Infectious Diseases

NICU Is Ideal Setting for Giving Parents Flu Vaccine

BY ERIK GOLDMAN Contributing Writer

OLD GREENWICH, CONN. — The neonatal intensive care unit is an ideal setting for delivering the trivalent influenza vaccine to parents of high-risk infants, Dr. Shetal Shah said at a meeting of the Eastern Society for Pediatric Research.

Flu vaccination rates among U.S. adults remain very low. Even among high-risk adult populations, such as the elderly or health care workers, full immunization rates run between 25% and 33%. Influenza is a very common and growing problem among infants in the NICU setting; the babies pick up the virus from adult caretakers. There has been a 10% increase in flu-related hospitalizations among vulnerable infants, said Dr. Shah of the Neonatal Intensive Care Unit at New York University, New York.

"If adults need to be immunized in order to protect their children, but they're

The hospital may be one of the few places to reach fathers. 'The ob.gyn. community is doing a much better job of getting the flu shots to women.'

not getting the vaccines, need to look at reasons the why," he said. Although vaccine shortages have played a role, by far the most common reason for failure to receive the shots is inconvenience. Busy parents are preoccupied

with so many other concerns that obtaining flu shots tends to fall to the bottom of the priority pile.

"Last year, we surveyed the parents of our NICU patients, and the flu vaccine rate was only around 33%. More than half who were not immunized cited inconvenience as the main reason." This prompted Dr. Shah and his colleagues to consider making the shots available right in the NICU.

It really is an ideal setting for this type of intervention. For one, we're dealing with the highest risk babies with the greatest need for protection. Most NICUs, like ours, are adopting a family-centered care model that actively engages the parents in the care of their babies. We have very liberal visiting hours at NYU: Parents can be in the NICU with their children for 22 out of every 24 hours. And we're open at times when other clinics or care delivery settings are closed," he said.

In addition, the NICU may be one of the few places to reach fathers. In general, men are far less likely than women to get regular medical checkups, and they tend to avoid physicians, hospitals, and clinics. "The ob.gyn. community is doing a much better job of getting the flu shots to women than the pediatric or family practice communities," Dr. Shah said at the meeting, cosponsored by the Children's Hospital of Philadelphia. In part, this has to do with the success of prenatal care programs, which seldom reach the fathers.

The NYU NICU team undertook a pilot project to provide trivalent flu vaccines for all NICU parents from November 2005 to March 2006. As part of the admission process, staff told parents that it was possible to get the flu vaccine, free of charge, right there in the NICU, and tried to get the parents to consent prior to delivery or soon thereafter. In addition, they also posted signs right on the babies' warmers, stating that the hospital strongly recommended that parents obtain the shot. They also posted reminders in common areas and breast-feeding rooms.

During the 4-month period, the NYU staff admitted 273 parents of 158 babies. They were able to counsel 220 about the importance of immunization and actually gave shots to 157 (71%). Fifty-two (24%) of the parents counseled had already been immunized, and 11 (5%) refused.

Of those vaccinated in the NICU, 61% got their shots within 2 days after their babies were admitted. "We got to most of them within 72 hours, and after that, it tended to drop off."

They also were most successful with parents of babies who were less than 28-32 weeks' gestational age at delivery. Dr. Shah attributed this to the much longer lengths of stay for this extremely premature subgroup.

Most of the parents accepted the importance of getting the shots, and Dr. Shah noted something of a peer-pressure effect. "People bond in the NICU with other people going through the same ordeal. So if one couple went for the shots, the others

CRITICAL INSIGHTS INTO THE NATURE OF **NICOTINE ADDICTION**

A SUMMARY OF KEY LEARNINGS TO DATE

With all the public awareness efforts that have been made, and with all the truths that have come to light over the last several decades about the dangers of smoking, one obvious question lingers: Why are people still smoking?

Understanding nicotine addiction

Smoking

is a chronic,

relapsing

condition

Most experts agree at this point that smoking is a chronic, relapsing condition—an addiction similar in nature to that seen in cocaine and heroin users.^{1,2} Following are 4 criteria the Surgeon General has used to define addiction, along with an explanation of how nicotinespecifically smoking—meets these criteria.2

1. Addiction leads to compulsive use, despite adverse consequences

According to a 1988 Surgeon General's report, "highly controlled or compulsive use indicates that drug-seeking and drug-taking behavior is driven by strong, often irresistible urges. It can persist despite a desire to quit or even repeated attempts to quit."2 Smoking statistics show that approximately 70% of current smokers report that they want to quit; however, only about 5% of smokers who try to quit without medical aid succeed.3,4 In fact, the average smoker has tried to quit 6 to 9 times.⁵ It is common for people to continue smoking despite known negative health consequences. In fact, smoking behavior often persists even after the presentation of comorbid conditions.^{2,6,7}

2. Addiction involves a psychoactive substance with reinforcing properties

The psychoactive (mood-altering) properties of nicotine are substantially related to its effect on the mesolimbic dopaminergic system. For delivery of nicotine, smoking is the most efficient mechanism. In a matter of seconds, nicotine from inhaled smoke crosses the blood-brain barrier and begins altering brain chemistry through binding to cholinergic receptors normally activated by acetylcholine. Dopamine is released in the nucleus accumbens, triggering central nervous system effects such as pleasure, relief of anxiety, better task performance, and improved memory. These rewards serve to reinforce smoking behavior.^{2,8-10}

Complicating this effect is that the routines associated with smoking, such as smoking in social environments, can also come to be reinforced through the pleasure response. Eventually, the pleasure associated with smoking in these settings acts as a subconscious trigger, making it hard for the smoker to dissociate the behavior from the addiction. This explains why successful quit attempts often require some degree of behavioral modification.^{2,11,12}

3. The addicted subject develops tolerance

Nicotine initiates its action by competitively binding at the nicotinic acetylcholine receptors (nAChRs), ligand-gated ion channels on the cell membrane. Compared with the endogenous agonist acetylcholine, nicotine causes a prolonged activation of nAChRs. The activation is followed by a desensitized

state in which the receptors are unresponsive to agonists. This process has been compared to tripping a circuit breaker. 10,11,13

Chronic use of nicotine leads to chronic desensitization of nAChRs. As more nicotine is consumed, and more receptors become desensitized, the user experiences a diminished pleasure effect with each subsequent cigarette smoked. As the response decreases, increasing levels of nicotine are required to achieve a consistent, desired effect.^{2,10-12} These are defining characteristics of tolerance.¹⁴

they had connected with often followed."

Despite the usually frantic pace of activity in the NICU, the staff took the flu shot endeavor quite seriously and managed to find the time to talk often with the parents. "We kept track of who was and who was not getting the shots, and we would talk to the ones who had not to see if they had any questions or concerns that we could address."

Among the 11 who refused the vaccine, 5 stated that they simply did not believe in immunization, and 2 said they feared that the shots might induce autism. Others cited religious objections or a reluctance to add anything else to whatever medical care they were already receiving. One cited an allergy to eggs, which is a legitimate concern because the vaccine contains some egg proteins.

Overall, the NYU NICU-based flu shot program was highly successful. Dr. Shah and colleagues hope to do a follow-up to see if the program had any impact on the rate of influenza among the neonates. He cautioned, however, that the sample size may be too small to support any definitive conclusion.

This pilot program did, however, prove that flu shots can be effectively distributed in the NICU setting to parents, who for a variety of reasons, had not previously gotten immunized. The program created very little additional strain on NICU physicians or nursing staff.

"Administration of the trivalent vaccine is very possible in a busy NICU, and implementation markedly increased compliance with recommendations aimed at protecting high-risk neonates," Dr. Shah told conference participants. "There will always be a small subset of parents who will refuse, no matter what. But we can get to many parents who are willing to take the shots." He added that this type of program is highly replicable and could be quickly implemented in any familycentered NICU.

4. An addictive substance causes physical dependence, as evidenced by withdrawal and relapse

The symptoms of nicotine withdrawal have been clearly identified and confirmed. For most smokers, these symptoms include at least one, if not several, of the following: craving, irritability, insomnia, headache, anxiety, depression, and impaired concentration. 11,14 These withdrawal symptoms have been identified as key contributors to relapse, as the smoker often "self-medicates" with nicotine to return to a perceived state of normalcy.12

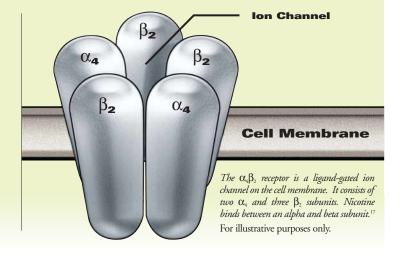
Additionally, chronic stimulation at the receptor site is believed to be responsible for upregulation (an increase) in the number of receptors expressed at the cell surface.8,10,12 This is likely a result of the brain compensating for the desensitization of existing receptors, as described earlier.

As nicotine leaves the system, however, desensitized receptors can return to an "open" state in which they are once again susceptible to stimulation.^{10,11} The combination of these factors—ie, a greater number of available, sensitized receptors may create "an excess excitability of the nicotinic cholinergic systems of smokers."12 This hyperexcitable state is believed to contribute to the smoker's motivation to smoke another cigarette (craving).9,12

Hyperexcitability may also explain why the first cigarette smoked following a period of abstinence provides a more intense pleasure response for the smoker.^{11,12} Note, for example, that most smokers derive the greatest pleasure from their first cigarette of the day. 10,12 In fact, smoking a single cigarette following a cessation attempt often prompts a complete relapse to heavy smoking. 10,111

The $\alpha_{4}\beta_{2}$ receptor

Recent evidence suggests that scientists have identified a specific nAChR in the brain that is believed to act as a primary mediator of the addictive properties of nicotine—the $\alpha_4\beta_2$ receptor.15-17 The isolation and characterization of this receptor is a significant advancement in the understanding of the neurobiology of smoking addiction.



Conclusion

Smoking is a chronic, relapsing condition. For most smokers, the compounding effects of behavioral, psychological, and physical triggers make overcoming their addiction extremely difficult. However, given the high morbidity and mortality related to smoking,^{3,8} getting smokers to quit is important. Proactive medical intervention for smokers may be beneficial.¹ Recent advancements in the study of nAChRs—specifically the identification and characterization of the $\alpha_4\beta_2$ receptor—represent a significant advancement in the understanding of the nature of nicotine addiction.

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Multidrug Resistance Stalls TB Drop

BY MIRIAM E. TUCKER Senior Writer

uberculosis cases reached an alltime low in the United States in 2005, but progress toward elimination of the disease has slowed, according to the Centers for Disease Control and Prevention.

Moreover, the number of multidrug-resistant (MDR) TB cases increased 13.3% from 2003 to 2004, marking the largest 1vear increase in such cases since 1993. A greater proportion of foreign-born patients than U.S.-born patients had MDR TB, the CDC said (MMWR 2006;55:305-8).

In 2005, a total of 14,093 TB cases was

More than half of the 7,656 foreign-born TB patients in 2005 were from Mexico, the Philippines, Vietnam, India, and China, the CDC said.

reported in the United States, representing a decline of 3.8% from 2004 and the lowest recorded rate (4.8 per 100,000 population) since national reporting began in 1953. However, the decline slowed from an average of 7.1%

per year during 1993-2000 to 3.8% per year during 2001-2005.

In 2005, the TB rate in foreign-born persons in the United States was 8.7 times that of U.S.-born persons. Although the total foreign-born population in the United States has increased 61.6% since 1993, the number of TB cases reported in this population hasn't changed substantially, resulting in a 36.0% decline in the TB rate among foreign-born persons. More than half of the 7,656 foreign-born TB patients in 2005 were from Mexico, the Philippines, Vietnam, India, and China, the CDC said.

Race/ethnicity data showed that TB rates in 2005 were increased 19.6 times among Asian Americans, 8.3-fold among blacks, and 7.3 times among Hispanics, compared with whites. But rates declined in almost all racial and ethnic populations from 2003 to 2005, with the most decline among American Indians/Alaska Natives (14.4%) and Asian Americans (14.1%).

The number of MDR TB cases increased from 113 cases in 2003 to 128 in 2004, the most recent year for which complete drugsusceptibility data are available. In 2004, 0.6% of U.S.-born and 1.6% of foreign-born TB patients had MDR TB. Approximately half of the foreign-born patients with MDR TB in 2004 were from Mexico, the Philippines, and Vietnam, the CDC said.

Also reported in the same MMWR issue, the first-ever data from the CDC and the WHO on rates of TB resistant to both firstand second-line antibiotics indicate that "extensively drug-resistant" TB accounted for 2% of all the MDR strains worldwide during the 2000-2004 period (MMWR 2006:55:301-5).