Studies of Autoimmune/Neurological **Diseases in Communities Concerned** about **Environmental Exposures**

Environmental Public Health Tracking Conference March 24, 2004



Overview

- Community concerns
- ATSDR funded studies
 - 1. Multiple sclerosis prevalence studies
 - 2. Multiple sclerosis case-control study
 - 3. MS/ALS prevalence studies
- Future plans



Community Concerns

- Residents living near hazardous waste sites expressed concern about elevated rates of autoimmune/neurological diseases and possible association with environmental exposures

 Multiple sclerosis, amyotrophic lateral
 sclerosis, hugus, etc
 - sclerosis, lupus, etc.



- Local, state and federal public health agencies receive calls to address concerns and conduct cluster investigations
- No population-based registries
- Cluster investigations are not fruitful



Multiple Sclerosis

- Most common disabling neurological disease in young people
- Typical onset 20 40 years
- Approximately twice as common in women
- Inflammation, destruction, and scarring of the myelin
- Typically considered non-fatal



MS Prevalence in US

- Wide range of national estimates for MS cases in the United States
 - 123,000 (Baum '81)
 - 250,000 350,000 (Anderson, '92)
 - 217,000 (Noonan, '02)
- Limited data available on age-, sex-, race-, and region-specific prevalence



1. MS Prevalence Studies

 Cooperative agreements to establish methods for determining MS prevalence

Ohio (Lorain County)
Jackson County, Missour
Texas (Lubbock)





Initial MS Concerns

• Ohio

- -Reports of 25 people with MS (4200 pop)
- -Local iron foundry and plastics plant

• Missouri

-Refinery 1904-1982

• Texas

- Previous El Paso investigation
- Determine background MS prevalence



Case Ascertainment

- Primary data source: neurologist offices and clinics
- Secondary data sources
 - General practitioners
 - -Nursing homes
 - Patient advocacy groups
 - -Death certificates
 - -Hospital discharge data



ICD Codes Considered

- All records with the following ICD codes were reviewed:
 - multiple sclerosis (340)
 - transverse myelitis (323.9)
 - optic neuritis (377.3)
 - other demyelinating disease (341.8 341.9)



Data Collection

- Medical records abstracted to collect:
 - -Descriptive data
 - Diagnosis and physician history
 - Laboratory/clinical data
 - -Attack history



Case Definition

- Residence within study area
- Physician visit during 3-year period (1/1/98 - 12/21/00)
- MS diagnosis confirmed by consulting neurologist
 - Poser '83 criteria
 - McDonald '01 criteria



Preliminary Results

	19	Jackson	Lorain
	counties	County,	County,
	Texas	Missouri	Ohio
Population	424,916	120,799	280,000
Cases	182	109	458
Period Prevalence (per 100,000; 1998-2000)	43	90	164



Conclusions

- MS prevalence increases with distance from equator, which is consistent with data from other parts of the world
- Hypothesized reasons for gradient:
 - Ancestry/genetics
 - UV exposure



2. MS Case-Control Study

Multi-site

Lorain County, Ohio
Jackson County, Missouri
Lubbock, Texas

Cases identified from prevalence studies



- Controls selected from same clinic as cases (1:2 matching cases:controls)
- All study participants will be asked to complete a questionnaire and provide a blood sample
- Examine joint role of environmental exposures, infectious disease and genetic susceptibility



Questionnaire

- Demographics
- Complete residential history
- Medical history
- Occupational history
- Reproductive history



Blood Sample

- Infectious disease EBV
- Genetic susceptibility
 - Human leukocyte antigen (HLA)
 - T cell antigen receptor (TCR)
 - Myelin basic protein (MBP)
 - Tumor necrosis factor (TNF)
 - Apolipoprotein E (APOE)
 - Vitamin D receptor



Status

- Obtained funding and OMB approval
- Obtaining IRB approval (12 IRBs)
- Developing CATI for questionnaire
- Begin data collection 2nd quarter 2004 (tentative)



3. MS/ALS Prevalence Studies

- Expansion of initial prevalence studies to include ALS
 - S.E. Massachusetts
 - Illinois (five communities)
 - Washington (Hanford)
 - Missouri (Herculaneum)
 - Texas (El Paso, Lubbock and San Antonio)



State	Time Period	Criteria	Expected cases
IL	1/1/98-	McDonald	MS 60
	12/31/02	El Escorial	ALS 2
MA	1/1/98-	Poser/McD	MS 700-900
	12/31/02	El Escorial	ALS 70-80
MO	1/1/98-	Poser	MS 250
	12/31/02	El Escorial	ALS 34
WA	1/1/98-	McDonald	MS 2500
	12/31/03	El Escorial	ALS ?
TX	1/1/98-	Poser	MS 400
	12/31/03	El Escorial	ALS 150-220

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Future Plans

- Increase sample size of MS case-control study by including individuals from additional MS prevalence study areas
- Develop ALS case-control study
- Recommend surveillance of select autoimmune/neurological diseases



Contact Information

Dee Williamson Division of Health Studies, ATSDR

> 404-498-0586 djw8@cdc.gov

