### Consequences of Alcohol on End-Organ Pathology in Liver Perspectives on a Proposed NIAAA-NIDA Merger

### Presentation to the SUAA Working Group of the SMRB September 23, 2009

#### Scott Friedman, MD

- Fishberg Professor of Medicine & Chief, Division of Liver Diseases,
  - Mount Sinai School of Medicine
  - President, American Assn for the Study of Liver Diseases
    - Member, NIAAA Advisory Council
      - Grantee, NIAAA & NIDDK

American Association for the Study of Liver Diseases



The Mount Sinai School of Medicine



## **Three Key Points**

- 1. Overall impact of alcoholic liver disease and its distinct features
- 2. Unique role of NIAAA in integrating alcohol research
- 3. Direct impact of an NIAAA-NIDA merger

### **Impact of Alcoholic Liver Disease**



- 2 million Americans suffer from alcoholic liver disease
- 30,000 deaths from alcoholic liver disease each year

Liver Transplant

# Impact of Alcoholic Liver Disease in the United States

- Alcohol use, together with hepatitis C (HCV) & hepatitis B (HBV) accounts for 70-90% of all cases of chronic liver disease in the Western world
- Up to 44% of patients with chronic HCV have a history of alcohol abuse
- Alcoholic liver disease accounts for ~ 20% of liver transplantations in the United States
- Alcohol abuse is an independent risk factor for liver cirrhosis and primary liver cancer (hepatocellular carcinoma)

## **Unique Features of Alcoholic Liver disease**

• The liver is the main site of alcohol metabolism



- Liver disease affects:
  - immunity
  - metabolism (protein, fat, and carbohydrate homeostasis)
  - bacterial clearance
  - drug detoxification

### Milestones in Alcoholic Liver Disease Leading to improved Health, supported by NIAAA

- Liver inflammation is precipitated by gut-derived pathogens due to altered intestinal permeability relevant to <u>ALL</u> forms of liver disease
- Alcoholic liver injury results from 'multiple hits'
- Studies of alcoholic liver injury uncovered new pathways and models relevant to fatty liver from obesity and other etiologies:
  - Role of oxidant stress and disordered fat metabolism
  - New models, e.g., zebrafish



Oil red O: lipid in liver

Work of K. Sadler-Edepli, PhD, NIAAA Grantee

#### The Multisystem and Inter-related Effects of Alcohol are Uniquely Integrated by NIAAA



# Proposed NIAAA-NIDA Merger: Concerns

- NIAAA has successfully and uniquely integrated the study of alcohol's effects on behavior, physiology, & pathology
- Integrated study of multi-organ damage has <u>not</u> been a focus of NIDA; this critical perspective may wither following a merger
- The impact of alcoholic liver injury and collateral effects on the health of Americans are significant and merit dedicated support
- It is uncertain what will be gained by an NIAAA-NIDA merger, but likely that much will be lost