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Executive Functions, Memory, and Social Cognitive Deficits and Recovery in Chronic Alcoholism: A Critical Review to Inform Future Research

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Abstract

Alcoholism is a complex and dynamic disease, punctuated by periods of abstinence and relapse, and influenced by a multitude of vulnerability factors. Chronic excessive alcohol consumption is associated with cognitive deficits, ranging from mild to severe, in executive functions, memory, and metacognitive abilities, with associated impairment in emotional processes and social cognition. These deficits can compromise efforts in initiating and sustaining abstinence by hampering efficacy of clinical treatment and can obstruct efforts in enabling good decision-making, success in interpersonal/social interactions, and awareness of cognitive and behavioral dysfunctions. Despite evidence for differences in recovery levels of selective cognitive processes, certain deficits can persist even with prolonged sobriety. Herein is presented a review of alcohol-related cognitive impairments affecting component processes of executive functioning, memory, and the recently investigated cognitive domains of metamemory, social cognition, and emotional processing; also considered are trajectories of cognitive recovery with abstinence. Finally, in the spirit of critical review, limitations of current knowledge are noted and avenues for new research efforts are proposed that focus on (1) the interaction among emotion-cognition processes and identification of vulnerability factors contributing to the development of emotional and social processing deficits and (2) the time line of cognitive recovery by tracking alcoholism's dynamic course of sobriety and relapse. Knowledge about the heterochronicity of cognitive recovery in alcoholism has the potential of indicating at which points during recovery intervention may be most beneficial.

Keywords

alcoholism; cognitive impairment; recovery; clinical implications; abstinence

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Conflict of Interest

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In Alcohol Use Disorder (AUD), individuals commonly exhibit a lack of recognition of their disease, minimizing its harmful consequences in life's physical, psychological, and social spheres. Supplanting former characterization of the alcoholic as morally flawed, it is now widely recognized that alcoholism is a complex, multi-dimensional, multi-determined disorder requiring adequate cognitive skills to cope with daily life high-risk situations that threaten maintenance of sobriety, whether it is through controlled drinking or complete abstinence.

To initiate behavioral change towards sobriety, an alcoholic patient often has to (1) recognize the reality of this harmful behavior by perceiving the effects of alcohol on the body (i.e., liver cirrhosis, brain damage), family, work, and community; (2) understand its causes and consequences, and match risks and benefits of behavioral change relative to no change by remembering past personal drinking, craving, and abstinence experiences; and (3) identify and assess new behavioral options, learn general information about alcohol dependence and coping strategies, choose adaptive behaviors, and monitor and control their drinking. The components of change and oversight require efficient cognitive skills involving multi-faceted neuropsychological functions to support abilities for self-assessment, social cognition and emotional processing that enable accurate perception and understanding of the social environment, and reliable memory and executive functioning to achieve optimal decision-making. This paper reviews quantitative studies of component cognitive functions, which are subject to compromise in chronic alcoholism, thereby having the potential to undermine efforts towards achieving and maintaining sobriety.

It has been consistently reported that treatment-seeking AUD individuals have detectable cognitive impairment, often involving executive dysfunction and memory deficits (see reviews by Oscar-Berman et al., 2014 and Sullivan et al., 2010). Recently, new considerations to expand the understanding of functional impairment in alcoholism have included the investigation of metacognition (the ability to accurately assess one's own cognitive abilities), especially metamemory processes, together with emotional and social cognitive abilities (Bora and Zorlu, 2016; Le Berre and Sullivan, 2016; Maurage et al., 2011a; Uekermann and Daum, 2008). In particular, research efforts are warranted for understanding the complex interactions among executive, memory, and social and emotional processing abilities. Also needing explications are how deficits in these processes influence the efficiency of clinical interventions (Blume et al., 2005; Le Berre et al., 2012), cognitive-behavioral therapy (Pitel et al., 2007b) and compromise efforts towards abstinence or controlled drinking (Bates et al., 2002; Bates et al., 2006).

The pattern and extent of cognitive deficits among individuals with chronic alcoholism vary widely, and not all alcoholics demonstrate measurable cognitive impairment (Fein et al., 1990). This heterogeneity is likely at least partly due to the dynamic course of AUD, which is generally marked by periods of withdrawal, abstinence, and relapse. Each of these periods is associated with different levels of functional recovery. Also influential are demographic and disease-related factors such as age, lifetime drinking patterns and amount, and number of withdrawals (cf., Duka et al., 2003). Indeed, recovery of cognitive functions, defined here as the process of returning toward a pre-morbid level of functioning associated with abstinence can occur; however, cognitive deficits can persist even with prolonged sobriety

(Fabian and Parsons, 1983; Fein et al., 1990; Pitel et al., 2009; Rosenbloom et al., 2007; Rourke and Grant, 1999; Stavro et al., 2013; Sullivan et al., 2000a; Yohman et al., 1985). Identifying the pattern, extent, and severity of recovered and persistent cognitive deficits associated with long-term chronic alcoholism with sobriety has the potential to inform brain structure-function plasticity and guide effective management and treatment of AUD.

Reflecting the complexity of normal cognitive functioning, successful performance on most neuropsychological tests requires multiple intact component processes. Parsing complex behavioral functions into their component cognitive processes, their functional building blocks, and examining how alcohol affects these basic processes can indicate which abilities are spared, impaired, recover, or persist with abstinence or continued drinking. Thus, to understand the underlying variation among alcoholism-related cognitive deficits requires a refined characterization of which specific component processes within the broad functional domains implicated are affected.

In the context of the presented overview, the objectives of this paper are three-fold: (1) review evidence for alcohol-related cognitive impairments in affecting component processes of executive functioning, memory, and the recently studied cognitive domains of metamemory, social cognition, and emotional processing; (2) investigate the possible interactions between these functional domains in relation to higher order cognitive processes such as decision making associated with initiation and maintenance of sobriety; and (3) report what is known and not known about the recovery trajectories of these cognitive deficits, and note limitations of our knowledge, thereby identifying avenues for future research.

I- Alcohol-related cognitive impairment

I-1: Executive Functions

Executive functions refer to a number of related but dissociable cognitive processes that enable one to plan, control, and monitor goal-directed and adaptive behaviors in response to novel or non-routine situations (Alvarez and Emory, 2006; Miyake et al., 2000). Specific component executive processes documented as impaired in chronic alcoholism using standard laboratory tasks, including attention, working memory, response inhibition, problem solving, deduction of rules, updating, cognitive flexibility and set shifting, and impulsivity [e.g., (Beatty et al., 1995; Beatty et al., 1993; Chanraud et al., 2007; Fama et al., 2004; Joyce and Robbins, 1991; Le Berre et al., 2012; Loeber et al., 2009; Moriyama et al., 2002; Nixon and Parsons, 1991; Noel et al., 2001; Oscar-Berman et al., 2009; Parsons, 1983; Pitel et al., 2007a; Ratti et al., 2002; Sullivan et al., 2002; Sullivan et al., 1993; Sullivan et al., 2000b; Sullivan et al., 1997; Tarter, 1973; Tarter and Parsons, 1971; Zinn et al., 2004)], each of which can potentially affect higher-order cognitive processes such as decision making and influence initiation and maintenance of sobriety.

Thus, decision-making has been consistently observed to be compromised in AUD under various conditions. These include ambiguous conditions, when the risk associated with a choice is not explicitly stated [e.g., Iowa Gambling Test, (Brevers et al., 2014; Dom et al., 2006; Fernandez-Serrano et al., 2010; Goudriaan et al., 2005; Le Berre et al., 2014; Mazas et

al., 2000; Miranda et al., 2009; Noel et al., 2007; Salgado et al., 2009)] and under risky conditions, when the risk associated with a choice is explicitly stated [e.g., Cambridge Gambling Test, (Lawrence et al., 2009); Cups and Coin Flipping Tasks, (Brevers et al., 2014); ambiguous coin task, (Jung et al., 2014)].

To understand the relative contribution of specific cognitive executive processes to higher order cognitive abilities such as decision-making in AUD, the construct of executive function needs to be deconstructed (Miyake et al., 2000). Severity variability in executive function impairment has been observed across studies likely owing to the complexity of these functions and the use of diverse tasks assessing different constellations of component processes. Indeed, most standard tasks assessing executive functioning are multidimensional and involve several executive function component processes.

Individuals with AUD have difficulty with visual-conceptual and visual-motor tracking skills that intersect with motor processing. Such processes involve psychomotor speed, divided attention, mental flexibility, and set shifting as assessed with the Trail Making Test part B (Chanraud et al., 2007; Davies et al., 2005; Loeber et al., 2010; Moriyama et al., 2002; Noel et al., 2012a; Noel et al., 2001; Nowakowska-Domagala et al., 2017; Oscar-Berman et al., 2009; Zinn et al., 2004). Cognitive inhibition is also particularly affected in chronic alcoholism as assessed with the Stroop Color Word test (Dao-Castellana et al., 1998; Le Berre et al., 2012; Noel et al., 2012a; Noel et al., 2001; Nowakowska-Domagala et al., 2017; Pitel et al., 2007a; Pitel et al., 2007b; Ratti et al., 2002; Tedstone and Coyle, 2004) and a semantic inhibition task, the Hayling test, ((Noel et al., 2012b; Noel et al., 2001). Individuals with AUD have difficulty with abstract thinking, cognitive flexibility, set-shifting, concept identification, working memory, problem solving, and ability to use feedback information assessed with measures such as the Wisconsin Card Sorting Test (Chanraud et al., 2007; Fama et al., 2004; Oscar-Berman et al., 2009; Ratti et al., 2002; Sullivan et al., 1993) and the Modified Card Sorting Test (Le Berre et al., 2012). Deficits in other component executive processes have also been associated with AUD, such as difficulties in organization and self-generation of strategies using verbal fluency tasks and the Ruff Figural Fluency task (Dao-Castellana et al., 1998; Fernandez-Serrano et al., 2010; Oscar-Berman et al., 2009; Pitel et al., 2007a), updating abilities using an N-back task (Noel et al., 2012a; Noel et al., 2001; Pitel et al., 2007a), and working memory using the Letter-Number Sequencing test (Chanraud et al., 2007).

Clinically, the ability to change behavioral schemes and make better choices and decisions in life entails the coordination of many component processes of executive functions. Critical ones include attending, consolidating, and retrieving information about change. With respect to AUD, inhibition of automatic drinking habits would enable change toward favoring new healthy behaviors, to resist temptation and make better choices in the face of high-risk situations and selecting and planning a constellation of behavioral avoidance strategies according to different life situations. From a clinical perspective, when transitioning from excessive drinking to sobriety or controlled drinking, alcoholic patients make different decisions to implement new behavioral schemes to maintain their abstinence or reduce alcohol consumption. The tendency to choose short-term gratification at the expense of long-term consequences suggests that alcoholics may suffer from myopia for the future

(Camchong et al., 2014; Le Berre et al., 2014). This ‘myopia’ may include patients’ awareness that the problems arise from their substance abuse and keep them in denial (Verdejo-Garcia and Perez-Garcia, 2008) or in a form of anosognosia (Le Berre and Sullivan, 2016) about their disorder.

I-2: Memory and Metamemory

Memory is not a unitary process but comprises a multitude of component mnemonic processes, not all of which have been extensively studied in chronic alcoholism (Squire, 1992; Squire, 2004). Over the last half-century, studies in alcoholism have highlighted impairments affecting episodic memory as well as semantic and cognitive procedural learning (Le Berre et al., 2010; Noel et al., 2012b; Pitel et al., 2007a; Pitel et al., 2007b). By contrast, visuomotor procedural and implicit perceptual learning and memory are relatively preserved (Fama et al., 2004; Fama et al., 2012). More recently, deficits in prospective (Griffiths et al., 2012), autobiographical (D’Argembeau et al., 2006; Nandrino et al., 2016) and source (Schwartz et al., 2002) memory have been reported in individuals with AUD. These component processes of memory are explicated next.

Episodic memory involves the mnemonic system founded on the processes of encoding, storage, and retrieval of personally experienced events, associated with a precise temporal and spatial context (Tulving, 2001; Tulving, 2002). Deficits in encoding and retrieval processes occur in recently abstinent alcoholics (Pitel et al., 2007a) and can affect learning of verbal and nonverbal information (Beatty et al., 1995; Everett et al., 1988; Kopera et al., 2012; Schaeffer and Parsons, 1987; Sherer et al., 1992; Sullivan et al., 1992; Sullivan et al., 2000b; Tivis et al., 1995). Episodic memory deficits have been related to executive dysfunction, with poor generation of spontaneous learning or retrieval strategies indirectly affecting free-recall performance (Noel et al., 2012a; Sullivan et al., 1992). A different perspective considers that a genuine episodic memory impairment exists in alcoholics even after accounting for the contribution of executive dysfunction (Pitel et al., 2007a).

Episodic memory is the foundation for conscious recollection of specific personal events from one’s past and the mental projection of anticipated events into one’s subjective future (Wheeler et al., 1997). Recollection of episodic events includes auto-noetic awareness, which is the feeling of re-experiencing or reliving the past and mentally traveling back in subjective time (Tulving, 2001). Sober alcoholics demonstrate a deficit of auto-noetic consciousness (Le Berre et al., 2010; Pitel et al., 2007a) associated with difficulties in retrieving the spatiotemporal context of encoding, with studies reporting compromise of temporal ordering ability (e.g., putting events in chronological order) and spatial context recognition (e.g., remembering where I was when I drank too much last time) (Pitel et al., 2007a; Salmon et al., 1986; Sullivan et al., 1997).

Despite evidence of episodic memory deficits, alcoholics as a group have a tendency to overestimate their memory skills (Le Berre et al., 2016; Le Berre et al., 2010). In particular, they have difficulty in accurately predicting how well they will perform on tasks requiring recognition of newly learned information [feeling-of-knowing, FOK (Hart, 1965)]. They are also likely to generate predictions about their cognitive abilities based on semanticized (implicit) and remote memories of self-ability and poor self-reflection (auto-noetic), and thus

maintain an outdated and unchanged concept of self (Mograbi et al., 2009). The lack of awareness for prospective mnemonic failures suggests a mild form of anosognosia (e.g., you don't know that you don't know) for episodic memory dysfunction and is considered a metamemory impairment (Le Berre and Sullivan, 2016). This metamemory deficit differs from retrospective confidence in memory ability, wherein alcoholics accurately judge how well they recognized newly experienced information [i.e., Retrospective Confidence Judgment, RCJ].

It may be that alcoholics fail to consolidate updated information about their level of memory performance into their personal long-term memory and instead base their predictions regarding current memory performance on outdated self-beliefs that their memory skills are good. A mnemonic anosognosia has been proposed to explain the pattern of metamemory impairment observed in alcoholics without the neurological complications associated with the profound memory impairment of Korsakoff's Syndrome (Hannesdottir and Morris, 2007; Le Berre and Sullivan, 2016; Morris and Mograbi, 2013). These results provide support for the hypothesis that mild mnemonic anosognosia occurs in chronic alcoholism. Overestimation of actual memory abilities can limit benefit from clinical treatment such as cognitive behavioral therapy (CBT) or educational-focused treatment. This recognition disability could place individuals at risk of laboring under the illusion that they have sufficiently consolidated and incorporated into their lexicon essential information acquired during CBT to enable maintenance of their abstinence or reduced alcohol consumption in their daily life.

Semantic memory refers to the ability to recall or recognize facts including personal information, concepts, and general knowledge about the external world, independent of personal experience and spatial/temporal context. In the context of alcoholism, individuals in treatment learn about alcohol and alcohol dependence, the medical and psychiatric consequences associated with excessive alcohol consumption and strategies and techniques to maintain sobriety. Procedural memory for cognitive and behavioral skills that operate at an automatic, unconscious level and independent of episodic memory could also be relevant for successful behavior modification. Over time, AUD individuals supplant new behavioral strategies and procedures to cope with urges and cravings for alcohol and previously entrenched habitual patterns. However, at treatment entry, alcoholic patients with cognitive impairment may exhibit difficulties in acquiring new semantic and procedural information, potentially hampering the efficiency—the essential ability—of cognitive-behavioral therapies (Pitel et al., 2007b), during which patients are taught to anticipate and recognize high-risk situations that could lead to relapse (Assanangkornchai and Srisurapanont, 2007; Berglund et al., 2003; Clay et al., 2008).

Other mnemonic systems impaired in alcoholism include prospective memory (Griffiths et al., 2012), which is the ability to remember to perform an action at a specific point in the future, and autobiographical memory (D'Argembeau et al., 2006), which is memory formed by different types of representation from specific personal events (episodic components) to general knowledge about oneself (semantic component) (Conway, 2001; Tulving et al., 1988). A specific autobiographical memory disorder affecting both the episodic dimension (i.e., long-term memories about specific personal experiences) and the semantic dimension

(i.e., general knowledge about past life events) was observed in recently abstinent alcoholic individuals. This deficit persisted after 6 months of abstinence, and was potentially explained by compromised encoding and consolidation of memories during drinking periods (Nandrino et al., 2016). Also potentially impaired is source memory for recently learned information, which is the ability to discriminate and recall the origin or source of information (Schwartz et al., 2002).

Individuals who labor at maintaining sobriety learn and integrate complex information requiring efficient abilities in a number of mnemonic processes. These include the mental re-experiencing and reliving of craving and emotional and personal states of mind during drinking and abstinence periods. Re-experiencing through mentally reliving episodic drinking experiences include spatial contexts (e.g., at home, a favorite bar) and temporal contexts (e.g., alone, with my 'drinking' friends, when under job related stressful situations).

I-3: Social cognition and emotional processing

Social cognition refers to processes contributing to the perception and understanding of social environments and social interactions (Frith, 2008). Alcoholic individuals can demonstrate deficits in decoding basic and complex emotional facial expressions (Castellano et al., 2015; D'Hondt et al., 2014; Donadon and Osorio Fde, 2014; Kornreich et al., 2001; Kornreich et al., 2002; Maurage et al., 2009; Maurage et al., 2008; Maurage et al., 2011a; Philippot et al., 1999; Uekermann and Daum, 2008). In particular, recognition of negative affect such as disgust and anger can be affected in alcoholism (Bora and Zorlu, 2016), such compromise having the potential to contribute to interpersonal problems in everyday life, for example, misperceiving a facial grimace for aggression (Kornreich et al., 2002).

Impairment in decoding affective prosody and body postures has also been observed in alcoholics (Maurage et al., 2009; Monnot et al., 2002; Monnot et al., 2001). Emotional prosody deficits may be exacerbated when the affective prosody does not match the semantic content in sentences or when trying to match affective prosody to facial expressions (Uekermann et al., 2005). Some alcoholic patients do not benefit from the crossmodal processing facilitation effect, that is, when affective information is conveyed through multiple sensory modalities [e.g., simultaneous auditory (voices) – visual (faces) processing] (Maurage et al., 2007; Maurage et al., 2013).

Alcoholism has been also associated with difficulties in processing humor and irony (Amenta et al., 2013; Uekermann et al., 2007), identifying and describing emotions, i.e., alexithymia, (Stasiewicz et al., 2012; Uzun et al., 2003), and experiencing empathy (Martinotti et al., 2009; Maurage et al., 2011b). In some alcoholics, the emotional component of empathy (i.e., experiencing and sharing emotional states of another person) can be impaired, while the cognitive component of empathy (i.e., understanding mental states of another person) can be relatively preserved (Maurage et al., 2011b).

Deficits in theory of mind (ToM) are consistently reported in alcoholics (Bosco et al., 2014; Maurage et al., 2015; Maurage et al., 2011b; Nandrino et al., 2014; Onuoha et al., 2016; Thoma et al., 2013; Uekermann et al., 2007). ToM enables individuals to predict, anticipate, and interpret the behavior of others (Frith and Frith, 1999; Premack and Woodruff, 1978).

ToM can be divided into (1) affective theory of mind, referring to thinking about affective states, feelings and emotions of others and (2) cognitive theory of mind, referring to thinking about cognitive states, beliefs, thoughts or intentions of others (Shamay-Tsoory et al., 2007). A dissociation between impaired affective ToM but preserved cognitive ToM in alcoholism has been observed (Maurage et al., 2016; Nandrino et al., 2014).

These deficits in social cognition and emotional processing, along with lack of awareness of these deficits, have the potential to contribute to family, social, and career related difficulties exhibited by many alcoholics (Kornreich et al., 2002; Philippot et al., 1999). Alcoholics can misunderstand their relatives' state of mind, leading to family strife, or have trouble recognizing negative affect such as anger from relatives regarding their drinking behavior (Maurage et al., 2009).

II- Recovery of alcohol-related cognitive impairment with abstinence

Sobriety can result in improvement in brain structure and function, indicative of either damage reversal (i.e., actual recovery) or compensatory mechanisms that can be identified with neuropsychological testing and quantitative structural or functional brain imaging. Tracking alcoholism's dynamic course of sobriety and relapse reveals the potential for recovery from and accommodation (i.e., compensation) to neural and neuropsychological insult. Functional imaging studies provide evidence for compensation by invoking non-normal sites and circuits to achieve normal performance on tasks typically impaired (Chanraud et al., 2013; Oscar-Berman and Marinkovic, 2007; Sullivan and Pfefferbaum, 2005), occurring at the cost of processing efficiency when patients perform in the normal range but need additional time to achieve this level (Nixon and Parsons, 1991; Sullivan and Pfefferbaum, 2005). Recovery from cognitive impairment in abstinent alcoholics is typically investigated with cross-sectional designs, comparing alcoholic groups with different lengths of sobriety varying from days to several years to each other or with a control group of healthy participants (e.g., Brandt et al., 1983; Hochla et al., 1982; Markowitsch et al., 1986; Munro et al., 2000; Reed et al., 1992). To assess within-subject change, longitudinal designs, retesting the same group of alcoholic patients and control participants at variable time intervals, are preferred (e.g., Fabian and Parsons, 1983; Glenn et al., 1994; Rosenbloom et al., 2004; Rourke and Grant, 1999; Yohman et al., 1985).

Select executive function component processes showed less impairment as a function of abstinence duration (cross-sectional studies) and demonstrate recovery (longitudinal studies) in alcoholics with several years of sobriety (Fein et al., 2006; Rourke and Grant, 1999) or even only a few months after drinking cessation (Loeber et al., 2010; Pitel et al., 2009). Specifically, inhibition, cognitive abstraction/flexibility, updating processes (Fein et al., 2006; Loeber et al., 2010; O'Leary et al., 1977; Pitel et al., 2009), attention (Fein et al., 2006; Loeber et al., 2010; O'Leary et al., 1977; Sullivan et al., 2000a), and short-term/working memory (Fein et al., 2006; Rosenbloom et al., 2004) show less impairment in long-term abstinent alcoholics compared with short-term abstinent alcoholics and exhibit recovery over time. Other studies, however, reported persistent executive impairment in AUD patients after long-term periods of abstinence from months to years (Munro et al., 2000; Nowakowska-Domagala et al., 2017; Yohman et al., 1985). Decision-making deficits

may also endure in long-term abstinent alcoholics (Ando et al., 2012; Fein et al., 2004); these deficits have been hypothesized to play a significant role in relapse. Stavro and colleagues (2013) highlighted the absence of studies that track the persistence and resolution of impulsive decision-making impairment in alcoholic individuals abstinent for many years.

Similar to persisting executive dysfunctions, cross-sectional studies report episodic memory deficits a few months to one year after drinking cessation (Munro et al., 2000; Parsons et al., 1990; Rosenbloom et al., 2005) and even after several years of sobriety (Brandt et al., 1983) in AUD patients relative to healthy controls. Short-term retention of verbal and nonverbal information was better in individuals with prolonged (5+ years) abstinence, compared with individuals with shorter durations of abstinence (Brandt et al., 1983); however, learning novel pairs of numbers and symbols was still impaired. By contrast, other studies reported improvement in episodic memory after several years of abstinence in AUD patients, who achieved then comparable performance to those of healthy controls (Fein et al., 2006; Reed et al., 1992; Rourke and Grant, 1999). Even in alcoholic patients with at least 6 months of sobriety, a longitudinal study showed normal levels of episodic memory performance when assessed with a selective reminding list learning test (Pitel et al., 2009). Although there is evidence for recovery in selective episodic memory processes such as list learning, other component episodic memory processes such as associative learning can remain impaired even with long-term abstinence (Brandt et al., 1983). To our knowledge, no study has been conducted to investigate cognitive recovery or improvement in semantic and cognitive procedural learning or in metamemory abilities with abstinence.

Deficits can persist with long-term abstinence in social cognition and emotional processing, which are abilities in decoding emotional facial expressions (Kornreich et al., 2001). Contradictory findings have been reported for processes related to ToM, with persistent deficits reported with prolonged abstinence in some (Bosco et al., 2014; Gizewski et al., 2013) but not all studies (Matyassy et al., 2006). Relapse associated with negative affect (e.g., depression) or social pressure represents approximately 70% of loss of sobriety after detoxification (Zywiak et al., 2003), suggesting that consideration of emotional and interpersonal difficulties is essential in clinical treatment for alcoholics. Additional studies are needed to specify the role of emotional and social cognition impairment in these types of relapses and how duration of sobriety contributes to scope and limits of recovery of emotional and social abilities.

In summary, although a number of cross-sectional and longitudinal studies provide evidence that abstinence is associated with improvement in cognitive functions, a meta-analysis of cognitive deficits in alcoholism suggested persistent dysfunctions in multiple cognitive processes even after weeks or months of abstinence (Stavro et al., 2013). The apparent slowdown in the course of recovery in cognitive functions could potentially be explained by factors such as age (Fein et al., 1990; Munro et al., 2000; Reed et al., 1992; Rourke and Grant, 1999), number of previous detoxifications (Loeber et al., 2010), use of cross-sectional design studies, and lack of consensus about the definition of interim drinking criteria used to classify abstainers and relapsers at follow-up.

III- Limitations of Knowledge to Date and Issues to Consider

Cognitive impairments commonly observed in alcoholism span deficits in executive functions and memory processes, including metacognitive difficulties such as metamemory impairment, and deficits in emotional and social cognitive abilities. Potential interactions among these deficits have yet to be fully appreciated or investigated. A new direction in the exploration of alcohol-related social and emotional processing impairments will further our understanding of the psychological processes underlying functional compromise characterizing alcoholism.

Within cognitive domains there remain debates concerning the varieties of component processes most affected in alcoholism. For example, are memory deficits in alcoholism primarily intrinsically mnemonic, or do they have their origin in executive dysfunction? Contradictory findings have emerged with episodic memory impairment in recently abstinent alcoholics not linked solely to executive dysfunction, suggesting genuine episodic memory deficits (Pitel et al., 2007a). By contrast, another study reported that episodic memory deficits were more related to impaired effortful executive processes in alcoholics than in controls (Noel et al., 2012a). To resolve this controversy, future studies need to investigate the interactions among component episodic memory and executive function processes in alcoholics and take account of alcohol history variables, including length of sobriety (cf., Fein et al., 2006), number of withdrawals (cf., Duka et al., 2003), and total lifetime alcohol consumption.

A few recent studies have explored interactions among component cognitive processes and provide evidence that episodic memory and executive component processes can affect higher-order abilities. Together, these processes may explain unawareness of memory impairment (i.e., metamemory decline) in AUD, as compromise of auto-noetic consciousness and strategic mnemonic search abilities were the principal cognitive mechanisms of metamemory decline in recently abstinent alcoholic patients (Le Berre et al., 2010). These compounded deficits could also hamper the learning of new complex semantic and procedural information in alcohol treatment entry and, therefore, potentially could have negative effect on efficiency—the essential ability—of cognitive-behavioral therapies during clinical treatment (Pitel et al., 2007b).

Few studies in AUD have attempted to specify which basic specific cognitive functions support decision-making skills. This ability is reliant on multiple cognitive component processes that can include working memory, inhibition, rule deduction, and reversal learning skills (Dunn et al., 2006), with emotional processes also being potentially influential [for a somatic marker hypothesis of addiction, see (Verdejo-Garcia and Bechara, 2009)]. Among the sparse studies in this realm is one report showing that impaired decision-making performance under conditions of ambiguity could be related to poor response inhibition (Noel et al., 2007). Another study revealed a link between a deficit in decision-making under risky conditions and poor working memory (Brevers et al., 2014). Yet how these compromises affect treatment outcome remains unknown.

With the exception of an identified role of autobiographical memory for ToM abilities (Nandrino et al., 2014) and working memory for humor (Uekermann et al., 2007), the exploration of interactions between cognitive deficits (executive functions and memory) and emotional and social impairments are lacking. Fulfillment of this knowledge gap requires examination with the aim of providing relevant information about cognitive processes underlying these emotional and social alterations. These impairments should be considered as fundamental factors in the emotional and interpersonal difficulties experienced by alcoholic individuals in everyday life and as a consequence in their relapse or maintenance of sobriety.

Regarding the question of cognitive recovery with abstinence in alcoholism, one critical limitation stems from the use of cross-sectional studies to address longitudinal questions. Cross-sectional studies only allow for inferences about cognitive recovery, whereas longitudinal studies provide direct information about cognitive recovery over time. Tracking alcoholism's dynamic course of sobriety and relapse is essential in revealing the potential and limits for recovery of cognitive abilities over time. Longitudinal studies offer the possibility to control for practice, aging, and sex effects when a matched control group is retested at comparable intervals and yield valuable comparisons between abstainers and relapsers. To our knowledge, only cross-sectional studies have been conducted on emotional and social cognition in alcoholic patients.

It has yet to be determined whether the course of recovery of emotional and social cognitive deficits is associated with duration of abstinence or whether the contribution of alcohol toxicity and risk factors are relevant to social cognitive impairment in alcoholism. Only longitudinal studies can impart new insight to these questions.

Further, interim drinking in itself is not clearly defined in the literature (Pitel et al., 2009) such that while some studies consider patients to still be abstainers (Johnson-Greene et al., 1997) having consumed a moderate amount of alcohol, others consider them as having relapsed (Rosenbloom et al., 2004; Rourke and Grant, 1999). Such a binary classification of relapse induces bias in subsequent observations and does not reflect the potential for recovery of relapsers, who have only resumed a limited amount of alcohol consumption without being at a dependent-level.

Whether cognitive deficits in AUD are the result of harmful consequences of excessive alcohol consumption, premorbid risk factors for addiction, or their combination remains unanswered. Family history studies have informed this area, with social cognitive deficits in high-risk individuals with a family history of alcoholism reported, suggesting that emotional and social impairments could be a risk factor in the development of AUD (Hill et al., 2007). Regarding decision-making deficits, a family history study (Lovallo et al., 2006) supported the premorbid vulnerability hypothesis, with individuals having a positive family history for alcoholism (FH+) demonstrating compromised decision-making abilities as assessed by the Iowa Gambling Task compared with individuals who did not have a family history of alcoholism (FH-). Although this finding suggests that deterioration or poor development of the decision-making processes can occur before the emergence of AUD, it does not negate the hypothesis that decision-making is also negatively affected as a result of chronic heavy

drinking. Not all deficits, however, show a relationship with a positive family history of alcohol; for example, ToM impairment was not more prevalent in children of alcohol-dependent parents (Kopera et al., 2014). Exploration of links between cognitive performance and alcohol use variables [i.e., length of alcoholism, usual daily alcohol intake or number of withdrawals (cf., Duka et al., 2003; Loeber et al., 2009) could also constitute other ways to investigate this question. However, alcohol consumption information can only be estimated through historical interview, which is limited by recall accuracy, possibly underlying why such relationships have often eluded detection in alcoholics. Again, longitudinal studies would best inform this area of inquiry.

Finally, metamemory paradigms have successfully highlighted the lack of awareness of memory deficits in alcoholics early in abstinence (Le Berre et al., 2016; Le Berre et al., 2010). Because metamemory is a complex and multidimensional higher-order cognitive function, involving processing components relevant for monitoring and controlling memory (Flavell, 1971; Flavell and Wellman, 1977; Nelson and Narens, 1990), a theoretical framework (Nelson and Narens, 1990) was proposed that involved control and monitoring processes invoked during acquisition, retention, and retrieval of information. The control component refers to regulation applied during a mnemonic activity to improve memory performance, such as selection and use of strategies or decisions on allocation of time and cognitive resources, depending on task demands. The monitoring component assesses the progress and the success of memory functioning. Heretofore, metamemory abilities were essentially investigated by monitoring measures (i.e., FOK and RCJ measures) focused on the memory retrieval phase. Therefore, the exploration of metamemory in alcoholism could be extended by using other monitoring measures such as the ‘judgment of learning’ assessed during the learning phase and by considering the control component of metamemory.

CONCLUSION

Alcoholism is a complex, dynamic disease punctuated by periods of abstinence and relapse, and influenced by multiple vulnerability and resilience factors. This review highlights the cognitive deficits in executive functions, memory, and metacognitive abilities, with associated impairment in emotional processes and social cognition that occur in some alcoholics, and the variable recovery that occurs over time with abstinence. Despite extensive study of cognitive impairment and recovery, knowledge lacunae abound. For example, it is now critical to investigate the emotional and social components contributing to functional impairment in chronic alcoholism. One focus might be on the interaction among emotionally-based cognition processes and identification of vulnerability factors that play a role in the development of emotional and social processing deficits. Treatment focusing on improving level of awareness about impaired and spared cognitive and emotional processing deficits may reveal how an alcoholic compensates for functional compromise, similar to the approach taken by traumatic brain injury (TBI) rehabilitation programs.

To date, studies on cognitive recovery have largely included individuals with long-term sobriety, whereas alcoholics who relapse, who are notoriously difficult to track, are less often studied. Relapses (i.e., the resumption of alcohol drinking following a period of abstinence), however, are a crucial part of the addiction process and deserve attention,

especially when after a period of abstinence, alcoholic patients who relapse may experience further decline in cognitive functioning (Pitel et al., 2009). Another factor seldom considered is the population studied. Specifically, most studies have examined treatment-seeking alcoholics, but this group reflects only about one-quarter of individuals with alcoholism (Smith and Fein, 2010).

An evolving aim of neuropsychology with respect to this multifactorial disease is the ability to identify alcoholics who are at particular risk of functional impairments in order to customize clinical treatment to increase the likelihood of sustained abstinence. Research focused on determinants or risk factors of cognitive deficits in alcoholics is even more urgent in light of the potential interactions and relations among vulnerability factors, alcohol consumption variables, and severity of cognitive and emotional impairment that have been elusive to or exclusive of quantitative, objective study. Prospective longitudinal studies, such as the NIH/NIAAA-supported National Consortium on Alcohol and NeuroDevelopment in Adolescence (NCANDA) (Brown et al., 2015), the Collaborative Studies on the Genetics of Alcoholism (COGA) (Begleiter, 1995) that study adolescents before initiating appreciable drinking, and now ABCD which is a longitudinal prospective study starting in preadolescence, can be particularly valuable by providing information to address the question of whether cognitive deficits observed in AUD are the harmful consequences of excessive alcohol consumption or a premorbid risk factor for addiction.

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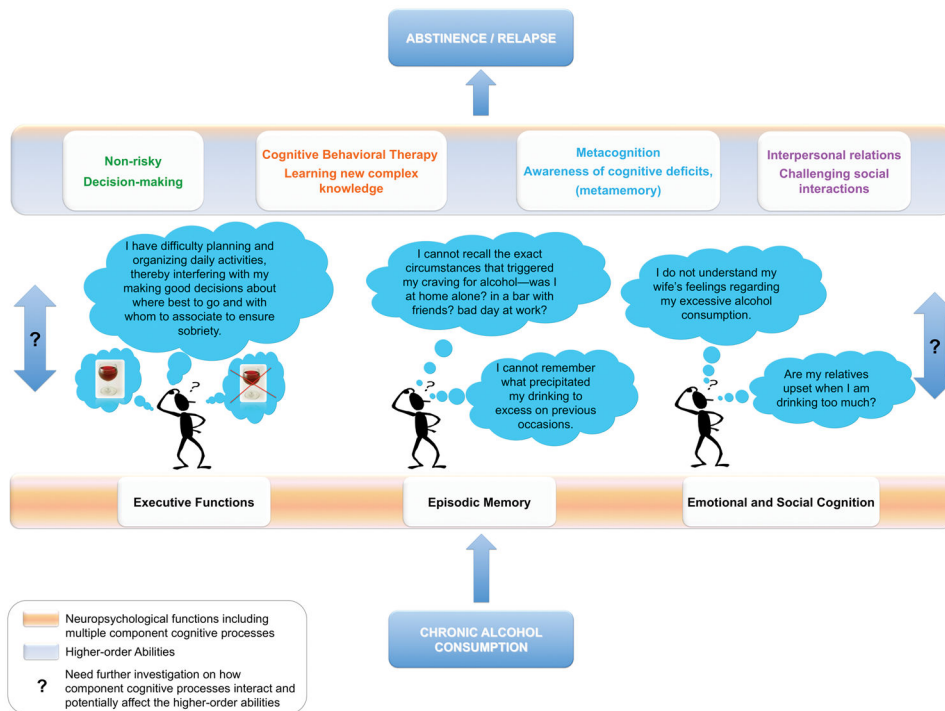


Figure. Neuropsychological functions affected in chronic alcoholism compromising efforts to initiate and maintain abstinence by potentially hampering efficacy of clinical treatment (e.g., cognitive-behavioral therapy requiring learning new complex knowledge) and potentially obstructing efforts in enabling other higher-order abilities such as non-risky decision-making, success in interpersonal/social interactions, and awareness of cognitive and behavioral dysfunctions (i.e., accurate metacognition). Blue arrows with question mark = Need of further investigation on how component cognitive processes interact and potentially affect the higher-order abilities.