

06/06/06 Overview, Approaches, & **Issues in the PGRN**

Rochelle M. Long, Ph.D. NIGMS



Research Sites





Background

"Networked" approach to pharmacogenetics research studies:

- Each group competed independently
- Each group had to be multi-disciplinary
- Groups could declare areas of interest; wanted breadth
- Group could have different research approaches; wanted range
- Many capitalizing on existing studies and trials
- Must make recommendations to represent <u>phenotypes</u>
- Must agree to work together to "raise field"
- Must be responsive to External Advisors ("more than sum...")



Data sharing was original to conception of PGRN:

- Must agree to share data in <u>www.PharmGKB.org</u>
- PharmGKB open to all scientists, upon registration
- Original identifying information remains at home site
- Information is organized by genes, drugs, diseases
- Hierarchy of phenotypes proposed: functional/cellular assays, pharmacokinetics, pharmacodynamics, outcomes
- <u>Pathways</u> are the bridge between genes and genome
- Shareable samples offered to Coriell repository



Range of studies, approaches underway:

- "Genotype-to-phenotype" studies
- "Phenotype-to-genotype" studies
- Diseases: CV, cancers, neuropsych., asthma
- Drug metabolism pathways (Phase I, Phase II, transport)
- Groups have discovered <u>common interests</u>, *e.g.*, DME, inflammation, adrenergic and serotonergic pathways
- Range of <u>scientific approaches</u>, including GWAS, some gene discovery in model organisms
- Sharing methods, technology, design, analysis (workshops)



Policies

Developed uniform policies through SC:

- Data deposits made at time of publication (when analyzed)
- <u>Model informed consent language</u> (being revised)
- Developed <u>recommendations on IP</u>, to not hinder research
- <u>Communications</u> with field (meetings, "white papers" coming on approaches, tests, ADRs, education areas)
- Positions on studies with *identified populations*
- Encourage <u>collaboration</u>, not competition



Plans

Future Plans:

- Promote development of PharmGKB, for data- and knowledge-sharing (interact with other dbs)
- Identify best, most appropriate technologies (Dec. mtg.)
- Encourage collaborations to maximize opportunities, *e.g.* with clinical trial designers, HMOs (May mtg.)
- Enhance translation of PG research (separate PA coming)
- **<u>Challenges</u>**: standardized vocabularies, lexicons, ontologies
- **<u>Challenges</u>**: HS protections, privacy, HIPAA
- **<u>Challenges</u>**: interactions with industry, and FDA