

**Department of  
Pharmacology and Toxicology  
University of Louisville  
School of Medicine**

**1998**

**Annual Report**

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## **I. Executive Summary**

1998 was a year of continued transition and program development for the Department of Pharmacology and Toxicology. The Department was deeply saddened by the unexpected death of Dr. Kee Chang Huang in November. Dr. Huang served on the Department faculty since 1953, and continued to be an active member of the Department as Professor Emeritus. A resolution in his honor was written and approved by the Department, the Medical Council, and the Faculty Senate. A copy is provided in the appendix. The Department initiated the Kee Chang Huang Outstanding Graduate Student Award and the Kee Chang Huang Memorial Lecture. The 1998 Kee Chang Huang Outstanding Graduate Student Award was awarded to Jian Cai. A second major event was the retirement of Dr. William J. Waddell as Professor on July 1. Dr. Waddell was appointed Professor Emeritus (he was appointed Chairman emeritus upon retirement as chair in 1997) and was honored with a reception and the initiation of a William J. Waddell seminar program. A third major event was the appointment of Joel A. Kaplan, M.D. as Vice President for Health Affairs/Dean for the School of Medicine/Senior Vice Provost for Academic Affairs effective July 1. Dr. Kaplan was previously Senior Vice-President for Clinical Affairs and Horace W. Goldsmith Professor and Chairman of the Department of Anesthesiology at Mount Sinai School of Medicine. Dr. Kaplan brought impressive credentials in research and an aggressive agenda for enhancement of research productivity in the School of Medicine and the Health Science Center.

The primary focus of the year was the recruitment of new faculty. A search committee consisting of Dr. David Hein (Chair), Dr. Fred Benz, Dr. Theresa Chen, Dr. Fred Hendler, Dr. James Kang, and Dr. Peter Rowell screened applications for two tenure track faculty positions. Over 100 applications were received and several candidates were invited for interviews. Three of the top candidates, Dr. Carrie Haskell-Leuvano, (Postdoctoral Fellow, Oregon Health Sciences University), Dr. Paul MacDonald, (Assistant Professor, Saint Louis University School of Medicine) and Dr. Donghai Wu (Assistant Professor, University of Kentucky College of Medicine) were offered a tenure track assistant or associate professor (Dr. MacDonald) position. Each declined, however, in order to accept new faculty positions at the University of Florida, Case-Western Reserve University, and the University of Florida, respectively. In addition to starting salary and lack of institutional research core facilities, a major stumbling block was the requirement for the new faculty to be placed in temporary research and office space prior to completion of the Baxter Biomedical Research Facility in 1999. Rather than continue on down the list of applicants, the Department initiated a new search in 1999 to coincide with the completion of the Baxter Biomedical Research Facility. The Department approved the joint appointment of Janice Sullivan, M.D. as assistant professor. Dr. Sullivan's primary appointment is in the Department of Pediatrics, and, prior to 1998, she had an associate faculty appointment in the Department of Pharmacology and Toxicology. Drs. Scott Gruber and Paul Epstein were appointed as adjunct faculty.

Highlights in program development included revision of the guidelines and curriculum for the graduate program in pharmacology and toxicology, which was approved by unanimous vote of the faculty. In addition, a Strategic Plan focusing on research and training strategies was initiated and approved by unanimous vote of the faculty. The research productivity of the Department increased significantly in 1998, and the Department contributed to the University of Louisville's transition to a premier metropolitan research University.

## **II. Mission Statement**

The Department of Pharmacology and Toxicology is committed to academic excellence and to the attainment of regional, national, and international recognition for the quality of its educational, research, and service activities. Guided by the University of Louisville Challenge for Excellence to become a preeminent metropolitan research university, the Department Strategic Plan will focus on accomplishment of five broad objectives:

- Provide instruction in pharmacology and toxicology of the highest quality for the education and preparation of medical, dental, nursing, and other health care professional students. Emphasis will be placed on the fundamental principles necessary for life-long learning and the essential knowledge required for rational, effective, and safe use of drug therapy.
- Advance biomedical knowledge through active contribution to high quality research and other scholarly activities, particularly in pharmacology and toxicology and other areas of focus within the University of Louisville Challenge for Excellence.
- Provide high quality research and educational experiences in pharmacology and toxicology for the education and training of future biomedical scientists who can provide and advance biomedical education, research, and service.
- Provide instruction of the highest quality in pharmacology and toxicology that is appropriate for students at the undergraduate, graduate, and postgraduate levels.
- Provide high quality service to the School of Medicine, the Health Sciences Center, the University, the people of Louisville and the surrounding region, the Commonwealth of Kentucky, professional organizations, the nation, and the world.

## **III. Personnel Listing**

### **A. Faculty with Primary Appointments**

**Benz, Frederick W.**, Professor; Ph.D., Pharmacology, University of Iowa (1970).

**Carr, Laurence A.**, Professor; Ph.D., Pharmacology, Michigan State University (1969).

**Chen, Theresa S.**, Professor; Ph.D., Pharmacology, University of Louisville (1971).

**Hein, David W.**, Peter K. Knoefel Professor and Chair; Ph.D., Pharmacology, University of Michigan (1982).

**Hurst, Harrell E.**, Professor; Ph.D., Toxicology, University of Kentucky (1978).

**Myers, Steven R.**, Associate Professor; Ph.D., Pharmacology, University of Kentucky (1986).

**Nerland, Donald E.**, Professor; Ph.D., Medicinal Chemistry, University of Kansas (1974).

**Pierce, William M., Jr.**, Professor; Ph.D., Pharmacology and Toxicology, University of Louisville (1981).

**Rowell, Peter P.**, Professor; Ph.D., Pharmacology and Therapeutics, University of Florida (1975).

**Waite, Leonard C.**, Professor and Vice Chairman; Ph.D., Pharmacology, University of Missouri (1969).

**Williams, Walter M.**, Professor; Ph.D., Pharmacology, University of Louisville (1970); M.D., University of Louisville (1974).

## **B. Faculty with Joint Appointments**

**Aronoff, George R.**, Professor of Medicine, and Pharmacology and Toxicology; M.D., Indiana University (1975).

**Delamere, Nicholas A.**, Professor of Ophthalmology and Visual Sciences, and Pharmacology and Toxicology; Ph.D., Membrane Physiology and Biophysics, University of East Anglia, Norwich, England (1976).

**Kang, Y. James**, Associate Professor of Medicine, and Pharmacology and Toxicology; Ph.D., Cell Biology and Zoology, Iowa State University (1989).

**Miller, Frederick N.**, Professor of Physiology and Biophysics, and Pharmacology and Toxicology; Ph.D., Pharmacology, University of Cincinnati (1971).

**Rodgers, George C., Jr.**, Professor of Pediatrics, and Pharmacology and Toxicology; Ph.D., Organic Chemistry, Yale University (1964); M.D., State University of New York (1975).

**Sullivan, Janice E.**, Assistant Professor of Pediatrics, and Pharmacology and Toxicology; M.D., University of Minnesota (1988).

**Tamburro, Carlo H.**, Professor of Medicine, and Pharmacology and Toxicology, and Chief, Division of Occupational Toxicology; M.D., Seton Hall (1962); M.P.H., Columbia University (1985).

**Wong, John L.**, Professor of Chemistry, and Pharmacology and Toxicology; Ph.D., Chemistry, University of California at Berkeley (1966).

**Zimmerman, Thom J.**, Professor and Chairman of Ophthalmology and Visual Sciences, and Pharmacology and Toxicology; Ph.D., Pharmacology, University of Florida (1976); M.D., University of Illinois (1968).

### **C. Faculty with Associate Appointments**

**Brier, Michael E.**, Associate Professor of Medicine; Ph.D., Industrial and Physical Pharmacy, Purdue University (1986).

**Edmonds, Harvey L.**, Professor of Anesthesiology; Ph.D., Pharmacology, University of California at Davis (1974).

**Jumblatt, James E.**, Professor of Ophthalmology and Visual Sciences; Ph.D., Biological Sciences, Columbia University (1975).

**Lang, Calvin A.**, Professor of Biochemistry and Molecular Biology; Sc.D., Biochemistry and Nutrition, Johns Hopkins University (1954)

**Rigor, Benjamin**, Professor and Chairman of Anesthesiology; M.D., University of the East Ramon Magsaysay Memorial Medical Center (1962).

**Ross, Mitchell P.**, Assistant Professor of Pediatrics; M.D., Oregon Health Sciences University (1985).

**Schurr, Avital**, Professor of Anesthesiology; Ph.D., Biochemical Pharmacology, Ben Gurion University, Beer Sheva, Israel (1977).

### **D. Faculty with Emeritus Appointments**

**Dagirmanjian, Rose**, Professor Emerita; Ph.D., University of Rochester (1960).

**Huang, Kee Chang**, Professor Emeritus; M.D. Columbia University (1953).

**Jarboe, Charles H.**, Professor Emeritus; Ph.D., University of Louisville (1956).

**Knoefel, Peter K.**, Emeritus Professor and Chairman; M.D., Harvard University (1931).

**Scharff, Thomas G.**, Emeritus Professor; Ph.D., University of Rochester (1956).

**Waddell, William J.**, Professor and Chair Emeritus; M.D., University of North Carolina (1955).

### **E. Faculty with Adjunct Appointments**

**Epstein, Paul N.**, Adjunct Associate Professor of Pharmacology and Toxicology; Ph.D., Baylor College of Medicine (1981).

**Friedman, Marvin A.**, Adjunct Professor of Pharmacology and Toxicology; Ph.D., Massachusetts Institute of Technology (1967).

**Gruber, Scott A.**, Adjunct Associate Professor of Pharmacology and Toxicology; M.D., SUNY Downstate Medical School (1983); Ph.D., University of Minnesota (1991).

**Hayes, A. Wallace**, Adjunct Professor of Pharmacology and Toxicology; Ph.D., Auburn University (1967).

**Holthouser, Michael G.**, Adjunct Assistant Professor of Pharmacology and Toxicology; M.D., University of Kentucky (1971).

**Lewis, Richard**, Adjunct Assistant Professor of Pharmacology and Toxicology; M.D., Case Western Reserve University (1980).

**Matyunas, Nancy**, Adjunct Instructor of Pharmacology and Toxicology; Pharm.D., University of Utah (1983).

**Nicholson, John A.**, Adjunct Assistant Professor of Pharmacology and Toxicology; D.M.D., University of Louisville (1979); Ph.D., University of Louisville (1968).

**Woodcock, Thomas M.**, Adjunct Professor of Pharmacology and Toxicology; M.D., Columbia University (1970).

## **F. Faculty with Emeritus Adjunct Appointments**

**Csaky, Tihamer Z.**, Emeritus Adjunct Professor of Pharmacology and Toxicology; M.D., University of Budapest (1939).

**Darby, Thomas D.**, Emeritus Adjunct Professor of Pharmacology and Toxicology; Ph.D., Medical College of South Carolina (1957).

**Gordon, Helmut A.**, Emeritus Adjunct Professor of Pharmacology and Toxicology; M.D., University of Budapest (1932).

## **G. Staff**

**Carpenter, Sharon**; Executive Secretary

**Clark, Denny**, Lab Tech III; Therapeutics and Toxicology Laboratory

**Corbett, Donna**; Lab Tech III

**Doll, Mark;** Research Associate

**Duggan, Deborah;** Lab Tech III

**Emery, Nicole;** Lab Tech III

**Jamison, Andrea;** Lab Tech II

**Metaxas, John;** Coordinator, Therapeutics and Toxicology Laboratory

**Greca, Edie;** Business Manager III

**Rubin-Teitel, Heddy;** Program Assistant III

**Smith, Ned;** Senior Research Technologist

**Srinivasan, Chandrika, Ph.D.,** Student Assistant

**VonderHaar, John,** Laboratory Research Technician III

**Wenger, Aaron;** Student Assistant, Therapeutics and Toxicology Laboratory

## **H. Graduate Students**

### **Doctoral Program**

**Alimozhin, Talgat**

**Brewer, Bradley G.**

**Cai, Jian**

**Fretland, Adrian**

**Fu, Xin**

**Hartford, April**

**Hennion, John**

**Jacobs, Al (Chip)**

**Lambert, Jason**

**Leff, Matthew**

**Li, Junyu**

**Nangju, Norma**

**Neale, Jason**

**Payne, Scot**

**Spinnato, Joe**

**Sun, Xichun**

### **Masters Program**

**Hasselbacher, David**

**Melo, Nicolas**

## **I. Postdoctoral Fellows**

**Devanaboyina, Udaya-sankar**

**Meng, Xianweng**

**Wang, Guang-wu**

**Wang, Ju-Feng**

**Wu, Huiyun**

**Webb, Stephanie**

**Xiao, Gong H.**

**Zhou, Zhan-Xiang**

## **J. Division of Occupational Toxicology**

**Buchanan, Stephanie**, B.S.N.; Clinical Research Coordinator

**Canchis, Willie**, M.D.; Research Fellow

**Cho, Lori**; Student Assistant

**Crawford, Robert**; Student Assistant

**Deeb, Lililane**, M.D.; Research Fellow

**Fortwengler, Philip**; Manager, Medical Surveillance

**Hall, Carl**, Ph.D.; Project Coordinator

**Hiser, Melissa**; Clerical Specialist III

**Jarman, Susanna**; Student Assistant

**Kapoor, Sandeep**, M.D.; Research Fellow

**Matthews, Sherri**; Executive Secretary

**Miller, Barbara L.**, B.S.N., M.S.; Senior Clinical Research Coordinator

**Morgenson, Melissa**; Secretary

**O'Connell, Kathleen**, Ph.D.; Research Facilitator

**Quiggins, Patricia**, Ph.D.; Research Associate

**Redferin, Lauren**; Student Assistant

**Reynolds, Eva**; Receptionist

**Reynolds, Lark**; Program Assistant III

**Richardson, Rita**; Secretary II

**Sherrill, Mike;** Student Assistant

**Sterry, Rachel;** Student Assistant

**Tamburro, Carlo H.,** M.D, M.P.H.; Division Chief

### **K. Faculty Appointments**

**Epstein, Paul N.,** Ph.D., Adjunct Associate Professor, November 1, 1998

**Gruber, Scott A.,** M.D., Ph.D., Adjunct Associate Professor, October 1, 1998

**Sullivan, Janice E.,** M.D., Assistant Professor (joint appointment), December 1, 1998

**Waddell, William J.,** M.D., Professor Emeritus, July 1, 1998

### **L. Staff Promotions**

**Carpenter, Sharon,** Promotion to Executive Secretary, effective August, 1998

**Greca, Edie,** Promotion to Business Manager III, effective August, 1998

### **M. New Graduate Student Appointments**

**Hartford, April**

**Lambert, Jason**

**Nangju, Norma**

**Neale, Jason**

**Sun, Xichun**

### **N. Graduations**

**David Hasselbacher** (M.S. awarded May 1998). Thesis title “The Development of a Dosing Model for the Anticoagulant Heparin Using Neural Networks.” Advisor: George R. Aronoff.

#### IV. Publications

1. Byrd, D.M., Allen, D.O., Beamer, R.L., Besch, H.R., Jr., Bylund, D.B., Doull, J., Fleming, W.W., Fries, A., Guengerich, F.P., Hornbrook, R., Lasagna, L., Lum, B.K., Michaelis, E.K., Morgan, E.T., Poland, A., Rozman, K.K., Smith, J.B., Swanson, H.I., Waddell, W.J., and Wilson, J.D. (1998). Letter to the Editor: The dose-response model for dioxin. *Risk Analysis*, **18**:1-2.
2. Chen, D., Waite, L.C., Pierce, W.M. Jr. (1998). In vitro bone resorption is dependent on physiological concentrations of zinc. *Biological Trace Element Research* **61**:9-18.
3. Deitz, A.C., Fretland, A.J., Leff, M.A., and Hein, D.W. (1998). Homo sapiens N-acetyltransferase-1 (NAT1) gene, NAT1\*26A allele, complete cds. *AF071552*.
4. Deitz, A.C., Fretland, A.J., Leff, M.A., Doll, M.A., and Hein, D.W. (1998). Homo sapiens N-acetyltransferase-1 (NAT1) gene, NAT1\*26B allele, complete cds. *AF067408*.
5. Delamere, N.A., Pierce, W.M., and Hou, Y. (1998). The influence of ascorbic acid on active sodium transport in cultured rabbit nonpigmented ciliary epithelium. *Investigative Ophthalmology and Visual Science* **39**(1):143-50.
6. Feng, Y., Rustan, T.D., Fretland, A.J., Cooley, A.M., Lee, K.J., Becker, W.K., and Hein, D.W. (1998). A rapid and sensitive method for detection of urinary bladder tumors in rodents. *Proceedings of the Western Pharmacology Society* **41**:11-13.
7. Fretland, A.J., Leff, M.A., Doll, M.A., and Hein, D.W. (1998). Homo sapiens N-acetyltransferase (NAT2) gene, NAT2\*5D allele, complete cds. *AF042740*.
8. Huang, K.C. *The Pharmacology of Chinese Herbs*, edition 2. (1998). Boca Raton: CRC Press.
9. Kang, Y. J. and Wang, J.-F. (1998). Cardiac protection by metallothionein against ischemia-reperfusion injury and its possible relation to ischemic preconditioning, in *Metallothionein IV* (C. Klaassen, ed), pp511-516, Birkhauser Verlag Basel/Switzerland.
10. Kemper, R.A., Elfarra, A.A., and Myers S.R. (1998). Metabolism of 3-butene-1,2-diol in B6C3F1 mice. Evidence for involvement of alcohol dehydrogenase and cytochrome p450. *Drug Metabolism and Disposition* **26**(9):914-20.
11. Leff, M.A., Doll, M.A., Feng, Y., Fretland, A.J., and Hein, D.W. (1998). Homo sapiens N-acetyltransferase (NAT2) gene, NAT2\*6D, complete cds. *AF055875*.
12. Leff, M.A., Doll, M.A., Fretland, A.J., and Hein, D.W. (1998). Homo sapiens N-acetyltransferase (NAT2) gene, NAT2\*14G, complete cds. *AF055874*.

13. Rowell, P.P. and Duggan, D.S. (1998). Long-lasting inactivation of nicotinic receptor function in vitro by treatment with high concentrations of nicotine. *Neuropharmacology* **37**:103-111.
14. Tsueda, K., Mosca, P.J., Heine, M.F., Loyd, G.E., Durkis, D.A.E., Malkani, A.L., and Hurst, H.E. (1998). Mood during continuous epidural infusion of morphine or fentanyl. *Anesthesiology* **88(4)**: 885-891.
15. Uthus, E. O. and Kang, Y. J. (1998). Effect of buthionine sulfoximine on the response to arsenic deprivation in female rats. *Journal of Trace Element and Experimental Medicine* **11**:29-36.
16. Waddell, W.J. "Evidence against environmental endocrines influencing cancer of the breast, uterus, prostate, and testicle" (1998). In: G.E. Dunaif, S.S. Olin, J.A. Scimeca, and J.A. Thomas (Eds.), *Human Diet and Endocrine Modulation: Estrogenic and Androgenic Effects* (Edition 1); Washington: ILSI Press, pp. 267-276.
17. Waddell, W.J. (1998). Epidemiological studies and effects of environmental estrogens. *International Journal of Toxicology*, **18**:197-215.
18. Wang, G.-W., Schuschke, D. A. and Kang, Y. J. (1998). Metallothionein overexpressing neonatal mouse cardiomyocytes are resistant to hydrogen peroxide toxicity. *American Journal of Physiology* **276**:H167-H175.
19. Wang, G.-W., Wu, H.-Y. and Kang, Y. J. (1998). Metallothionein and its importance relative to glutathione in cardiac protection against doxorubicin toxicity, in *Metallothionein IV* (C. Klaassen, ed), pp517-522, Birkhauser Verlag Basel/Switzerland.
20. Wong, J.L., Zheng, Y.T., Li, J., Tamburro, C.H., and Benz, F.W. (1998). Immunoassay of haemoglobin-acrylonitrile adduct in rat as a biomarker of exposure. *Biomarkers* **3**:317-326.
21. Wrightson, W.R., Myers, S.R., and Galandiuk, S. (1998). HPLC analysis of mitomycin C in tissue and serum with ultraviolet detection. *Biochemical and Biophysical Research Communication* **706(2)**:358-61.
22. Wu, H.-Y. and Kang, Y. J. (1998). Inhibition of buthionine sulfoximine-enhanced doxorubicin toxicity in metallothionein overexpressing transgenic mouse heart. *Journal of Pharmacology and Experimental Therapeutics* **287**:515-520.
23. Wu Q, Pierce, W.M., Jr., and Delamere, N.A. (1998). Cytoplasmic pH responses to carbonic anhydrase inhibitors in cultured rabbit nonpigmented ciliary epithelium. *Journal of Membrane Biology* **162**:31-38.
24. Yin, X., Wu, H.-Y., Chen, Y. and Kang, Y. J. (1998). Induction of antioxidants by adriamycin in mouse heart. *Biochemical Pharmacology* **56**:87-93.

## **A. Additional Publications of Faculty with Joint Appointments.**

1. Aronoff, G.R. (1998). Informatics and end-stage renal disease. *Advances in Renal Replacement Therapy* **5**:292-298.
2. Aronoff, G.R., Erbeck, K.M., and Brier, M.E. (1998). "Prescribing drugs for dialysis patients." In: W. L. Henrich (Ed.), *Principles and Practice of Dialysis*. Williams and Wilkins.
3. Ward, R.A., and Aronoff, G.R. (1998). "Anticoagulation strategies during hemodialysis procedures." In: W. L. Henrich (Ed.), *Principles and Practice of Dialysis*. (Edition 2); Baltimore: Williams & Wilkins, pp. 70-78).
4. Delamere, N.A., Manning, R.E., Jr., Liu, L., Moseley, A.E., and Dean, W.L. (1998). Na, K-ATPase polypeptide upregulation responses in lens epithelium following an episode of cytoplasmic ATP depletion. *Investigative Ophthalmology and Visual Science* **39(5)**:763-768.
5. Kuwahara, S., Chin, S., and Delamere, N.A. (1998). Partial inhibition of Na, K-ATPase activity in cultured rabbit nonpigmented ciliary epithelium following an episode of cytoplasmic ATP depletion. *Acta Physiologica Scandinavica* **164**:13-20.
6. Ehringer, W.D., Edwards, M.J., Wintergerst, K.A., Cox, A., and Miller, F.N. (1998). An increase in endothelial intracellular calcium and F-actin precedes the extravasation of interleukin-2 activated lymphocytes. *Microcirculation* **5**:71-80.
7. Heinzelmann, M., Mercer-Jones, M.A., Flodgaard, H., and Miller, F.N. (1998). Heparin-binding protein (CAP37) is internalized in monocytes and increases LPS-induced monocyte activation. *Journal of Immunology* **160**:5530-5536.
8. Heinzelmann, M., Platz, A., Flodgaard, H., and Miller, F.N. (1998). Heparin binding protein (CAP37) is an opsonin for staphylococcus aureus and increases phagocytosis in monocytes. *Inflammation* **22**:493-507.
9. Heinzelmann, M., Polk, H.C., Jr., and Miller, F.N. (1998). Modulation of lipopolysaccharide-induced monocyte activation by heparin-binding protein and fucoidan. *Infection and Immunity* **66(12)**:5842-5847.
10. Lentsch, A.B., Yoshidome, H., Cheadle, W.G., Miller, F.N., and Edwards, M.J. (1998). Chemokine involvement in hepatic ischemia/reperfusion injury in mice: roles for macrophage inflammatory protein-2 and KC [corrected and republished article originally printed in *Hepatology* 1998 Feb;27(2):507-12]. *Hepatology* **27(4)**:1172-1177.

11. Lentsch, A.B., Miller, F.N., and Edwards, M.J. (1998). Mechanisms of leukocyte-mediated tissue injury induced by interleukin-2. *Cancer Immunology and Immunotherapy* **47**:243-248.
12. Lentsch, A.B., Edwards, M.J., Sims, D.E., and Miller, F.N. (1998). N-nitro-L-arginine methyl ester inhibits inflammatory liver injury induced by interleukin-2. *Journal of Leukocyte Biology* **63**:22-30.
13. Yoshidome, H., Lentsch, A.B., Cheadle, W.G., Miller, F.N., and Edwards, M.J. (1998). Enhanced pulmonary expression of CXC chemokines during hepatic ischemia/reperfusion-induced lung injury in mice. *Journal of Surgical Research* **81**:33-37.
14. Rodgers, G.C., and Matyunas, N.J. (1998). Hazardous materials guide for first responders. Rodgers, G. C. and Matyunas, N. J. (Ed.), *Federal Emergency Management Agency: United States Fire Administration*.
15. Sullivan, J.E., and Rees, A. (1998). Recurrent bacterial endocarditis in a patient with repaired congenital heart disease. *Intensive Thoughts* 1 Edition; 5-5. Louisville: Kosair Children's Hospital/University of Louisville.
16. Galbreath, K.C., Zygarlicke, C.J., Huggins, F.E., Huffman, G.P., and Wong, J.L. (1998). Chemical Speciation of Nickel in Residual Oil Ash. *Energy and Fuels* **12**:818-822.
17. Liu, A., and Wong, J.L. (1998). Nickel Speciation of Fly Ash by Combining Phase Separation with Carbon Paste Electrode Direct Analysis. Preprints: *American Chemical Society, Division of Environmental Chemical Symposia* **38(2)**:197-199.

## **V. Abstracts**

1. Benz, F.W., Cai, J., Nerland, D.E., Li, J., Corbett, D., and Hurst, H.E. The nonlinear dose-dependent toxicodynamics of acrylonitrile can be explained by the similar dose-dependence of its toxicokinetics. *Toxicological Sciences* 42: No. 1-Supplement, 212, 1998.
2. Cai, J. and Hurst, H.E. Use of GC/MS to investigate mechanisms of N-(carboxymethyl)-valine adduct formation in hemoglobin. 46<sup>th</sup> ASMS Conference on Mass Spectrometry and Allied Topics, Orlando, FL, May 31 – June 4, 1998. Proceedings 46<sup>th</sup> ASMS Conference on Mass Spectrometry and Allied Topics, p.1138 (1998).
3. Cai, J. and Hurst, H.E. Use of GC/MS to investigate mechanisms of N-(carboxymethyl)-valine adduct formation in hemoglobin. UofL Health Sciences Center 24<sup>th</sup> Annual Student Research Day, October 27, 1998.
4. Cai, J. and Hurst, H.E. Use of GC/MS to investigate mechanisms of N-(carboxymethyl)-valine adduct formation in hemoglobin. Ohio Valley Society of Toxicology Regional Meeting, Dayton, OH, October 30, 1998.

5. Feng, Y., Finley, J.W., Davis, C.D., Fretland, A.J., Korynta, E., and Hein, D.W.: The protective effect of selenium on the aberrant crypt formation in rats administered 3,2'-dimethyl-4-aminobiphenyl. *Proceedings of the American Association for Cancer Research* 39:390, 1998.
6. Feng, Y., Finley, J.W., Davis, C.D., Fretland, A.J., Korynta, E., Hein, D.W., and Becker, W.K.: Inhibition of 3,2'-dimethyl-4-aminobiphenyl-induced aberrant crypt formation in the colon of rats by selenium. *Proceedings of the Eighteenth Annual Frank Low Research Day*, p. 53, 1998.
7. Feng, Y., Fretland, A.J., Rustan, T.D., Becker, A.M., Becker, W.K., and Hein, D.W.: Role of CYP1A2 and NAT2 polymorphism in colon carcinogenesis of Syrian hamsters administered 3,2'-dimethyl-4-aminobiphenyl. *Proceedings of the Seventh International Conference of Carcinogenic/Mutagenic N-Substituted Aryl Compounds*, Nagoya, Japan, p. 32, 1998.
8. Feng, Y., Fretland, A.J., Rustan, T.D., Becker, A.M., Becker, W.K., and Hein, D.W.: Formation of colonic aberrant crypt foci by 3,2'-dimethyl-4-aminobiphenyl in rapid and slow acetylator congenic hamsters requires metabolic activation by CYP1A2 and NAT2. *Proceedings of Research Louisville 1998*, November, 1998.
9. Feng, Y., Rustan, T.D., Doll, M.A., Leff, M.A., Cooley, A.M., Fretland, A.J., Grazziotin, M.U., Luthra, G., Becker, W.K., and Hein, D.W.: Human N-acetyltransferase-1 (NAT1) and -2 (NAT2) genotype/phenotype determinations in cytosolic preparations from surgical human colon specimens. *Proceedings of the First International Workshop on the Arylamine N-Acetyltransferases*, Kuranda, Queensland, Australia, p. 23, 1998.
10. Fretland, A.J., Feng, Y., and Hein, D.W.: 2-Amino-1-methyl-6-phenylimidazo[4,5-b]pyridine (PhIP) DNA adduct formation in homozygous rapid and slow acetylator congenic hamsters. *Toxicological Sciences* 42:Number 1-S:281, 1998.
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40. Spatola, A.F., Romanovskis, P.J., and Pierce, W.M., Jr. Cyclic Pseudopeptides. Presented at International Peptide Symposium, Vienna, Austria, Fall, 1998.
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## VI. Invited Scientific Presentations and Seminars

1. Benz, Frederick W. Presentation to Chemistry Class at Jefferson Community College, at the invitation of Mr. Steve Pruett, Chemistry Instructor, 4/20/98.
2. Hein, David W. Molecular Pharmacogenetics of Acetylation: Its Possible Role in Genetic Predisposition to Cancer from Aromatic and Heterocyclic Amine Carcinogens. Department of Pharmacology and Graduate Center for Toxicology, University of Kentucky, Lexington, Kentucky, February 1998.
3. Hein, David W. Role of N-Acetyltransferase Polymorphisms in Genetic Predisposition to Cancer. Symposium on the Role of Genetic Polymorphisms and Repair Deficiencies in Environmental Disease, 37<sup>th</sup> Annual Meeting of the Society of Toxicology, Seattle, Washington, March 1998.
4. Hein, David W. Metabolic Activation of N-Hydroxy-2-Amino-1-Methyl-6-Phenylimidazo-[4,5-b]Pyridine (N-OH-PhIP) by Twenty-Three Recombinant Human NAT2 Allozymes. Genetics of Cancer and Exposure Susceptibility Mini-symposium, Eighty-Ninth Annual Meeting of the American Association for Cancer Research, New Orleans, Louisiana, March 1998.
5. Hein, David W. Structure/Function Studies on Metabolic Activation of Aromatic Amine Carcinogens by Recombinant Human N-Acetyltransferases. Student Research Opportunity Program, University of Louisville, Louisville, Kentucky, July 1998.
6. Hein, David W. Human N-Acetyltransferase-1 (NAT1) and -2 (NAT2) Genotype/Phenotype Determinations in Cytosolic Preparations from Surgical Human Colon Specimens. First International Workshop on the Arylamine N-Acetyltransferases, Kuranda, Queensland, Australia, October 1998.
7. Kang, Y.J. London Regional Cancer Center, Univ. Western Ontario, London, Canada, "Metallothionein and adriamycin cardiotoxicity," Nov. 17, 1998
8. Kang, Y.J. Depts. Pathology, and Pharmacology and Toxicology, Univ. Western Ontario, London, Canada, "Metallothionein in myocardial ischemic preconditioning," Nov. 16, 1998.
9. Kang, Y.J. Institute of Toxicology, National Taiwan University, Taiwan, "Transgenic approach to antioxidant function of catalase in the heart," Apr. 24, 1998.
10. Kang, Y.J. Drug Development Division, Development Center for Biotechnology, Taiwan, "Catalase provides protection against oxidative cardiotoxicity," Apr. 24, 1998
11. Kang, Y.J. Institute of Biomedical Sciences, Academia Sinica, Taiwan, "Catalase in cardiac protection against oxidative injury," Apr. 23, 1998.

12. Kang, Y.J. Department of Cardiology, Veterans General Hospital in Taipei, Taiwan, "Metallothionein provides cardiac protection against ischemia-reperfusion injury," Apr. 23, 1998.
13. Kang, Y.J. Department of Cardiology, Veterans General Hospital in Taichung, Taiwan, "Cardiac catalase and oxidative injury," Apr. 20, 1998.
14. Kang, Y.J. The Keynote Speaker of the Toxicology Society of Taiwan, The Thirteenth Joint Annual Conference of Biomedical Sciences, National Taiwan University, Taipei, "Transgenic approaches to cardiac protection against oxidative injury," Apr. 19, 1998.
15. Kang, Y.J. Department of Human Nutrition and Food Management, Ohio State University, Columbus, OH. "Cardiac specific overexpression of metallothionein protects against oxidative injury," Mar. 11, 1998.
16. Myers, S.R. Biomarkers of Nitroaromatic Munitions. National Exposure Research Laboratory, Cincinnati, Ohio, April, 1998.
17. Pierce, William M. Jr. Bone Targeted Estrogens. Presentation at Kowa Research Institute, San Jose, CA, November 9, 1998.
18. Rowell, Peter P. Annual Nicotine Research Conference, Duke University, Nov. 4, 1998.

## VII. Research Grants and Contracts

### A. Research Grant and Contract Proposals Submitted

	<u>Agency</u>	<u>Budget Requested</u>
<b>Dr. Frederick W. Benz</b>		
Vinyl Monomers: Detoxified by Glutathione S-Transferases (Co-I); D.E. Nerland (PI) 02/01/00 – 01/31/04	NIH	\$510,287
<b>Dr. Laurence A. Carr</b>		
Tobacco Use and Attention Deficit Disorder (Co-I); C. Mitchell (PI) 10/01/98 – 09/30/02)	NIH	\$1,717,523
<b>Dr. David W. Hein</b>		
Pharmacogenetics of Drug and Carcinogen Metabolism (cont.) (P.I.) 07/01/98 – 06/30/99	NIH-NCI	\$361,947
Molecular Epidemiology of Colon and Stomach Cancers (PI) 09/01/98 – 06/30/00	NIH-NCI	\$177,550
Center for Children's Environmental Health (Co-I); S.R. Myers (PI) 08/01/98 – 06/31/02	NIH/NIEHS	\$3,368,844
Pharmacogenetics of Fetal Tobacco Smoke Exposure (Co-I); S.R. Myers (PI) 12/01/98 – 11/30/03	NIH	\$1,840,554
Role of Genetic Polymorphism In Liver Toxicity (Co-I); T.S. Chen (PI) 09/01/98 – 08/31/99	UofL (CGeMM)	\$24,325
Pharmacogenetics of Fetal Tobacco Smoke Exposure (Co-I); S.R. Myers (PI) 07/01/99 – 06/30/04	NIH/NIEHS	\$2,330,585
Analysis of Environmental Carcinogens in Urine of Exposed Children and Adults (Co-I); S.R. Myers (PI) 10/01/99 – 09/30/02	USEPA	\$750,986
Pediatric Pharmacology Research Unit (Co-I); J.E. Sullivan (PI) 01/01/99 – 12/31/03	NIH/NICHD	\$1,579,652

	<u>Agency</u>	<u>Budget Requested</u>
<b>Dr. Harrell E. Hurst</b>		
The Effect of Ozone on the Production of Insect Repellant Compounds in Plants (Co-I); A. senGupta (PI) 08/01/98 – 07/31/01	US EPA	\$250,756
Pharmacogenetics of Fetal Tobacco Smoke Exposure (Co-I); S.R. Myers (PI) 12/01/98 – 11/30/03	NIH	\$1,840,554
Pediatric Pharmacology Research Unit (Co-I); J.E. Sullivan (PI) 01/01/99 – 12/31/03	NIH/NICHD	\$1,579,652
Analysis of Environmental Carcinogens in Urine of Exposed Children and Adults (Co-I); S.R. Myers (PI) 10/01/99 – 09/30/02	US EPA	\$750,986
<b>Dr. Y. James Kang</b>		
Butylated hydroxyanisole-mediated hypercholesterolemia (PI), submitted 02/01/98	NIH-NIDDK	\$650,000
Metallothionein and adriamycin cardiotoxicity, submitted 07/01/98	NIH-NHLBI	\$623,000
Metallothionein and adriamycin cardiomyopathy, submitted 07/15/98	Jewish Hospital Foundation	\$50,000
<b>Dr. Steven R. Myers</b>		
Pediatric Pharmacology Research Unit (Co-I); J.E. Sullivan (PI) 01/01/99 – 12/31/03	NIH-NICHHD	\$1,579,652
Children's Health Center for Children's Environmental Health and Disease Prevention (PI) 08/01/98 – 07/31/02	NIH	\$3,368,844
Analysis of Environmental Carcinogens in Urine of Exposed Children and Adults (PI) 10/01/99-09/30/02	USEPA	\$750,986
Pharmacogenetics of Fetal Tobacco Smoke Exposure (PI) 12/1/98 – 11/30/03	NIH	\$1,840,554

	<u>Agency</u>	<u>Budget Requested</u>
<b>Dr. Steven R. Myers (cont.)</b>		
Pharmacogenetics of Fetal Tobacco Smoke Exposure (P.I.) 7/1/99 – 6/30/04	NIH	\$2,330,585
<b>Dr. Donald E. Nerland</b>		
Vinyl Monomers: Detoxified by Glutathione S-Transferases (PI) 02/01/00 – 01/31/04	NIH	\$722,647
<b>Dr. William M. Pierce, Jr.</b>		
Neutrophil Respiratory Burst in Azotemia (Co-I); R. Ward (PI) 1999 - 2002	NIH	\$300,000
Estrogens: Bone Blood Flow and Bone Mechanical Properties (Co-I); J. Fleming (PI) 1999 - 2000	NIH	\$50,000
Regulation of Neutrophil Activation (Co-I); K. McLeish (PI) 1999-2002	VA RAGS	\$300,000
Tyrosine Phosphorylation of Platelet Ca <sup>2+</sup> -ATPase (Co-I); W. Dean (PI) 1998 – 2001	American Heart	\$200,000
Aging Induced Renal Dysfunction: G Protein Activity (Co-I); J.C. Passmore (PI) February 1, 1999 – January 31, 2001	NIH	\$268,731
Glycosphingolipid Metabolism and Trafficking (Co-I); W.W. Young (PI) 1999-2002	NIH	\$300,000
CYP Isoforms and Cardiovascular Function (Co-I); A. Adeagbo (PI) 02/01/99-01/31/2001	NIH	\$70,000
Undergraduate Research Training (Mentor); R. Wittebort (PI) 1998-2001	Howard Hughes Medical Institute	
Riverine Food Webs and their Roles in Contaminant Transport (Co-I); P. Bukaveckas (PI) 1998-1999	EPA-EPSCoR	\$168,500

	<u>Agency</u>	<u>Budget Requested</u>
<b>Dr. Peter P. Rowell</b>		
Functional Activity of Mesolimbic Nicotinic Receptors (PI) 07/01/99 – 06/30/99	NIH	\$207,991
<b>Dr. Walter M. Williams</b>		
Role of Genetic Polymorphism in Liver Toxicity (Collaborator); T.S. Chen (PI) 09/01/98 – 08/31/99	IDEA Pilot Proposal	\$24,325

## B. Research Grants and Contracts in Force

	<u>Agency</u>	<u>Project Award</u>
<b>Dr. Frederick W. Benz</b>		
Acute Acrylonitrile Intoxication: Antidotal Assessment (PI) 09/01/98 – 08/31/99	NIH-NIEHS	\$108,088
Acute Acrylonitrile Intoxication: Antidotal Assessment (PI) 09/01/97 – 08/31/98	NIH-NIEHS	\$103,930
<b>Dr. Laurence A. Carr</b>		
An Immunomodulatory Role for Brain Dopamine Systems (PI) 09/01/98-09/01/99	SOM Research Committee	\$7,920
<b>Dr. Theresa S. Chen</b>		
Acute Acrylonitrile Intoxication: Antidotal Assessment (Co-I); F.W. Benz (PI) 09/01/97 – 08/31/98	NIH-NIEHS	\$105,432
Acute Acrylonitrile Intoxication: Antidotal Assessment (Co-I); F.W. Benz (PI) 09/01/98 – 08/31/99	NIH-NIEHS	\$108,088
Glutathione Status and Lipid Peroxidation in Chronic Toxicity (P.I.) 06/01/96 – 03/31/98	ULCHES-NIEHS	\$16,000
Role of Genetic Polymorphism in Liver Toxicity (P.I.) 09/01/98 – 08/31/99	IDEA Pilot Proposal	\$8,077
<b>Dr. David W. Hein</b>		
Pharmacogenetics of Drug and Carcinogen Metabolism (P.I.) 09/01/97 – 06/30/02	NIH-NCI	\$1,859,936
Metabolic Basis of Sulfonamide Toxicity in AIDS patients (P.I. on subproject) 09/01/97 – 05/31/00	NIH-Subcontract from Wayne State	\$80,349

	<u>Agency</u>	<u>Project Award</u>
<b>Dr. David W. Hein (cont.)</b>		
Role of genetic polymorphism in liver toxicity (Co-I); T.S. Chen (PI)	IDEA Pilot Proposal	\$8,077
<b>Dr. Harrell E. Hurst</b>		
The Center for the Study of Environmental Hazards to Reproductive Health (Co-I); F.J. Hendler (PI) 01/01/97 - 12/31/99	Department of Veterans Affairs	\$949,100
Acute Acrylonitrile Intoxication – Antidotal Assessment (Co-I); F.W. Benz (PI) 09/01/98 - 08/31/99	NIEHS	\$108,088
HPLC Electrospray Mass Spectrometer (Co-I); W.M. Pierce (PI) 06/01/97 – 05/31/98	NIH/NCRR	\$307,954
Contract for Drug Screening, (Director) 07/01/97 – 06/30/98	UofL Athletics Association	\$37,500
Freshness Phase I: Odor & Ethylene Removal (Co-I); C. Staff (PI) 08/17/98 – 11/30/98	General Electric Company	\$60,177
<b>Dr. Y. James Kang</b>		
Effect of Catalase on Adriamycin Cardiotoxicity (PI) 08/01/95-07/31/00	NIH-NCI	\$507,500
Suppression of Cardiac Ischemia-Reperfusion Injury by Overexpression of metallothionein (PI) 01/01/97-12/31/00	American Heart Association	\$300,000
The role of Catalase in Protection Against Copper Deficiency-Induced Heart Damage in Transgenic Mice (PI) 09/01/95-10/31/98	USDA	\$140,000
Metallothionein and Alcoholic Liver Injury (PI) 12/01/98-10/31/99	UofL Medical School	\$15,000

	<u>Agency</u>	<u>Project Award</u>
<b>Dr. Donald E. Nerland</b>		
Acute Acrylonitrile Intoxication: Antidotal Assessment (Co-I); F.W. Benz (PI) 09/01/97-08/31/98	NIH-NIEHS	\$103,930
Acute Acrylonitrile Intoxication: Antidotal Assessment (Co-I); F.W. Benz (PI) 09/01/98-08/31/99	NIH-NIEHS	\$108,088
<b>Dr. William M. Pierce, Jr.</b>		
Bone Targeting of Pharmaceutical Agents for Osteoporosis – Phase 2 (P.I.) 09/1/97 – 07/31/98	KOWA	\$255,148
Bone Targeting of Pharmaceutical Agents for Osteoporosis – Phase 3 (P.I.) 08/01/98 – 09/30/98	KOWA	\$32,692
Bone Targeting of Estrogens for Osteoporosis (P.I.) 10/1/98 – 9/30/00	KOWA	\$316,708
HPLC Electrospray Mass Spectrometer (P.I.) 6/1/97 – 05/31/98	NIH SIG	\$307,954
Blood Lead Monitoring (P.I.) 01/01/98 – 12/31/98	Service to Regional Practitioners	\$20,113
Atomic Absorption Spectrophotometer (P.I.) 08/01/98 – 07/31/99	WHAS Crusade for Children	\$50,000
<b>Dr. Peter P. Rowell</b>		
Functional Activity of Presynaptic Nicotinic Receptors (PI) 06/01/95-05/31/99	NIH	\$185,070
<b>Dr. Leonard C. Waite</b>		
Bone Targeting of Pharmaceutical Agents for Osteoporosis – Phase 2 (Co-I); W.M. Pierce (PI) 09/01/97 – 07/31/98	KOWA	\$255,148

	<u>Agency</u>	<u>Project Award</u>
<b>Dr. Leonard C. Waite (cont.)</b>		
Bone Targeting of Pharmaceutical Agents for Osteoporosis – Phase 3 (Co-I); W.M. Pierce (PI) 08/01/98 – 09/30/98	KOWA	\$32,692
Bone Targeting of Estrogens for Osteoporosis (Co-I); W.M. Pierce (PI) 10/01/98 – 09/30/00	KOWA	\$316,708

## **VIII. Summary of Teaching Activities**

### **A. School of Medicine**

The Department team-taught the Medical Pharmacology course to second year medical students. Dr. Mike Williams served as course director. In addition, Dr. Laurence Carr served as co-director of the interdisciplinary Clinical Neuroscience course. Individual faculty contributions are identified in the Appendix.

### **B. School of Dentistry**

The Department team-taught the Dental Pharmacology and Therapeutics course to second year dental students. Dr. Leonard Waite served as course director. Individual faculty contributions are identified in the Appendix.

### **C. School of Nursing**

The Department team-taught a Basic Pharmacology for Nursing course to second year nursing students. Dr. William Pierce served as course director. Individual faculty contributions are identified in the Appendix.

The Department team-taught an Advanced Pharmacology course to graduate nursing students. Dr. Leonard Waite served as course director. Individual faculty contributions are identified in the Appendix.

### **D. School of Allied Health**

The Department team-taught a Pharmacology course to second year students in the Dental Hygiene Program. Dr. Leonard Waite served as course director. Individual faculty contributions are identified in the Appendix.

### **E. Graduate School**

The Department team-taught several courses for graduate students. The individual courses and course directors were as follows:

Principles of Drug Action (Dr. Frederick Benz)

Research Methods (Dr. William Pierce)

Pharmacology Seminar (Dr. Donald Nerland)

Individual faculty contributions to these courses are identified in the Appendix.

Individual faculty served on a number of graduate student committees as identified in the Appendix.

## **IX. Summary of Service Activities**

Faculty provided service to the Department, the School of Medicine, the University of Louisville, the profession, the nation, and the community in many ways. Individual faculty service activities are identified in the Appendix.

## **X. Honors and Awards**

**Jian Cai** received the Dr. Kee Chang Huang outstanding graduate student award, and received the Second Place Research Poster Award at the annual meeting of the Ohio Valley Society of Toxicology.

**Adrian Fretland** received a Travel Award to present his research at the annual meeting of the Society of Toxicology.

**David Hein** received the Second Place Research Award "Potential for Major Clinical Applications" from the Jewish Hospital Foundation at Research!Louisville.

**James Kang** was the keynote speaker of the Toxicology Society of Taiwan.

**Matthew Leff** received the First Place Research Poster Award, at the annual meeting of the Ohio Valley Society of Toxicology, received a Travel Award to present his research at the annual meeting of the Society of Toxicology, and received an Honorable Mention Research Poster Award at the University of Louisville Health Sciences Center Student Research Day

**William J. Waddell** was appointed Professor and Chair Emeritus.

**Mike Williams** received the University of Louisville School of Medicine-Gender Equity Award from the American Medical Women's Association and received the Golden Apple Award for Teaching from the University of Louisville School of Medicine Class of 2000.

## **XI. Standing Committee Membership**

### **Graduate Program Committee**

Dr. Len Waite (Chair)

Dr. Peter Rowell (2000)

Dr. Fred Benz (1999)

Dr. William Pierce (1998)

### **PBSI/Grievance Committee**

Dr. Peter Rowell (Chair)  
Dr. Mike Williams (2000)  
Dr. Don Nerland (1999)  
Dr. Harrell Hurst (1998)

### **Teaching Evaluation Committee**

Dr. Larry Carr (Chair)  
Dr. Mike Williams (2000)  
Dr. Len Waite (1999)  
Dr. Fred Benz (1998)

### **Seminar Committee**

Dr. Don Nerland (Chair)  
Dr. Theresa Chen (2000)  
Dr. Harrell Hurst (1999)  
Dr. Steve Myers (1998)

### **Core Laboratories/Research Development Committee**

Dr. William Pierce (Chair)  
Dr. Steve Myers (2000)  
Dr. Larry Carr (1999)  
Dr. Theresa Chen (1998)