

# Integrated Care of Alcohol-Related Liver Disease

Gerald S. Winder<sup>\*,†,‡</sup>, Anne C. Fernandez<sup>\*</sup>, Jessica L. Mellinger<sup>§</sup>

<sup>\*</sup>Department of Psychiatry, University of Michigan, Ann Arbor, MI, USA, <sup>†</sup>Department of Surgery, University of Michigan, Ann Arbor, MI, USA,

<sup>‡</sup>Department of Neurology, University of Michigan, Ann Arbor, MI, USA and <sup>§</sup>Department of Internal Medicine, University of Michigan, Ann Arbor, MI, USA



**Background/Aims:** Alcohol-related liver disease (ALD) is the medical manifestation of alcohol use disorder, a prevalent psychiatric condition. Acute and chronic manifestations of ALD have risen in recent years especially in young people and ALD is now a leading indication of liver transplantation (LT) worldwide. Such alarming trends raise urgent and unanswered questions about how medical and psychiatric care can be sustainably integrated to better manage ALD patients before and after LT. **Methods:** Critical evaluation of the interprofessional implications of broad and multifaceted ALD pathophysiology, general principles of and barriers to interprofessional teamwork and care integration, and measures that clinicians and institutions can implement for improved and integrated ALD care. **Results:** The breadth of ALD pathophysiology, and its numerous medical and psychiatric comorbidities, ensures that no single medical or psychiatric discipline is adequately trained and equipped to manage the disease alone. **Conclusions:** Early models of feasible ALD care integration have emerged in recent years but much more work is needed to develop and study them. The future of ALD care is an integrated approach led jointly by interprofessional medical and psychiatric clinicians. (J CLIN EXP HEPATOL 2022;12:1069–1082)

Alcohol remains a primary etiology of liver disease worldwide<sup>1</sup> and a major source of global disease burden.<sup>2</sup> Alcohol-related liver disease (ALD) commonly presents in more advanced stages<sup>3</sup> and its morbidity and mortality are increasing including among young people.<sup>4</sup> At the core of ALD is alcohol use disorder (AUD), a prevalent, chronic, stigmatized, and deadly psychiatric condition which is often severe among ALD patients and comorbid with other psychiatric disorders of mood, anxiety, and personality.<sup>5–8</sup> This broad and serious medical-psychiatric pathophysiology ensures that no single discipline is adequately trained or equipped to manage ALD. This raises important questions about how different models of integrated care can be developed, implemented, and maintained.

Other authors recognize and call for multidisciplinary care to address such a breadth of serious and chronic ALD problems.<sup>8–13</sup> There is a newer and growing literature about the actual implementation and maintenance of integrated ALD care before and after liver transplantation (LT) as well as the pitfalls that can disrupt the work.<sup>7,14,15</sup> While integrated ALD care has obvious application in

cirrhosis and LT where preventing further alcohol-related injury is paramount, it is also relevant and desirable in earlier and less severe disease stages where patients' course toward decompensation and transplant may be delayed or avoided altogether. Care integration is also indicated for at-risk groups, vulnerable populations, and ALD patients with disadvantaged LT access.

This article seeks not only to recapitulate key integrated care principles, preparations, processes, and pitfalls but also to depict in clear and practical fashion the discrete aspects of ALD that require an integrated approach thus equipping the reader with actionable insights and tools to begin their own ALD integration process.

## PRINCIPLES OF CARE INTEGRATION

Blending the expertise of clinicians from different disciplines is an established idea. The integration of mental health services into general medical care, so-called collaborative care, is a model that has been well studied in its feasibility and effectiveness.<sup>16,17</sup> Collaborative care is governed by several key principles: team-driven, population-focused, measurement-guided, evidence-based, and accountability and quality improvement (Figure 1). These same principles apply to integrating psychiatric and addiction services into liver disease management. These ideas also appear in a 2016 joint report from the American Psychiatric Association and the Academy of Consultation-Liaison Psychiatry (formerly Academy of Psychosomatic Medicine) regarding the dissemination of integrated care within adult primary care settings.<sup>18</sup> This report cites studies examining the relative contributions of different roles on a collaborative team (Figure 2).

**Keywords:** integrated, interprofessional, alcohol, multidisciplinary, liver  
 Received: 10.12.2021; Accepted: 22.1.2022; Available online 31 January 2022  
 Address for correspondence: Gerald Scott Winder, Clinical Associate Professor, Departments of Psychiatry, Surgery, and Neurology, University of Michigan, F6319 University Hospital South, 1500 E. Medical Center Dr., SPC 5259, Ann Arbor, MI 48109, USA.  
 E-mail: gwinder@med.umich.edu

**Abbreviations:** ALD: alcohol-related liver disease; AUD: alcohol use disorder; LT: liver transplantation; QI: quality improvement; SUD: substance use disorder

<https://doi.org/10.1016/j.jceh.2022.01.010>

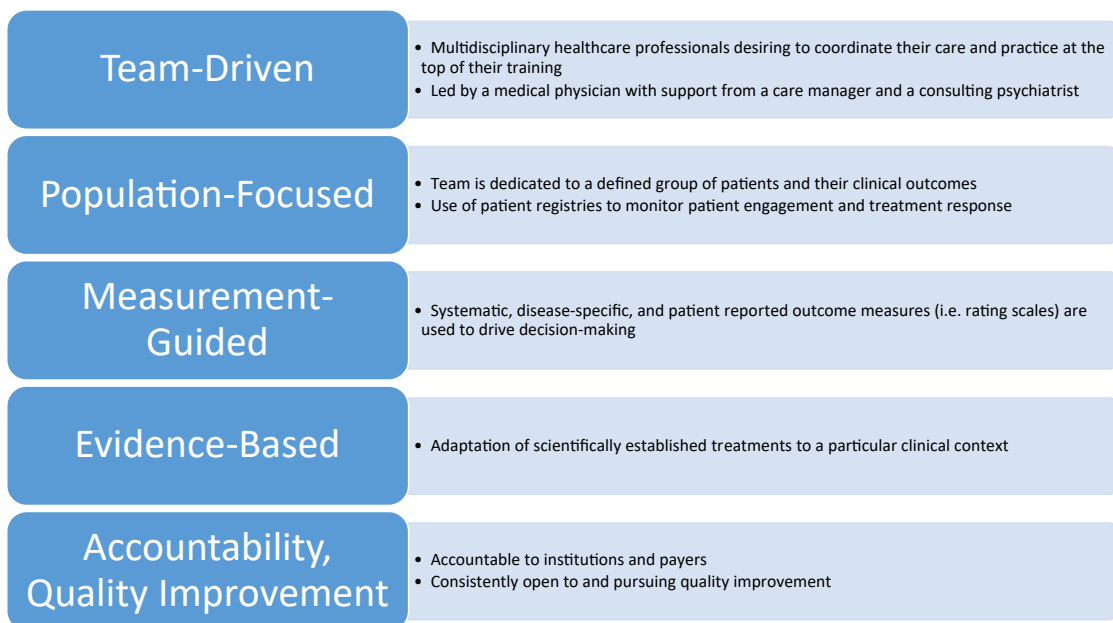


Figure 1 Essential elements of collaborative care (adapted from Vanderlip et al).<sup>18</sup>

### INTERPROFESSIONAL IMPLICATIONS OF ALD PATHOPHYSIOLOGY

Articles have characterized medical and psychiatric facets of ALD pathophysiology;<sup>2,9</sup> it is clear ALD has far-reaching effects on patients and families and places unique demands on care teams and clinicians. Less has been written about the numerous interprofessional implications of ALD (Table 1) or the complicated logistics of integrated and interprofessional treatment models required to address such breadth and complexity.

Psychiatry and addiction clinicians are accustomed to dealing with a wide array of overlapping psychological disorders and symptoms just as hepatology and other medical clinicians regularly manage pathophysiology spanning multiple organ systems. ALD is unique and challenging in the way its clinical treatment requires a holistic view of its interprofessional medical-psychiatric complexity, as a unified core of ALD pathophysiology, rather than parsed or categorized components which can simply be delegated out to diverse specialists. Few would argue that an ideal liver transplant team is one where surgeons, hepatologists, and nurses rarely collaborated. Similarly, a growing number of clinicians in medicine and psychiatry are acknowledging how optimal ALD care models require integration.

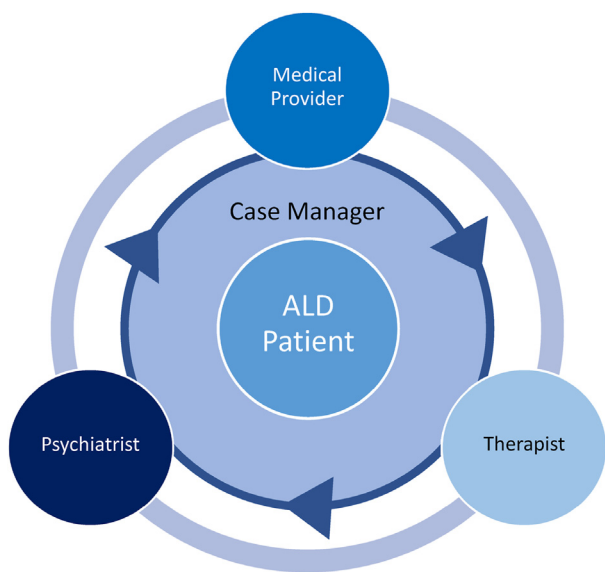


Figure 2 Collaborative care team model on which integrated ALD care models can be based.

### PREPARATIONS FOR ALD CARE INTEGRATION

#### Personal Relationships, Personnel Recruitment, and Team Culture

The foundation of any interprofessional collaboration is a strong *a priori* set of personal and professional relationships shared among clinicians. The importance of constructive interprofessional team dynamics for complex and multifactorial diseases like ALD can be likened to the crucial natures of hand hygiene and surgical sterility for quality bedside care and procedures. Trust, patience, rapport, and good faith catalyze the complicated work of building and maintaining complex and chronic care models for a challenging ALD patient population. Such quality relationships must be an early objective for integrated care and they will not form automatically or easily; they must be deliberately

**Table 1 Interprofessional Implications of ALD Pathophysiology Necessitating Integrated Care.**

Clinical Domain	Description and Context	Interprofessional Implications
Acute alcohol-related hepatitis	<ul style="list-style-type: none"> <li>• AAH has high short-term mortality<sup>23</sup> and any resumed drinking after severe AAH risks substantial mortality<sup>24</sup></li> <li>• A standardized, consensus psychosocial approach to AAH evaluation and ongoing management is still emerging<sup>9</sup></li> <li>• OLT performed in AAH patients have recently increased<sup>25</sup> where psychosocial evaluations are crucial and team collaboration essential</li> </ul>	<ul style="list-style-type: none"> <li>• Hepatology training may not adequately prioritize addiction and mental health and psychiatric training may not provide adequate preparation for end-stage disease and OLT signaling the need for substantial, ongoing interprofessional education</li> <li>• Expert, at-the-ready psychiatric consultation may be sparse for liver teams</li> <li>• Post-discharge AUD treatment must be prioritized and integrated into follow-up planning given relapse and mortality risks</li> <li>• Psychosocial OLT evaluations should be prioritized in AAH; unfavorable influences on psychosocial evaluation include:                         <ul style="list-style-type: none"> <li>○ Prevalent patient AMS</li> <li>○ Patient and family incentives to conceal or minimize psych history due to OLT access concerns</li> <li>○ Compressed transplant and disease timetables</li> </ul> </li> </ul>
Gastrointestinal bleeding	<ul style="list-style-type: none"> <li>• Commonly prescribed SRI medications may increase bleeding risk due to platelet function effects<sup>26</sup> prompting some to advise SRI discontinuation before liver procedures<sup>27</sup></li> </ul>	<ul style="list-style-type: none"> <li>• Risks and benefits of SRI medication use VS discontinuation must be carefully analyzed given competing medical and psychiatric needs                         <ul style="list-style-type: none"> <li>○ High psychiatric comorbidity in AUD/ALD (see later) ensures this will be a common clinical consideration for liver teams</li> <li>○ <i>Either</i> choice (ongoing Rx use, therapeutic interchange, or taper/discontinuation) will necessitate interprofessional follow-up and coordination</li> </ul> </li> </ul>
Hepatic encephalopathy	<ul style="list-style-type: none"> <li>• Insomnia is common in cirrhosis patients<sup>28</sup> and in alcohol users<sup>29</sup> increasing the likelihood that sedating medications may be prescribed to ALD patients which may, in turn, increase HE risk<sup>30</sup></li> <li>• Cirrhosis patients are commonly prescribed BZD and opioids<sup>31</sup> which are HE risk factors<sup>30,32,33</sup></li> <li>• HE is a common and preventable reason for expensive cirrhosis hospital readmissions<sup>34</sup></li> </ul>	<ul style="list-style-type: none"> <li>• Collaboration is needed weighing the mental health risks of ongoing insomnia against the medical risks of using sedating medications</li> <li>• Opioids and BZD have numerous medical and psychiatric implications in terms of dose reduction and discontinuation</li> <li>• Lactulose titration problems are a common and preventable cause of expensive cirrhosis hospital readmissions<sup>34</sup> which could be precipitated by psychiatric or addictive disorders which can unfavorably affect treatment adherence</li> </ul>
Visible Cirrhosis Stigmata (Ascites, Jaundice, Spider Angiomas)	<ul style="list-style-type: none"> <li>• Ascites is a common reason for cirrhosis hospital re-admission<sup>34</sup></li> <li>• The discomfort and appearance of an ascitic abdomen is a common cause for psychological distress and physical pain which ALD patients may not be well-equipped to cope with and manage</li> <li>• Patients may report that others know when they have been drinking based on outward signs of their liver disease</li> </ul>	<ul style="list-style-type: none"> <li>• Fluid imbalances, poor diuretic management, and poor paracentesis planning are common preventable reasons for expensive cirrhosis hospital readmissions<sup>34</sup> which could be provoked by alcohol relapses and/or other psychiatric or addictive disorders which unfavorably affect treatment adherence</li> <li>• Visible signs of ALD may induce patient embarrassment and shame which can impact AUD treatment engagement which is already low<sup>21</sup></li> </ul>
Neuropathy	<ul style="list-style-type: none"> <li>• Neuropathy is a common comorbidity in AUD and ALD patients<sup>35</sup></li> <li>• Untreated neuropathy could lead to physical and psychological distress which, left untreated, could unfavorably influence relapse risks and mental health</li> </ul>	<ul style="list-style-type: none"> <li>• Gabapentin has literature supporting its off-label use for alcohol use disorder and related insomnia and negative affect;<sup>36</sup> it can be used in milder forms of alcohol withdrawal<sup>37</sup></li> <li>• Pregabalin has some evidence for alcohol use disorder treatment and related anxiety; it has shown mixed results in withdrawal treatment<sup>38</sup></li> <li>• Duloxetine is a commonly used Rx in neuropathy patients whose use in severe liver and kidney insufficiencies is discouraged</li> <li>• Thoughtful Rx selection can cover multiple clinical needs with fewer drugs</li> </ul>

(Continued on next page)

Table 1 (Continued)

Clinical Domain	Description and Context	Interprofessional Implications
Overeating, obesity, and eating disorders	<ul style="list-style-type: none"> <li>Metabolic and alcohol-related liver injuries are closely related<sup>39</sup></li> <li>AUD and eating disorders are commonly comorbid<sup>40</sup></li> <li>Bariatric surgery patients are at risk for AUD and ALD<sup>41</sup></li> <li>Many sober ALD patients find themselves newly drawn to unhealthy foods and feeding behaviors</li> </ul>	<ul style="list-style-type: none"> <li>In their commitment to treating liver disease and compulsive behaviors, ALD clinicians must monitor not only SUD but also eating behaviors and weight</li> <li>Topiramate, with its off-label use in AUD,<sup>42</sup> renal clearance, and tendency to induce weight loss, may be an Rx consideration in ALD patients requiring AUD treatment amidst feeding and weight challenges</li> </ul>
Polypharmacy	<ul style="list-style-type: none"> <li>The broad nature of ALD pathophysiology and its comorbidities increases the likelihood of patient polypharmacy</li> <li>Number of medications is predictor of cirrhosis hospital readmissions<sup>34</sup></li> </ul>	<ul style="list-style-type: none"> <li>ALD patients may receive Rx from numerous clinicians including PCPs, psychiatrists, and hepatologists, among others; without coordination, these regimens may be redundant and excessive</li> <li>Peer-to-peer outreach and coordination among clinicians caring for ALD patients take extra effort and may reduce patient Rx side effect burden</li> </ul>
Alcohol withdrawal and AUD relapses	<ul style="list-style-type: none"> <li>Short- and long-term AUD relapse rates vary between 20% and 80%<sup>43</sup></li> <li>Alcohol relapses occur in varying trajectories after OLT<sup>44</sup></li> <li>Access to specialized AUD treatment and Rx is low in ALD patients<sup>45,46</sup></li> <li>Some AUD medications should not be used in liver disease while others must be carefully monitored given risks in end-stage disease and LT<sup>47</sup></li> <li>Many hepatologists do not feel comfortable prescribing AUD medications<sup>48</sup></li> <li>Shorter periods of pre-OLT sobriety (&lt;6 months) have been shown to predict post-OLT drinking,<sup>49,50</sup> but “6-month rules” are discouraged as numerous other factors should be considered<sup>51</sup></li> </ul>	<ul style="list-style-type: none"> <li>Withdrawal can precipitate or complicate any medical clinic and hospital ALD patient presentation</li> <li>ALD teams will need to decide whether BZD (which carry risks of abuse, cognitive and psychomotor side effects) are indicated for withdrawal or if other agents should be used<sup>52</sup></li> <li>Selection and follow-up of AUD Rx is optimal when psychiatric and medical professionals coordinate care given dosing adjustments in liver and kidney disease and OLT-related considerations<sup>47</sup></li> <li>ALD clinicians may understandably feel anger and resentment toward relapsing patients, sentiments which should be appropriately suppressed during clinical encounters and processed elsewhere if needed<sup>15</sup></li> <li>MI is an invaluable communication skillset<sup>53</sup> to mobilize ALD patients toward change behavior; ALD patients uniquely trust their liver clinicians<sup>21</sup> making their use of MI particularly important</li> <li>Liver patients may conceal substance use, particularly if they require OLT; questionnaires and addiction consultation increase liver teams' accurate detection of drinking<sup>54</sup></li> <li>Along with clinical interviews, toxicology, used in accordance with guidelines,<sup>55</sup> is an important tool to corroborate patient reports since liver patients may misrepresent their use<sup>56</sup></li> </ul>
Comorbid non-alcohol SUD	<ul style="list-style-type: none"> <li>Liver disease patients commonly have a history of polysubstance use<sup>57</sup></li> <li>Smoking is common in liver patients and may be increasing; tobacco use has been shown to predict post-LT alcohol relapse<sup>49,50,58</sup></li> <li>Marijuana use is common in ALD patients, increasing in the liver disease population, and associates with other psychosocial risk factors<sup>7,59</sup></li> </ul>	<ul style="list-style-type: none"> <li>ALD teams will frequently need to personalize SUD treatment to include Rx and psychotherapeutic interventions for substances other than alcohol</li> <li>Several SUD medications may be used in ALD patients and require careful consideration in end-stage disease and OLT<sup>47</sup></li> <li>Substance policies in OLT vary widely<sup>60</sup> making interprofessional collaboration key to promote ethical treatment and organ allocation</li> <li>Use of other toxicological markers may be indicated as part of ALD care</li> </ul>
Medical and recreational marijuana	<ul style="list-style-type: none"> <li>Distinction between medical and recreational MJ use can be blurry<sup>61</sup></li> <li>ALD patients have medical foci for pain, psychiatric comorbidities, and SUD meaning the risks and benefits of MMJ will be challenging to discern</li> <li>Many OLT recipients use MJ and CBD<sup>61</sup></li> <li>Transplant center policies regarding MJ are heterogeneous<sup>60</sup></li> </ul>	<ul style="list-style-type: none"> <li>Clinicians should have a working knowledge of MMJ formulations, dosing, and interactions and integrate the topic into their interviews and treatment plans</li> <li>In cases where OLT patients are using MMJ, teams may opt to consult colleagues in the corresponding medical discipline (i.e. neurology when patients are treating neuropathy) to query if an MMJ regimen is optimal</li> </ul>

Table 1 (Continued)

Clinical Domain	Description and Context	Interprofessional Implications
Prescription benzodiazepines and opioids	<ul style="list-style-type: none"> <li>BZD and opioids (alone and in combination) are commonly prescribed in cirrhosis patients including in high doses<sup>31</sup></li> <li>BZD and opioids have been associated with HE<sup>30,32,33</sup></li> </ul>	<ul style="list-style-type: none"> <li>ALD clinicians must not only assess the medical necessity and risks of BZD and opioids but also the impact these drugs have on sobriety and other SUD risks</li> <li>Any alteration or discontinuation of these drugs will have medical and psychiatric implications requiring careful interprofessional coordination and follow-up</li> </ul>
Insomnia	<ul style="list-style-type: none"> <li>Insomnia is prevalent in the end-stage liver disease population<sup>62</sup></li> <li>Insomnia can cause or exacerbate any number of psychiatric conditions and SUD and yet its pharmacological treatment risks worsening HE<sup>30</sup> or drug–drug interactions</li> </ul>	<ul style="list-style-type: none"> <li>Treatment of insomnia will require ongoing interprofessional discussion given the ever-changing risks and benefits of Rx treatment given the dynamic natures of ALD and AUD</li> <li>If available, non-Rx therapies, like CBTi may be prioritized for ALD patients<sup>53</sup></li> </ul>
Suicidal ideation	<ul style="list-style-type: none"> <li>Alcohol is a predictive factor in developing suicidal ideation and is implicated in a high percentage of suicides<sup>64,65</sup></li> </ul>	<ul style="list-style-type: none"> <li>Integrated ALD care entails screening for and responding to self-injurious behavior, SI, and SA meaning teams will benefit from suicide education and assessment training</li> </ul>
Anxiety and mood disorders	<ul style="list-style-type: none"> <li>Anxiety and mood disorders are commonly comorbid with AUD<sup>6</sup></li> <li>Depressive disorders have been shown to impact post-OLT mortality<sup>66</sup></li> <li>Psychiatric comorbidity may predict post-OLT alcohol relapse in ALD patients<sup>50</sup></li> </ul>	<ul style="list-style-type: none"> <li>As stated earlier, SRI medications treating mood and anxiety will need to have their benefits and risks scrutinized alongside other medical and psychiatric factors</li> <li>Alcohol cessation and the stress of liver disease (and OLT) may unmask or worsen mood and anxiety which, in turn, risks alcohol relapse making ongoing psychiatric evaluation and treatment a priority</li> </ul>
Physical, sexual, and emotional trauma	<ul style="list-style-type: none"> <li>PTSD patients are at higher risk of developing AUD; the two conditions are highly comorbid and treatment-seeking rates are low<sup>67,68</sup></li> <li>Many traumatized patients wait years to disclose past trauma which delays treatment and increases distress<sup>69,70</sup></li> </ul>	<ul style="list-style-type: none"> <li>Integrated ALD treatment will often include patients disclosing their history of abuse and/or trauma, sometimes for the first time; it is crucial that clinicians receiving such disclosures attentively listen, believe the patient, and authentically provide support<sup>71</sup></li> <li>Trauma could be a primary etiology and perpetuating factor of ALD making its ongoing pharmacological and psychotherapeutic treatment a priority</li> </ul>
Personality disorders	<ul style="list-style-type: none"> <li>AUD co-occurs with challenging PD characterized by impulsivity and affective dysregulation<sup>72</sup></li> <li>The patient-provider therapeutic alliance in OLT is already complicated by the need for clinicians to advocate for patients while maintaining stewardship over donor organs</li> </ul>	<ul style="list-style-type: none"> <li>PD in ALD can be challenging and may require teams to regularly meet for mutual support and emotional processing, stress and burnout management, and clinical collaboration, similar to a DBT consultation team<sup>73</sup></li> <li>Interprofessional education around PD and their management may be required</li> <li>Addressing PD earlier in the ALD course may increase the likelihood of future patient success in OLT given the challenges PD pose in medical and transplant care<sup>74,75</sup></li> </ul>
Patient deception, defensiveness, ambivalence, denial, and poor insight	<ul style="list-style-type: none"> <li>Misrepresentation of substance use in OLT candidates has been documented<sup>56,76</sup></li> <li>Fear related to transplant access and death may motivate OLT patients to conceal their psychiatric and substance use history</li> <li>Deception can elicit strong emotions in clinicians and may unfavorably affect how they perceive, interact with, and advocate for patients</li> <li>ALD patients tend to have moderate to severe AUD<sup>7</sup> which may correlate with lower levels of alcohol insight and higher defensiveness</li> <li>Ambivalence is a common obstacle in any person's journey toward change</li> <li>Clinicians desire to fix problems and commonly feel a “righting reflex”<sup>53</sup> which can manifest in the overuse of an overly directive style which is often ineffective or counterproductive</li> <li>“Insight” is a commonly used term and parameter in ALD and OLT clinical decision making but remains a poorly defined term</li> </ul>	<ul style="list-style-type: none"> <li>MI training is a valuable skillset whose “spirit”<sup>53</sup> is a helpful guide for rapidly allying with ALD patients:                         <ul style="list-style-type: none"> <li><u>Partnership</u>—collaborative conversation and joint decision-making recognizing that change is done “with” the person, not “to” them</li> <li><u>Acceptance</u>—empathy, affirmation, valuing, and unconditional positive regard for the patient as they are</li> <li><u>Compassion</u>—actively promote the other's welfare and give priority to their needs</li> <li><u>Evocation</u>—patients often already have much of what is needed to change, clinicians must aid in activating their strengths and resources</li> </ul> </li> <li>Observation and critique of teammates' interviews can train, hone skills, and avoid pitfalls like blaming, “gotcha,” or overly paternalistic approaches</li> <li>ALD and OLT teams should agree on a shared meaning of subjective terms like “insight” given its potential impact on decision-making and outcomes</li> <li>A warm team culture facilitates clinicians can acknowledge their frustration with patients, seek and receive colleague support, and accept feedback or correction on their approach to deceptive patients</li> </ul>

(Continued on next page)

**Table 1** (Continued)

Clinical Domain	Description and Context	Interprofessional Implications
Low social support	<ul style="list-style-type: none"> <li>Social problems arising from alcohol use impact QoL,<sup>77</sup> AUD recovery,<sup>78</sup> and aspects of transplant adherence and outcomes<sup>79</sup></li> <li>Poor social support may predict post-OLT alcohol relapse<sup>49,50</sup></li> </ul>	<ul style="list-style-type: none"> <li>ALD teams may recommend that family members consider their own MH treatment and/or seeking support in communities like Al-Anon</li> <li>The 12 steps of AA address aspects of repairing and strengthening relationships</li> <li>Teams must be mindful of particularly mindful of relationship dynamics as ALD patients seek LDLT from their social networks</li> </ul>
Stigma and terminology	<ul style="list-style-type: none"> <li>Stigma among clinicians and the public toward alcohol patients needing OLT has been documented<sup>80</sup></li> <li>The liver community has called for nomenclature adjustments to reduce stigma<sup>9</sup></li> <li>Concerns about alcohol relapse and graft damage have been shown to be the public's main concern about OLT in AAH patients<sup>81</sup></li> <li>A majority of survey respondents were at least neutral in their perceptions of AAH patients receiving OLT<sup>81</sup></li> </ul>	<ul style="list-style-type: none"> <li>ALD clinicians can model respectful terminology in their case discussions and documentation</li> <li>With warmth and patience as well as a robust understanding of current scientific literature, ALD clinicians can work to correct stigmatized attitudes and policies they encounter in their institutions and workplaces</li> <li>ALD clinicians will encounter alcohol and addiction stigma in themselves and the minds of patients and families which they can carefully address as part of clinical care</li> </ul>

AA, alcoholics anonymous; AAH, acute alcohol-associated hepatitis; ALD, alcohol-related liver disease; AMS, altered mental status; AUD, alcohol use disorder; BZD, benzodiazepines; CBD, cannabidiol; CBTi, cognitive behavioral therapy for insomnia; DBT, dialectical behavioral therapy; HE, hepatic encephalopathy; LDLT, living donor liver transplantation; MH, mental health; MI, motivational interviewing; MJ, marijuana; MMJ, medical marijuana; OLT, orthotopic liver transplant; PCP, primary care physician; PD, personality disorders; QoL, quality of life; Rx, prescription; PTSD, post-traumatic stress disorder; SA, suicide attempts; SI, suicidal ideation; SRI, serotonergic reuptake inhibitor; SUD, substance use disorder; VS, versus.

sought out, built, and maintained. ALD clinicians who minimize the importance of such interpersonal factors do so at their own risk due to numerous clinical and logistical challenges (see later) that accompany ALD care integration. Disagreements, adverse clinical outcomes, and miscommunication inevitably occur on any clinical team and may be more pronounced and impactful amidst complex problems with high stakes like ALD. The culture of an integrated ALD team should be one where problems can be discussed openly and respectfully on the way toward a mutually satisfactory resolution.

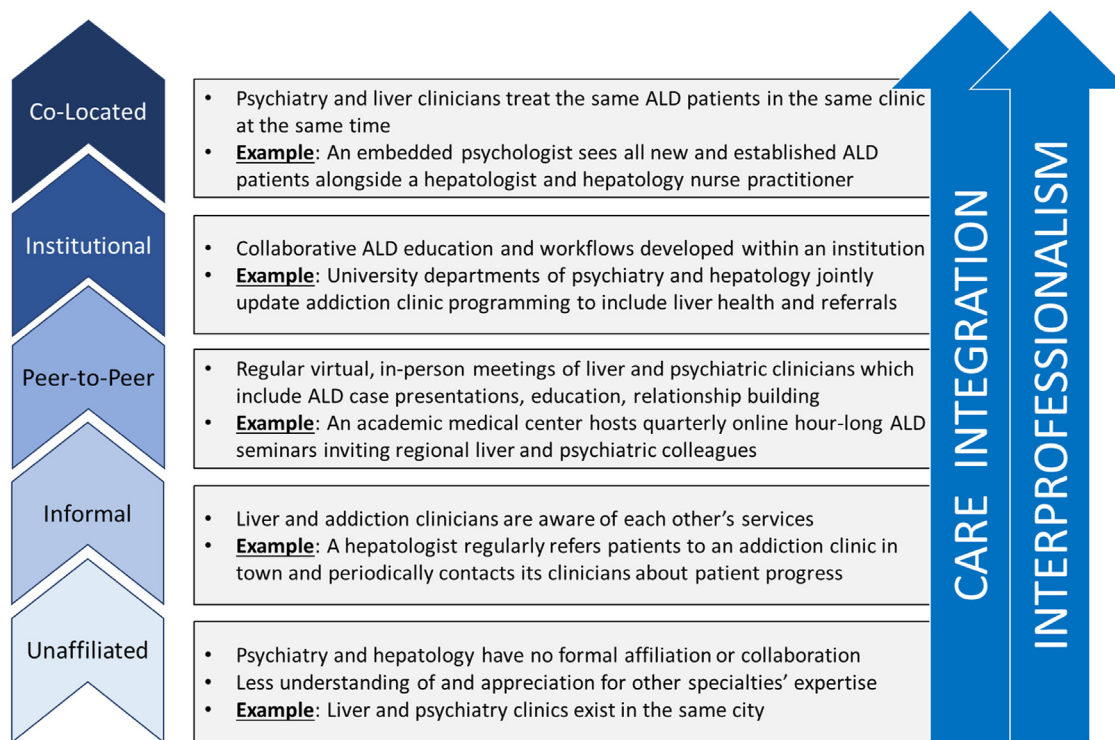
Given alarming ALD epidemiological trends and widespread clinical needs, integrated ALD care models are likely to rapidly grow and require eventual additional multidisciplinary clinician recruitment. Team expansion is an opportunity to deliberately acquire individuals with the interpersonal capacity to participate in and promote interprofessional work alongside their primary clinical and research skillsets. By cultivating a strong spirit of collaboration and interprofessional curiosity, a unique and powerful collaborative culture can arise which can stand out to patients, rotating learners and trainees, and colleagues seeking consultation. All clinical and/or research achievements and recognition should benefit the careers and salaries of every clinician on the team.

### ALD Clinical Model Design and Data Management

Depending on an institution's resources, personnel, and priorities, different models of ALD care may be more

feasible than others. **Figure 3** depicts ascending tiers of ALD care integration and interprofessionalism which clinicians might consider establishing to incrementally improve their clinical services from their current state. Unaffiliated clinicians may simply seek to contact their medical or psychiatric colleagues as the first step to future integration; inviting a colleague to attend seminar to present a talk is a good early strategy. Clinicians who already share some informal connections (periodically reciprocally referring patients to each other's clinics and providing some clinical updates, for example) may wish to build more substantive relations and opt to begin an open-door quarterly seminar where ALD cases are discussed, interprofessional education offered with continuing medical education credits, open discussions held, and relationships developed.

Multidisciplinary clinicians already united within a single institution's mission, facilities, and professional culture may seek to further refine specialty ALD care by developing new specialized connections or workflows among distinct departments and service lines. This could take several forms such as liver clinicians assisting an addiction clinic in developing a curriculum which contains liver topics and hepatology referrals or a psychologist dedicating a half-day per week in her clinic to referrals generated from liver colleagues with whom she regularly corresponds. Arguably, co-location is the epitome of ALD care integration given the high-grade, coordinated specialty care available to ALD patients receiving care from cross-trained clinicians who can easily and rapidly



**Figure 3** Ascending tiers of ALD care integration and interprofessionalism.

combine, adjust, and personalize comprehensive medical and psychiatric treatment plans. **Figure 4** depicts how a morning evaluation clinic might run with each patient remaining in a single examination room while team members rotate. Brief time spent between evaluations allows team members to personalize and fine tune evaluations in interdisciplinary fashion, a capacity unparalleled in other ALD care environments.

Integrated ALD care generates large amounts of multi-modal data including rich qualitative psychosocial data from serial clinical interviews, psychometrics, imaging, vital signs, lab values including toxicology, medical history, and physical examination findings. Attention to data acquisition and management strategies (**Table 2**) in the design phase will ensure that teams can obtain and review a wide array of data to guide clinical care and facilitate research. A case manager role can be an effective way to coordinate integrated ALD care inside and outside the clinic, collate clinical data, and lead multidisciplinary case review meetings (see later). In our era of open notes where patients and families have access to the medical record, teams must be attentive to their verbiage around sensitive topics to maintain therapeutic alliances and honor the spirit of “do no harm.”

### Institutional Leadership Support and Facilities

Integrated ALD care will require resources from the health system in terms of time, money, facilities, and

personnel. The initial group of clinicians seeking to integrate care may consider preparing a written proposal detailing their development plan and then meet face to face with relevant leadership stakeholders (i.e. psychiatry, nursing, transplant, social work, internal medicine) about the next steps. Such meetings are also effective ways to build or strengthen interdepartmental relationships.

It would not be unexpected for an integrated and co-located ALD clinic, for example, to use the physical facilities and clerical resources of a transplant clinic while staffing it with internal medicine physicians, nurses, and advanced practice providers alongside several psychiatry clinicians. Given ongoing severe psychiatric and addiction clinician shortages, salary support flowing from medicine or transplant surgery to psychiatry may be used to secure and protect ALD time allocation. ALD teams may request that physical space be optimized to facilitate integrated care (i.e. aesthetically appropriate interview rooms within a medical clinic, medical equipment, and exam space within an addiction clinic). Such diverse mixtures of resources and personnel will only be possible with ongoing leadership support. Teams will also want to develop workflows around how to refer patients for clinical services that may not be offered within their care venue (i.e. calling ambulances for medical or psychiatric emergencies, referrals to residential rehabilitation or intensive outpatient addiction programming, etc.).

	Patient 1	Patient 2	Patient 3	Patient 4
8:00 AM	Check-in & Rooming	Check-in & Rooming	Check-in & Rooming	
	Hepatology	Psychiatry	Psychology / Social Work	
	Brief Team Discussions	Brief Team Discussions	Brief Team Discussions	Check-in & Rooming
9:00 AM	Psychology / Social Work	Hepatology		Psychiatry
	Brief Team Discussions	Brief Team Discussions		Brief Team Discussions
10:00 AM	Psychiatry		Hepatology	Psychology / Social Work
	Brief Team Discussions		Brief Team Discussions	Brief Team Discussions
11:00 AM		Psychology / Social Work	Psychiatry	Hepatology
		Brief Team Discussions	Brief Team Discussions	Brief Team Discussions
12:00 PM	Wrap up, final treatment planning, and patients sent to lab.			

Figure 4 Sample half-day co-located ALD clinic new patient evaluation schedule.

## THE PROCESS OF ALD CARE INTEGRATION

### Initiation

Precise early steps and processes of ALD integration will vary greatly according to team composition; individual and institutional relationships, resources, and timelines; and the tier and model design selected (Figure 3). There are numerous permutations of how ALD care can incrementally become more integrated from its present state. Several examples illustrate potential integration initiation points and trajectories:

1. A hepatologist presently unaffiliated with any psychiatric or addiction clinicians is increasingly concerned about the size of her ALD patient census and her patients' ongoing drinking. She begins

asking more drinking-related interview questions, dedicates a section of her clinic note to alcohol, and pilots use of a validated alcohol screening questionnaire (Alcohol Use Disorders Identification Test). She submits a written proposal to department leadership for an ALD nurse case manager (Figure 2) to monitor her ALD patients' medical and psychiatric care including acting as a liaison among their various clinicians.

2. A hepatologist perceives discontinuities and inefficiencies in the way her ALD patients access AUD treatment services within her institution. To explore integrative remedies, she contacts one of the health system's psychiatrists to informally brainstorm over coffee about problems and possible solutions.



**Table 2 Data Acquisition and Management Strategies for Integrated ALD Care.**

Information Type	Strategies
Interview data and past medical and psychiatric histories	<ul style="list-style-type: none"> <li>• Agreed-upon divisions in history gathering duties may boost efficiency and increase predictability as to the location and storage of key data points</li> <li>• Well-designed note templates and well-constructed prose ensure that information can be located efficiently for clinical and research purposes</li> </ul>
Laboratory values (including toxicology) and imaging	<ul style="list-style-type: none"> <li>• Teams may opt to design risk-stratified protocols for varying levels of data gathering personalized to medical and psychiatric acuity</li> <li>• Standing orders may increase likelihood that data are reliably obtained</li> <li>• Both medical and psychiatric clinicians can include reminders for labs and imaging in their patient interactions</li> </ul>
Psychometrics	<ul style="list-style-type: none"> <li>• Questionnaires can be pushed to patient through the electronic portal before and between clinic visits</li> <li>• Integrate questionnaires into phone calls between clinic visits where patient status and treatment progress are queried and updated</li> <li>• Collate psychometric data and display in ALD Clinic dashboards alongside medical and toxicological data for review during team meetings</li> </ul>
Psychotherapies and mutual support communities	<ul style="list-style-type: none"> <li>• Elicit and record patient treatment preferences and any past successes, sobriety periods, and unsuccessful treatment</li> <li>• Teams should solicit interval updates from outside therapists and facilities treating ALD patients</li> <li>• Ask patients to keep logs or collect signatures for each recovery community meeting (i.e. Alcoholics Anonymous) attended</li> </ul>
Patient behavior and adherence	<ul style="list-style-type: none"> <li>• Psychosocial clinicians should be documenting and updating meaningful mental status examinations</li> <li>• Documented patient adherence to clinic visits, lab draws and imaging appointments, and clinic communications can be important data guiding treatment planning</li> <li>• In rare instances where patient behavior may impact their ability to continue to access care services, neutral, succinct, and timely clinician documentation should be in the chart</li> </ul>

ALD, alcohol-related liver disease.

3. A hepatologist working alongside psychiatrists, psychologists, and social workers in an established co-located ALD clinic reviews and implements treatment guidelines for AUD management in ALD from the American Association for the Study of Liver Diseases<sup>19</sup> and European Association for the Study of the Liver.<sup>20</sup> She also enrolls in an upcoming motivational interviewing training to improve her alcohol-related conversations with patients and families.

### Scopes-of-Practice

The massive breadth of ALD pathophysiology and its high clinical stakes demand a diverse care team with proportionally broad scopes of practice which overlap and complement each other. For example, both psychiatrists and hepatologists may prescribe AUD medications. Social workers and psychologists both have psychotherapeutic expertise. To reduce frustrating redundancy and confusion and to optimize team operation, team roles must be clearly delineated agreed upon by all parties.

There are many training backgrounds that can fill the general roles and duties pertinent to ALD care. Table 3 reveals not only how multiple training backgrounds can fill similar roles but also the flexible combinations of professionals that could be integrated into innovative ways in different successful models. As more clinicians collaborate with complimentary clinical skillsets and blended scopes of practice, it is easy to conceptualize how new ALD care models can emerge which are simultaneously comprehensive enough to take on a severe and chronic disease like ALD while being flexible enough to meet individual patient needs. ALD patients trust their medically trained clinicians about alcohol and liver matters<sup>21</sup> meaning these clinicians can play an essential role in reducing stigma around mental health and substance use thus facilitating patient interaction with psychiatric colleagues.

The process of further expanding a clinician's scope of practice can be daunting particularly for specialists who have already spent years accumulating and refining

**Table 3 Integrated ALD Care Scopes-of-Practice by Training Background.**

Training Background	Clinical Duties					
	Liver Disease Management	Psychotropic Medication Management	AUD and Psychosocial Evaluation	Motivational Interviewing and AUD Treatment Engagement	SUD and Psychiatric Disorder Psychotherapy	Case Management and Outreach
Hepatologist	X	X		X		
Hepatology APP	X	X		X		
Hepatology RN	X			X		X
Social Worker			X	X	X	X
Psychologist			X	X	X	
Psychiatric APP		X	X	X	X	
Psychiatrist		X	X	X	X	

APP, advanced practice provider; AUD, alcohol use disorder; RN, registered nurse; SUD, substance use disorder.

niche expertise. ALD's vast scope requires its clinicians to "re-generalize" their specialty skillsets to include relevant elements of medical or psychiatric training. It is a sizable request to ask transplant hepatologists and addiction psychiatrists, for example, to pursue additional alcohol and liver training, respectively, for which they will receive no additional pay or licensure designation and amidst all their other personal and professional endeavors. Yet there simply may be no other way to build the interprofessional and integrated teams that ALD care requires.

### Interprofessional Education and Staff Training

Integrated ALD clinicians benefit from additional training by colleagues from other disciplines. Such training can be formal didactics provided in local case conferences or during professional society annual meetings. More often, however, interprofessional education is provided clinician-to-clinician while collaborating on clinical care or research projects. This person-to-person information transfer highlights the unique value in strong relationships, close clinical proximity, and clear and frequent communication. Several scenarios show the methods and value of additional team training:

1. After the announcement of an integrated ALD clinic, clerical and call center staff in hepatology have concerns about receiving calls from ALD patients with psychiatric emergencies trying to reach their psychiatric or addiction clinicians. The ALD psychologist and psychiatrist hold a lunch hour meeting reviewing relevant institutional policy and procedure and invite open discussion. They also encourage staff to contact them directly when additional assistance is needed.
2. Hepatology's more frequent use of alcohol biomarkers raises several questions with medical nurses

tasked with following up on the tests and their results. Nursing leadership invites psychiatry and social work to present on technical and practical aspects of the use of toxicology in patients with addiction. Time is set aside at the end of the meeting for candid discussion.

3. An addiction clinic recently launched a service line dedicated to ALD patients. Psychiatric clinicians are uncertain how to respond to patients presenting with altered mental status. Time is set aside during a weekly team meeting where hepatology presents on hepatic encephalopathy and medical emergencies. A protocol is established by which medical consultation can be obtained when needed.

### Multidisciplinary Team Meetings

An efficiently run and well-attended meeting is the backbone of integrated ALD operations. Two scenarios depict how efficiently-run and well-attended multidisciplinary team meetings can favorably impact ALD care:

1. The case manager of a co-located ALD clinic contacts a patient by phone who missed recent hepatology and psychiatry visits. The psychometric and interview data she gathers during the call indicates the patient is in psychological distress (elevated depression and anxiety questionnaire scores, missing work due to not feeling well). During that week's team meeting, the entire ALD team reviews these data against the backdrop of recent lab trends which include worsening liver function tests and a note from the lab that patient declined toxicology testing. Concerned about a relapse, the team contacts the patient that day for urgent appointments planning to adjust medication and psychotherapy regimens.
2. Academic clinicians from psychiatry, addiction medicine, transplant surgery, and hepatology hold quarterly lunch hour case conferences which discuss ALD/LT

patient cases accompanied by interprofessional didactics. Talks are also streamed online to invited regional colleagues and archived for reference. With her new ALD training and acquaintances, a hepatologist asks a psychiatrist about collaborating on inpatient addiction consultation for the university hospital's GI/liver service.

### Patient Education Materials

Integrated ALD care should be reflected in the education materials that patients receive. Online or written materials orienting patients to their ALD care should feature psychological and behavioral topics alongside those from hepatology and pharmacy. [Supplemental Figure A](#) shows sample pages from an actual patient ALD booklet which moves seamlessly from detailed discussions of liver decompensation, diet, and medications to psychological topics like cycles of habit, coping strategies, and decision-making principles, among others. Mirroring the pathophysiological realities of ALD, patients should ideally perceive little separation between their liver care and their mental health in what they read and experience in clinic.

### Connection to the Transplant Center and Health System

One of the goals of improving ALD care integration is to intervene adequately earlier in the disease process so that patients will not require LT. Many patients, however, will still require LT evaluation and integrated ALD clinics can play a large role in maximizing chances of successful transplant in several ways. Three main LT obstacles and delaying factors for ALD patients are (1) adequate AUD treatment, (2) accumulated sober time, and (3) treatment of psychiatric comorbidities. ALD patients who have previously received integrated care are much more likely to have had these matters addressed prior to LT and have treatment plans in place which satisfy LT listing criteria thus easing the downstream burden on the transplant center. Integrated ALD care elicits and documents large amounts of multimodal data including toxicology which, if accessible to the transplant center, expedites transplant clinicians' understanding of ALD patients' medical and psychiatric pathology enabling them to more adeptly treatment plan and risk stratify. The degree to which ALD clinicians acquire their own relevant transplant knowledge bases and skillsets only further facilitates ALD patient success in transplant.

Integrated ALD clinicians should remain aware of and involved in other addiction needs and initiatives in their health systems. Other institutional initiatives may proffer additional opportunities for ALD care model expansion such as AUD care in less severe liver disease in the primary care setting or in vulnerable populations. Another example

would be a new inpatient addiction medicine consultation service benefitting from ALD-specific training which in turn generates numerous referrals for an integrated ALD outpatient clinic.

The need for integrated addiction care is not confined to AUD treatment in hepatology and other medical or psychiatric colleagues may turn to integrated ALD clinicians for ideas for improving the treatment of other challenging medical substance use disorder (SUD) populations (i.e. patients with comorbid endocarditis and severe opioid use disorder). In fact, lessons learned from ALD care integration could be a return on investment presented to the health system and departmental leadership during planning stages.

### Quality Improvement, Research, and Future Directions

Integrated ALD care is a growing field within hepatology and will benefit from close attention to quality improvement (QI) and study. Several multidisciplinary outcomes and metrics ([Table 4](#)) may be worth tracking as more institutions adopt various types of integrated ALD care models. As described earlier, an open and warm team culture undergirded by strong interpersonal relationships will facilitate QI information becoming rapidly integrated into team interactions and clinical operations toward problem resolution and improvement.

There are several future directions for clinical development and research regarding integrated ALD care.<sup>22</sup> The COVID-19 pandemic has brought about the widespread usage of virtual visits which may play a role in improving integrated ALD care access and adherence. More work is needed to innovate and understand what team professional compositions (i.e. hepatologist + psychiatric nurse practitioner + addiction social worker) and which visit types (in-person, virtual, phone) and frequencies are optimal. Eventual controlled and multicenter studies will be required to properly evaluate novel integrated care models and establish standards of care.

### ALD CARE INTEGRATION PITFALLS

A detailed discussion of potential pitfalls to efforts integrate psychiatric, social, and medical care in end-stage disease and transplant and their possible remedies exist elsewhere.<sup>15</sup> Additional ALD-specific integration pitfalls to consider include over-ambitious patient recruitment resulting in care access problems in a medically ill liver population, vague referral criteria resulting in an overly heterogenous population (i.e. primary non-alcohol SUD, mild liver disease not requiring hepatology care, patients unwilling to speak to psychiatric clinicians), or ill-conceived and/or forced integration efforts which supersede actual clinician

**Table 4 Potential Quality Improvement and Outcome Metrics for Integrated ALD care.**<sup>14,22</sup>

Psychiatric	<p>Psychometric score improvement (QofL, anxiety, depression, sleep, et cetera)</p> <p>Negative toxicology (alcohol, other drugs, nicotine) and rates of discordance with patient reports</p> <p>Reduced alcohol consumption per subjective report and quantitative biomarkers (i.e. PEth)</p> <p>Rates of alcohol treatment engagement, retention, and completion (residential rehabilitation, intensive outpatient, group and/or individual psychotherapy, mutual support group milestones [i.e. AA sobriety coins])</p> <p>Reduced alcohol cravings (subjective report or validated questionnaire)</p> <p>Rates of regained sobriety and alcohol treatment reengagement after relapse</p>
Medical	<p>Improved LFTs (AST, ALT, Tbili, albumin) and MELD scores</p> <p>Reduced healthcare utilization (hospital admissions, ER visits)</p> <p>Rates of improved symptoms of decompensated cirrhosis and averted LT evaluations</p> <p>Reduced mortality</p> <p>Expedited LT timetables (i.e. accelerated time-to-listing)</p> <p>Improved LT outcomes (lower rates of de-listing, rates of pre- and post-transplant relapse, graft failure, hospital readmission, post-transplant return-to-function)</p>
Other	<p>Patient and family satisfaction scores and feedback for in-person and/or virtual visits</p> <p>Clinical and administrative personnel satisfaction scores and feedback regarding teamwork quality and clinical operations</p> <p>Satisfaction scores and feedback from intra- and extramural referring clinicians</p> <p>Cost savings generated from reduced healthcare utilization rates (hospital admissions, ER visits)</p> <p>Patient access and utilization rates (no-shows, cancellations, intra- and extramural referrals, et cetera)</p> <p>Reimbursement and revenue generation</p>

AA, alcoholics anonymous; ALT, alanine aminotransferase; AST, aspartate aminotransferase; ER, emergency room; LFT, liver function tests; LT, liver transplant; MELD, model for end-stage liver disease; PEth, phosphatidylethanol; QofL, quality of life; Tbili, total bilirubin.

buy-in and collaborative capacity. Ideally, ALD clinicians are aware of these humanistic and logistical obstacles early in preparation phases, address them in proposals submitted to leadership, and work to minimize or prevent them during integrated care implementation.

ALD care integration exists in some form in many transplant centers but is much less common elsewhere in hepatology. Given the concerning alcohol epidemiological trends and unique aspects of ALD pathophysiology, the field of hepatology is recognizing the benefits of integrating psychiatric and addiction treatment into liver care. Early models of feasible care integration have emerged in recent years but much more work is needed to develop and study them. If the future of ALD care is an integrated approach, it will be jointly led by medical and psychiatric clinicians who seek to provide collaborative and comprehensive care while seeking their own lifelong interprofessional education and training.

### CREDIT AUTHORSHIP CONTRIBUTION STATEMENT

Dr. Winder—conception of the article, drafting of the original and revising manuscript; Dr. Fernandez—conception of the article, reviewing and editing the manuscript; Dr. Mellinger—conception of the article, reviewing and editing the manuscript.

### CONFLICTS OF INTEREST

The authors have none to declare.

### FINANCIAL SUPPORT

J. L. M. is supported by an NIAAA K23 AA02633301 Career Development Award, and A. C. F. is supported by an NIAAA K23 AA023869 Career Development Award.

### REFERENCES

1. Rehm J, Samokhvalov AV, Shield KD. Global burden of alcoholic liver diseases. *J Hepatol*. 2013;59:160–168.
2. Seitz HK, Bataller R, Cortez-Pinto H, et al. Alcoholic liver disease. *Nat Rev Dis Prim*. 2018;4:16.
3. Shah ND, Ventura-Cots M, Abinales JG, et al. Alcohol-related liver disease is rarely detected at early stages compared with liver diseases of other etiologies worldwide. *Clin Gastroenterol Hepatol*. 2019;17:2320–2329. e12.
4. Tapper EB, Parikh ND. Mortality due to cirrhosis and liver cancer in the United States, 1999-2016: observational study. *BMJ*. 2018;362:k2817.
5. Grant BF, Goldstein RB, Saha TD, et al. Epidemiology of DSM-5 alcohol use disorder: results from the national epidemiologic survey on alcohol and related conditions III. *JAMA Psychiatr*. 2015;72:757–766.
6. Hasin DS, Stinson FS, Ogburn E, Grant BF. Prevalence, correlates, disability, and comorbidity of DSM-IV alcohol abuse and dependence in the United States: results from the national epidemiologic survey on alcohol and related conditions. *Arch Gen Psychiatr*. 2007;64:830–842.

7. Mellinger JL, Winder GS, Fernandez AC, et al. Feasibility and early experience of a novel multidisciplinary alcohol-associated liver disease clinic. *J Subst Abuse Treat*. 2021;108396.
8. Singal AK, Mathurin P. Diagnosis and treatment of alcohol-associated liver disease: a review. *JAMA*. 2021;326:165–176.
9. Asrani SK, Trotter J, Lake J, et al. Meeting report: the Dallas consensus conference on liver transplantation for alcohol associated hepatitis. *Liver Transpl*. 2020;26:127–140.
10. Addolorato G, Mirijello A, Barrio P, Gual A. Treatment of alcohol use disorders in patients with alcoholic liver disease. *J Hepatol*. 2016;65:618–630.
11. Weinrieb RM, Van Horn DH, Lynch KG, Lucey MR. A randomized, controlled study of treatment for alcohol dependence in patients awaiting liver transplantation. *Liver Transpl*. 2011;17:539–547.
12. Addolorato G, Mirijello A, Leggio L, et al. Liver transplantation in alcoholic patients: impact of an alcohol addiction unit within a liver transplant center. *Alcohol Clin Exp Res*. 2013;37:1601–1608.
13. Im GY, Mellinger JL, Winters A, et al. Provider attitudes and practices for alcohol screening, treatment, and education in patients with liver disease: a survey from the American association for the study of liver diseases alcohol-associated liver disease special interest group. *Clin Gastroenterol Hepatol*. 2021;19:2407–2416.e8.
14. Winder GS, Fernandez AC, Klevering K, Mellinger JL. Confronting the crisis of comorbid alcohol use disorder and alcohol-related liver disease with a novel multidisciplinary clinic. *Psychosomatics*. 2020;61:238–253.
15. Winder GS, Clifton EG, Fernandez AC, Mellinger JL. Interprofessional teamwork is the foundation of effective psychosocial work in organ transplantation. *Gen Hosp Psychiatr*. 2021;69:76–80.
16. Archer J, Bower P, Gilbody S, et al. Collaborative care for depression and anxiety problems. *Cochrane Database Syst Rev*. 2012.
17. Katon WJ, Lin EH, Von Korff M, et al. Collaborative care for patients with depression and chronic illnesses. *N Engl J Med*. 2010;363:2611–2620.
18. Vanderlip E, Rundell J, Avery M, et al. *Dissemination of Integrated Care within Adult Primary Care Settings: The Collaborative Care Model*. Washington DC: American Psychiatric Association, Academy of Psychosomatic Medicine; 2016.
19. Crabb DW, Im GY, Szabo G, Mellinger JL, Lucey MR. Diagnosis and treatment of alcohol-associated liver diseases: 2019 practice guidance from the American Association for the Study of Liver Diseases. *Hepatology*. 2020 Jan;71(1):306–333.
20. European Association for the Study of the L. EASL clinical practice guidelines: management of alcohol-related liver disease. *J Hepatol*. 2018;69:154–181.
21. Mellinger JL, Winder GS, DeJonckheere M, et al. Misconceptions, preferences and barriers to alcohol use disorder treatment in alcohol-related cirrhosis. *J Subst Abuse Treat*. 2018;91:20–27.
22. Singal AK, Kwo P, Kwong A, et al. Research methodologies to address clinical unmet needs and challenges in alcohol-associated liver disease. *Hepatology*. 2021 Sep 8 <https://doi.org/10.1002/hep.32143>. Epub ahead of print. PMID: 34496071.
23. Mathurin P, Moreno C, Samuel D, et al. Early liver transplantation for severe alcoholic hepatitis. *N Engl J Med*. 2011;365:1790–1800.
24. Louvet A, Labreuche J, Artru F, et al. Main drivers of outcome differ between short term and long term in severe alcoholic hepatitis: a prospective study. *Hepatology*. 2017;66:1464–1473.
25. Cotter TG, Sandökcö B, Paul S, et al. Liver transplantation for alcoholic hepatitis in the United States: excellent outcomes with profound temporal and geographic variation in frequency. *Am J Transplant*. 2021;21:1039–1055.
26. de Abajo FJ. Effects of selective serotonin reuptake inhibitors on platelet function. *Drugs and Aging*. 2011;28:345–367.
27. Starlinger P, Pereyra D, Hackl H, et al. Consequences of perioperative serotonin reuptake inhibitor treatment during hepatic surgery. *Hepatology*. 2021;73:1956–1966.
28. Hamdan A-J, Al Enezi A, Anwar AE, et al. Prevalence of insomnia and sleep patterns among liver cirrhosis patients. *J Circadian Rhythms*. 2014;12.
29. Chakravorty S, Chaudhary NS, Brower KJ. Alcohol dependence and its relationship with insomnia and other sleep disorders. *Alcohol Clin Exp Res*. 2016;40:2271–2282.
30. Tapper EB, Henderson JB, Parikh ND, Ioannou GN, Lok AS. Incidence of and risk factors for hepatic encephalopathy in a population-based cohort of Americans with cirrhosis. *Hepatol Commun*. 2019;3:1510–1519.
31. Konerman MA, Rogers M, Kenney B, et al. Opioid and benzodiazepine prescription among patients with cirrhosis compared to other forms of chronic disease. *BMJ Open Gastroenterol*. 2019;6e000271.
32. Moon AM, Jiang Y, Rogal SS, Tapper EB, Lieber SR, Barritt IV AS. Opioid prescriptions are associated with hepatic encephalopathy in a national cohort of patients with compensated cirrhosis. *Aliment Pharmacol Therapeut*. 2020;51:652–660.
33. Grünbæk L, Watson H, Vilstrup H, Jepsen P. Benzodiazepines and risk for hepatic encephalopathy in patients with cirrhosis and ascites. *Unit Eur Gastroenterol J*. 2017;6:407–412.
34. Volk ML, Tocco RS, Bazick J, Rakoski MO, Lok AS. Hospital readmissions among patients with decompensated cirrhosis. *Am J Gastroenterol*. 2012;107:247–252.
35. Julian T, Glasgow N, Syeed R, Zis P. Alcohol-related peripheral neuropathy: a systematic review and meta-analysis. *J Neurol*. 2019;266:2907–2919.
36. Mason BJ, Quello S, Shadan F. Gabapentin for the treatment of alcohol use disorder. *Expert Opin Invest Drugs*. 2018;27:113–124.
37. Leung JG, Hall-Flavin D, Nelson S, Schmidt KA, Schak KM. The role of Gabapentin in the management of alcohol withdrawal and dependence. *Ann Pharmacother*. 2015;49:897–906.
38. Guglielmo R, Martinotti G, Clerici M, Janiri L. Pregabalin for alcohol dependence: a critical review of the literature. *Adv Ther*. 2012;29:947–957.
39. Boyle M, Masson S, Anstee QM. The bidirectional impacts of alcohol consumption and the metabolic syndrome: cofactors for progressive fatty liver disease. *J Hepatol*. 2018;68:251–267.
40. Bulik CM, Klump KL, Thornton L, et al. Alcohol use disorder comorbidity in eating disorders: a multicenter study. *J Clin Psychiatr*. 2004;65, 0.
41. Mellinger JL, Shedden K, Winder GS, et al. Bariatric surgery and the risk of alcohol-related cirrhosis and alcohol misuse. *Liver Int*. 2021;41:1012–1019.
42. Guglielmo R, Martinotti G, Quatralo M, et al. Topiramate in alcohol use disorders: review and update. *CNS Drugs*. 2015;29:383–395.
43. Moos RH, Moos BS. Rates and predictors of relapse after natural and treated remission from alcohol use disorders. *Addiction*. 2006;101:212–222.
44. DiMartini A, Dew MA, Day N, et al. Trajectories of alcohol consumption following liver transplantation. *Am J Transplant*. 2010;10:2305–2312.
45. Mellinger JL, Fernandez A, Shedden K, et al. Gender disparities in alcohol use disorder treatment among privately insured patients with alcohol-associated cirrhosis. *Alcohol Clin Exp Res*. 2019;43:334–341.
46. Rogal S, Youk A, Zhang H, et al. Impact of alcohol use disorder treatment on clinical outcomes among patients with cirrhosis. *Hepatology*. 2020;71:2080–2092.
47. Winder GS, Shenoy A, Dew MA, DiMartini AF. Alcohol and other substance use after liver transplant. *Best Pract Res Clin Gastroenterol*. 2020:101685.

48. Cotter TG, Ayoub F, King AC, Reddy KG, Charlton M. Practice habits, knowledge, and attitudes of hepatologists to alcohol use disorder medication: sobering gaps and opportunities. *Transplant Direct*. 2020;6:e603–e.
49. Dew MA, DiMartini AF, Steel J, et al. Meta-analysis of risk for relapse to substance use after transplantation of the liver or other solid organs. *Liver Transplant*. 2008;14:159–172.
50. Chuncharunee L, Yamashiki N, Thakkestian A, Sobhonslidsuk A. Alcohol relapse and its predictors after liver transplantation for alcoholic liver disease: a systematic review and meta-analysis. *BMC Gastroenterol*. 2019;19:150.
51. Shawcross DL, O'Grady JG. The 6-month abstinence rule in liver transplantation. *Lancet*. 2010;376:216.
52. Ait-Daoud N, Malcolm RJ, Johnson BA. An overview of medications for the treatment of alcohol withdrawal and alcohol dependence with an emphasis on the use of older and newer anticonvulsants. *Addict Behav*. 2006;31:1628–1649.
53. Miller WR, Rollnick S, Ebock L. *Motivational Interviewing: Helping People Change*. New York ; London: Guilford Press; 2013:482, 1 online resource (xii).
54. Donnadieu-Rigole H, Olive L, Nalpas B, et al. Follow-up of alcohol consumption after liver transplantation: interest of an addiction team? *Alcohol Clin Exp Res*. 2017;41:165–170.
55. Baxter Sr L, Brown DL, Hurford DM, et al. Appropriate use of drug testing in clinical addiction medicine. *J Addiction Med*. 2017;11:1–56.
56. Webzell I, Ball D, Bell J, et al. Substance use by liver transplant candidates: an anonymous urinalysis study. *Liver Transplant*. 2011;17:1200–1204.
57. Dimartini A, Dew MA, Javed L, Fitzgerald MG, Jain A, Day N. Pre-transplant psychiatric and medical comorbidity of alcoholic liver disease patients who received liver transplant. *Psychosomatics*. 2004;45:517–523.
58. Likhitsup A, Hassan A, Mellinger J, et al. Impact of a prohibitive versus restrictive tobacco policy on liver transplant candidate outcomes. *Liver Transplant*. 2019;25:1165–1176.
59. Likhitsup A, Saeed N, Winder GS, Hassan A, Sonnenday CJ, Fontana RJ. Marijuana use among adult liver transplant candidates and recipients. *Clin Transplant*. 2021e14312.
60. Zhu J, Chen P-Y, Frankel M, Selby RR, Fong T-L. Contemporary policies regarding alcohol and marijuana use among liver transplant programs in the United States. *Transplantation*. 2018;102:433–439.
61. Yan K, Forman L. Cannabinoid use among liver transplant recipients. *Liver Transpl*. 2021 Nov;27(11):1623–1632. <https://doi.org/10.1002/lt.26103>. Epub 2021 Jul 20. PMID: 34018308.
62. Peng JK, Hepgul N, Higginson IJ, Gao W. Symptom prevalence and quality of life of patients with end-stage liver disease: a systematic review and meta-analysis. *Palliat Med*. 2019;33:24–36.
63. Bruyneel M, Sersté T. Sleep disturbances in patients with liver cirrhosis: prevalence, impact, and management challenges. *Nat Sci Sleep*. 2018;10:369.
64. Bagge CL, Littlefield AK, Conner KR, Schumacher JA, Lee H-J. Near-term predictors of the intensity of suicidal ideation: an examination of the 24 h prior to a recent suicide attempt. *J Affect Disord*. 2014;165:53–58.
65. Kaplan MS, McFarland BH, Huguét N, et al. Acute alcohol intoxication and suicide: a gender-stratified analysis of the national violent death reporting system. *Inj Prev*. 2013;19:38–43.
66. Dew MA, Rosenberger EM, Myaskovsky L, et al. Depression and anxiety as risk factors for morbidity and mortality after organ transplantation: a systematic review and meta-analysis. *Transplantation*. 2015;100:988–1003.
67. Blanco C, Xu Y, Brady K, et al. Comorbidity of posttraumatic stress disorder with alcohol dependence among US adults: results from national epidemiological survey on alcohol and related conditions. *Drug Alcohol Depend*. 2013;132:630–638.
68. Debell F, Fear NT, Head M, et al. A systematic review of the comorbidity between PTSD and alcohol misuse. *Soc Psychiatr Psychiatr Epidemiol*. 2014;49:1401–1425.
69. Ruggiero KJ, Smith DW, Hanson RF, et al. Is disclosure of childhood rape associated with mental health outcome? Results from the National Women's Study. *Child Maltreat*. 2004;9:62–77.
70. Smith DW, Letourneau EJ, Saunders BE, Kilpatrick DG, Resnick HS, Best CL. Delay in disclosure of childhood rape: results from a national survey. *Child Abuse Neglect*. 2000;24:273–287.
71. Campbell R, Ahrens CE, Sefl T, Wasco SM, Barnes HE. Social reactions to rape victims: healing and hurtful effects on psychological and physical health outcomes. *Violence Vict*. 2001;16:287–302.
72. Helle AC, Watts AL, Trull TJ, Sher KJ. Alcohol use disorder and antisocial and borderline personality disorders. *Alcohol Res*. 2019;40. arcr.v40.1.05.
73. Noll LK, Lewis J, Zalewski M, et al. Initiating a DBT consultation team: conceptual and practical considerations for training clinics. *Train Educ Prof Psychol*. 2020;14:167.
74. Sansone RA, Bohinc RJ, Wiederman MW. Borderline personality symptomatology and compliance with general health care among internal medicine outpatients. *Int J Psychiatr Clin Pract*. 2015;19:132–136.
75. Laederach-Hofmann K, Bunzel B. Noncompliance in organ transplant recipients: a literature review. *Gen Hosp Psychiatr*. 2000;22:412–424.
76. Bholah H, Bate J, Rothwell K, Aldersley M. Random blood alcohol level testing detects concealed alcohol ingestion in patients with alcoholic liver disease awaiting liver transplantation. *Liver Transplant*. 2013;19:782–783.
77. Lee SB, Chung S, Seo JS, Jung WM, Park IH. Socioeconomic resources and quality of life in alcohol use disorder patients: the mediating effects of social support and depression. *Subst Abuse Treat Prev Pol*. 2020;15:13.
78. Dobkin PL, Civita MD, Paraherakis A, Gill K. The role of functional social support in treatment retention and outcomes among outpatient adult substance abusers. *Addiction*. 2002;97:347–356.
79. Ladin K, Daniels A, Osani M, Bannuru RR. Is social support associated with post-transplant medication adherence and outcomes? A systematic review and meta-analysis. *Transplant Rev*. 2018;32:16–28.
80. Neuberger J. Public and professional attitudes to transplanting alcoholic patients. *Liver Transplant*. 2007;13.
81. Stroh G, Rosell T, Dong F, Forster J. Early liver transplantation for patients with acute alcoholic hepatitis: public views and the effects on organ donation. *Am J Transplant*. 2015;15:1598–1604.

### SUPPLEMENTARY DATA

Supplementary data to this article can be found online at <https://doi.org/10.1016/j.jceh.2022.01.010>.