

one group adhered, the less likely they were to drink heavily by the end of the first month.

This association was not seen in the placebo group.

Generally side effects were reported more often in the naltrexone group than in the placebo group, but not among the heavy drinkers.

The one exception was that nausea was reported more often among the naltrexone patients in the heavy-drinking group.

It is possible that naltrexone-related nausea may be involved in the reduc-

tion in heavy drinking, but the current study design did not allow for distinguishing between nonspecific nausea and nausea related to drinking alcohol, the investigators commented.

Levels of the liver enzymes aspartate aminotransferase (AST) and alanine transaminase (ALT) did not differ between the naltrexone and placebo group at baseline or at 1 month of treatment, nor did the percentage of individuals with levels out of the normal range: 3.6% of the naltrexone group and 2.7% of the placebo group were outside the normal range for AST at

week 2, as were 10.7% and 18.9%, respectively, for ALT. "These findings support the hepatic safety of naltrexone in people with nicotine dependence and may be reassuring to treatment providers concerned about potential liver toxicity with naltrexone," Dr. King and her associates said.

This study was funded by the National Institute on Alcohol Abuse and Alcoholism and the University of Chicago's Cancer Research Center and General Clinical Research Center.

The authors reported having no conflicts of interest. ■

Alcohol Abuse May Lead to Depression

BY MARY ANN MOON

Alcohol abuse and dependence appear to lead to major depression, rather than vice versa, according to a collection of statistical analyses reported in the Archives of General Psychiatry.

Researchers used data from the ChristChurch Health and Development Study, a cohort of 635 boys and 630 girls born in urban New Zealand in 1977 and followed through age 25, to examine the well-known relationship between alcohol abuse or dependence and depression. They used several advanced statistical modeling methods to explore possible causal pathways between the two disor-

Subjects who abused alcohol were nearly twice as likely to fulfill criteria for major depression as were those who did not abuse alcohol, the statistical analyses showed.

ders, said David M. Fergusson, Ph.D., and his associates at the University of Otago, Christchurch.

At age 24-25 years, approximately 14% of the sample met DSM-IV criteria for alcohol dependence (6%) or abuse (8%), and 14% met criteria for major depression.

At all ages, there were clear and significant trends for alcohol abuse to be associated with depression, such that subjects who abused alcohol were nearly twice as likely to fulfill criteria for major depression as were those who did not abuse alcohol. In contrast, major depression did not appear to predispose subjects to alcohol abuse.

In addition, the relationship between alcohol abuse and depression was not found to arise from some common factor underlying both disorders, Dr. Fergusson and his colleagues said (*Arch. Gen. Psych.* 2009;66:260-6).

It is possible that the use of alcohol may trigger a response in genetically susceptible individuals that raises the risk of depression. Other research also suggests that alcohol's action as a depressant may lead to periods of depressed affect among drinkers, the investigators added.

The researchers noted that these results contradict those of previous studies, some of which have suggested that the causal relationship moves in the opposite direction because some people with depressive symptoms self-medicate with alcohol.

The current study might be more able to detect the true direction of the causal relationship because of its use of numerous complex statistical methods as well as its use of repeated structured mental health assessments over time, Dr. Fergusson and his associates said. ■

Defining the role of alpha-2A receptors within ADHD

New preclinical science suggests that stimulation of alpha-2A receptors located throughout the prefrontal cortex (PFC) strengthens executive function including working memory, which is thought to play an important role within ADHD.¹⁻³

Our current understanding of ADHD treatment includes, in part, increasing levels of norepinephrine that act at the alpha-2A receptor.¹ Directly engaging these receptors is thought to exert a positive effect on cognitive functioning, such as behavioral inhibition and impulse control.^{1,4}

As we continue to learn more about ADHD, we must consider the emerging role of the alpha-2A receptor—**it's big.**