflammatory cytokines.

### • Cytokines and Liver Cell Death:

- o Higher levels of cytokines (IL-1β, IL-8, TNFα) were found in severe AAH patients.
- o These cytokines, along with HPMA, were strongly linked to liver cell death.

# Conclusion

This study suggests that measuring HPMA levels in urine can help identify severe liver damage in patients with acute alcoholic hepatitis. High HPMA levels are associated with increased inflammation and liver cell death. These findings could lead to better ways of diagnosing and treating severe liver conditions caused by alcohol use. Increasing our understanding of acrolein's role in liver damage may also lead to new treatments to protect the liver.

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