

Shanghai Health Study Overview

Goal: Conduct benzene health research in China.

Purpose: To gain an accurate understanding of the true relationship between benzene exposure and the risk of certain hematopoietic diseases. This study is an important element in the product stewardship programs of participating companies. It can also lead to improved benzene-driven, health-based regulatory (e.g., ambient air, water & occupational) standards worldwide, by providing the regulatory and scientific communities with information for more accurate assessment of the risks from exposure to benzene.

A unique set of circumstances exists at this point in time in Shanghai to conduct meaningful benzene research. These include a database of occupational exposures in a different regulatory environment, and willing and qualified co-investigators influential in local scientific medical, and regulatory establishments. Therefore, questions that have plagued benzene health risk assessment for decades appear to be answerable in Shanghai.

The Studies: The project involves three inter-related studies:

- Case-Control Study - A hospital-based case control study to investigate the potential relationship between benzene exposure and both Non-Hodgkins Lymphoma (NHL) and Acute Myelogenous Leukemia (AML), including an attempt to quantify the potency of benzene for the AML response.
- Disease Progression Study - A hospital/clinic-based investigation of the relationship between non-cancer diseases of the bone marrow and AML (cancer) to determine if effects on bone marrow progress through non-cancer stages to an ultimate leukemia. This study would also attempt to define the shape of the dose response for possible “precursor” diseases and could result in characterization of unique identifiers of benzene-induced AML.
- Molecular Epidemiology Study - Investigate if there is a quantitative relationship between benzene exposure and potential early indicators of both exposure and effect in a focused population of workers.

Investigators are leading experts in their field coming from such well-known institutions as the University of Colorado Health Sciences Center and Fudan University Medical Center. The combined studies apply cutting-edge technologies in new ways that will not only advance our understanding of benzene-induced hematopoietic disease, but may also identify new scientific approaches. [approaches to what? This seems an unfinished thought]

Benefits: Information gained from this project will have a direct relevance for worldwide regulatory standards designed to protect workers and the general population from benzene-induced hematopoietic disease. These include air and water quality standards, as well as hazardous waste and site remediation standards. These studies will assist contributing companies in achieving the highest product stewardship goals and in oversight of their risk management programs.

Shanghai Health Study Overview

Challenges: The proposed studies meet several technical, ethical, and cultural/political challenges.

- **Technical** - In order to stand up to scientific scrutiny, studies must be large, well-managed, and incorporate the latest technology. State-of-the-art research is absolutely necessary to make real progress in developing sound benzene risk assessments. To that end, a Scientific Review Panel made up of leading experts from a number of fields relevant to these studies has reviewed and endorsed the research protocols.
- **Ethical** - To address the ethical issues, the study protocols have all undergone a thorough analysis to ensure that they meet international standards for research involving humans, including that both the subjects and participating communities benefit from having participated in the study; that confidentiality is preserved, and that researchers are working with local regulatory authorities. An Ethics Panel is part of the oversight for the project and includes a Chinese bioethicist.
- **Cultural/Political** - A key to this project's success is the partnerships with local medical and regulatory entities such as the Medical School at Fudan University, the Institute for Public Health Supervision, and the Shanghai Municipal Centers for Disease Control and Prevention (The latter two are responsible for occupational exposure monitoring and enforcement).

Costs: The estimated costs for the project were derived after careful consideration of all of the technical, ethical, and political issues. (inflation has been incorporated into the estimate).

Item	Cost Estimate
Case-Control Study	\$2,283,100
Disease Progression & Molecular Epid. Studies	\$16,702,000
Total Research Costs	\$18,985,100
Scientific Review Panel	\$648,000
Ethics Review Panel	\$216,000
External Pathology & Cytogenetics Workgroups	\$130,000
Total Panel Costs	\$994,000
Outside Counsel	\$125,000
Public Affairs	\$1,000,000
QA/QC Activities	\$225,000
Miscellaneous Costs	\$1,350,000
Subtotal	\$21,329,100
Administrative Overhead (5%)	\$1,066,500
GRAND TOTAL	\$22,395,600

Funding: Funding for the project will be handled by a consortium of interested stakeholders, with shares determined by size of the benzene related operations of the contributing organization. The proposed funding formula was devised based on the assumption that most of the support will come from the petroleum and petrochemical

Shanghai Health Study Overview

industries. It is anticipated to take five years to complete the project, and that the costs will be highest in the first years of the project.