



# Evaluation of Pain Medications in Patients with Liver Disease

Jeremiah Vance Pharm.D. Candidate  
Carrie Vogler Pharm.D., BCPS

SOUTHERN ILLINOIS  
UNIVERSITY EDWARDSVILLE  
SCHOOL OF PHARMACY

## BACKGROUND

- Pain is reported in 82% of patients with liver disease and cirrhosis<sup>1</sup>
- No guidelines or protocols exist for the treatment of pain in patients with liver disease
- Cirrhosis is responsible for approximately 770,000 deaths annually<sup>2</sup>
- Acetaminophen dosing is 2-3 grams in 24 hours for patients with liver disease (2 grams in cirrhosis)<sup>3</sup>
- Opioids undergo extensive dosage adjustments in liver disease

## OBJECTIVES

- To observe pain locations, pain scores, liver disease severity (Child Pugh or MELD Score), and pain medications/doses that are being used in patients with liver disease

## METHODS

### Study Design:

- Single Center, Retrospective Chart Review
- 500 bed teaching hospital in Springfield, Illinois

### IRB Approval:

- The Springfield Committee for Research Involving Human Subjects (SCRIHS)

### Inclusion Criteria:

- Patients aged 18 to 89 years of age
- ICD 10 codes K70 Alcoholic Liver Disease and K74 Fibrosis and cirrhosis of liver
- Must have received at least one dose of analgesic medication within 72 hours of admission

### Exclusion Criteria:

- Patients end of life or hospice
- Hospital stay < 24 hours
- Patients who presented with drug-induced overdose

### Data Analysis:

- Descriptive statistics:
  - Average, mean, median, mode of data, and range with standard deviation

## Results

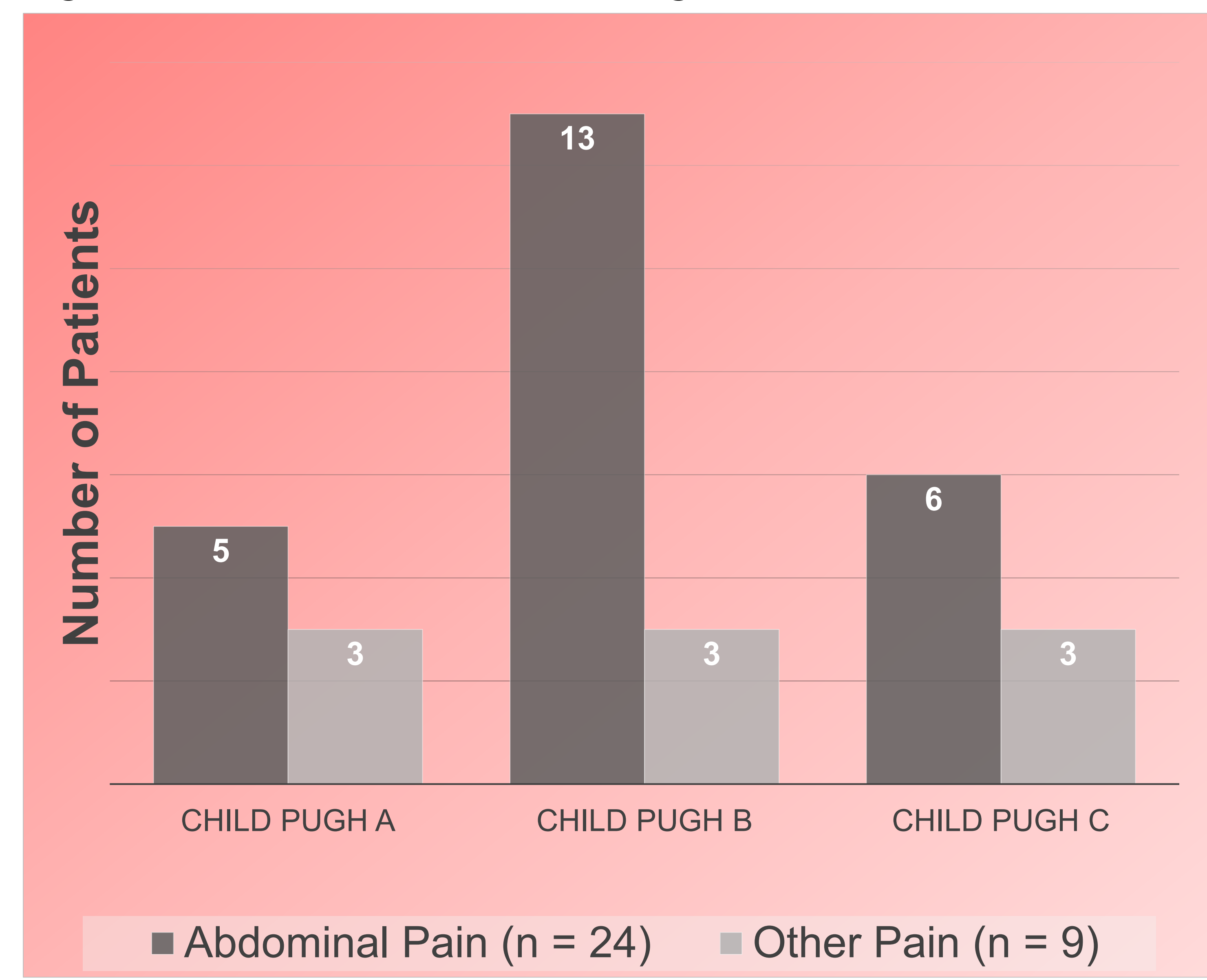
Table 1. Baseline Characteristics

| Variable         | Mean ± SD, n (%) |
|------------------|------------------|
| Age, years       | 52 ± 13.2        |
| Height, in       | 68 ± 3.8         |
| Weight, kg       | 98.9 ± 28.7      |
| MELD Score       | 23 ± 6.7         |
| Sex, male        | 20 (60.6)        |
| Reported race    |                  |
| White            | 29 (87.8)        |
| African American | 4 (12.2)         |
| Child Pugh Score |                  |
| A                | 8 (24)           |
| B                | 16 (49)          |
| C                | 9 (27)           |

Table 2: Medication administrations over 72 hours (n = 33)

| Medication                   | Administrations n = 232 |
|------------------------------|-------------------------|
| Morphine IV                  | 79 doses                |
| Hydrocodone/acetaminophen PO | 61 doses                |
| Acetaminophen PO             | 24 doses                |
| Hydromorphone IV             | 19 doses                |
| Tramadol PO                  | 16 doses                |
| Morphine PO                  | 10 doses                |
| Oxycodone PO                 | 10 doses                |
| Fentanyl IV                  | 5 doses                 |
| Aspirin PO                   | 3 doses                 |
| Ketorolac IV                 | 3 doses                 |
| Ibuprofen PO                 | 1 dose                  |
| Tramadol/acetaminophen PO    | 1 dose                  |

Figure 1. Pain Location and Child Pugh Score



## Results

Table 3. First Pain Medication by Child Pugh Score and Pain Score

| Medication and Route*          | Child Pugh Score (A, B, C) | n = Number of Patients | Pain Score Pre-administration avg±SD (Range)# | Pain Score Post-administration avg±SD (Range) |
|--------------------------------|----------------------------|------------------------|---|---|
| Morphine IV                    | A                          | 6                      | 8.6 ± 1.12 (4-10)                             | 5.8 ± 3.14 (0-10)                             |
|                                | B                          | 6                      |   |   |
|                                | C                          | 4                      |   |   |
| Fentanyl IV                    | A                          | 0                      | 8.8 ± 1.26 (9-10)                             | 6.2 ± 3.03 (2-10)                             |
|                                | B                          | 3                      |   |   |
|                                | C                          | 2                      |   |   |
| Hydrocodone/acetaminophen PO** | A                          | 0                      | 7.5 ± (6-10)                                  | 6   |
|                                | B                          | 4                      |   |   |
|                                | C                          | 0                      |   |   |
| Acetaminophen PO               | A                          | 1                      | 6.75 ± 2.47 (4-10)                            | 7 ± 2.16 (5-10)                               |
|                                | B                          | 2                      |   |   |
|                                | C                          | 1                      |   |   |

# Pain Scores are average of all Child Pugh score per medication

Abbreviations: Intravenous (IV) By Mouth (PO)

\* Medication irrespective of dose

\*\* Post administration pain score was not reported in 3/4 patients

Table 4: Pain Medication Doses and Child Pugh Score

| Medication                   | Total Number of Doses | Number of Patients Receiving Doses (n = 36) |
|------------------------------|-----------------------|---|
| Morphine IV                  | 79                    | 18  |
| Child Pugh A                 | 27                    | 4   |
| Child Pugh B                 | 46                    | 9   |
| Child Pugh C                 | 6                     | 5   |
| Hydrocodone/acetaminophen PO | 61                    | 12  |
| Child Pugh A                 | 13                    | 2   |
| Child Pugh B                 | 39                    | 8   |
| Child Pugh C                 | 9                     | 2   |
| Hydromorphone IV             | 19                    | 6   |
| Child Pugh A                 | 16                    | 4   |
| Child Pugh B                 | 3                     | 2   |
| Child Pugh C                 | 0                     | 0   |

\*n = 36 does not reflect study sample size. Patients could have received more than 1 medication

## Discussion

- There was a large variability in the pain medications including initial drug selected and dosages (Table 2 and 3). Individual dosages were not assessed
- The most common medications used were hydrocodone/acetaminophen and IV morphine. Literature recommends the use of hydromorphone and fentanyl for acute pain due to limited dose reductions and less accumulation in renal dysfunction
- In this study, there was no identifiable trend of prescribing certain pain medications in relation to pain location, pain score, or Child Pugh score.

## Limitations

- Site was a single center location
- Retrospective study design
- Study had a small sample size
- Self conducted chart review was done to gather data

## CONCLUSION

- Patient specific factors such as Child Pugh score and pain scores, used in conjunction with drug pharmacokinetics, is an area to expand research. This could lead to the design of treatment protocols for this patient population through collaboration of physicians treating pain in patients with liver disease and pharmacists providing knowledge in drug pharmacodynamics and pharmacokinetics.

### References:

1. Cirrhosis of the liver: What is it, symptoms, causes & stages. Copyright info: 2022 Cleveland Clinic. <https://my.clevelandclinic.org/health/diseases/15572-cirrhosis-of-the-liver>. Accessed December 20, 2021.
2. Lim YS, Kim WR. The global impact of hepatic fibrosis and end-stage liver disease. *Clin Liver Dis*. 2008 Nov;12(4):733-46, vii.
3. Watkins PB, Kaplowitz N, Slattery JT, et al. Aminotransferase elevations in healthy adults receiving 4 grams of acetaminophen daily. A randomized controlled trial. *JAMA* 2006; 296:87-93.
4. Lewis JH, Stine JG. Review article: Prescribing medications in patients with cirrhosis – a practical guide. *Aliment Pharmacol Ther* 2013; 37:1132.
5. Dean M. Opioids in renal failure and dialysis patients. *Journal of Pain and Symptom Management*. 2004;28(5):497-504. [j.jpainsymman.2004.02.021](https://doi.org/10.1016/j.jpainsymman.2004.02.021)