

Keynote Speech

<Keynote Speech Topic>

Studies on the effect of pesticides on reproductive and developmental toxicity and endocrine disrupting activity in rats

Shui-Yuan Lu

Taiwan Agricultural Chemicals and Toxic Substances Research Institute, Council of Agriculture, Executive Yuan

11, Kuang-Ming Rd., Wufeng District, Taichung City, 41358, Taiwan

Phone: 04 2330 2101 ext. 505; 501

Fax: 04 2332 3073

e-mail: lusueyen@tactri.gov.tw



Education:

8/06 Ph.D. National Taiwan University College of Medicine, Graduate Institute of Toxicology, Taipei, Taiwan

Major: Reproductive and developmental toxicity and endocrine disrupting activity in mammals

Minor: Embryology

8/92 M.S. Tunghai University, Taichung, Taiwan, R.O.C.

Major: Animal Science

6/86 B.S. Tunghai University, Taichung, Taiwan, R.O.C.

Major: Animal Science

Research experience:

1988~1989: Research Assistant at Taiwan Pig Research Institute

1991~1992: Research Assistant at Livestock Research Institute, Council of Agriculture, Executive Yuan

1992~1993: Assistant Researcher at Taiwan Agricultural Chemicals and Toxic Substances Research Institute, Council of Agriculture, Executive Yuan

1993~1997: Assistant Researcher at Taiwan Agricultural Chemicals and Toxic Substances Research Institute, Council of Agriculture, Executive Yuan

1998~ 2007.08: Assistant researcher at Taiwan Agricultural Chemicals and Toxic Substances Research Institute, Council of Agriculture, Executive

Keynote Speech

Yuan

2007.08~2020.09.01 : Associate Researcher at Taiwan Agricultural Chemicals and Toxic Substances Research Institute, Council of Agriculture, Executive Yuan

2020.09.02~2021.07.15 : Researcher at Taiwan Agricultural Chemicals and Toxic Substances Research Institute, Council of Agriculture, Executive Yuan

2021.07.16~ : Researcher and Chief of Applied Toxicology Division at Taiwan Agricultural Chemicals and Toxic Substances Research Institute, Council of Agriculture, Executive Yuan

Speaker Biographies :

Selected publications

1. 呂水淵、楊德威、李坤雄、劉世華、王家宇。1992. 短暫舒解熱緊迫對荷蘭女牛受胎率和生殖內泌素之影響。中國畜牧學會會誌 21 : 369-383.
2. 王家宇、呂水淵、胡淵欽、楊德威。1993. 夏季冷卻處理對荷蘭母牛泌乳及生殖性狀之影響。中國畜牧學會會誌 22 (2) : 163-173.
3. 王家宇、楊德威、楊忠亮、呂水淵。1995. 預先餵飼盤固乾草對早泌乳牛葉炎發生率之影響。中國畜牧學會會誌 24 (3): 311-328.
4. 呂水淵、林宏偉、王順成。1995. 氨基甲酸鹽農藥免賴得 (Benomyl) 對大鼠胚胎畸形性之探討。中華民國獸醫學會會誌 20 (4) : 348-356.
5. 呂水淵、林宏偉、王順成。1995. 殺菌劑貝芬替 (Carbendazim) 對鼠致胚胎畸形性評估。植保會刊 37 : 331-338.
6. 呂水淵、王順成。1996. 五種農藥對大鼠子代外觀畸形與骨骼之影響。中華民國獸醫學會會誌 22 (6) : 402-412.
7. 呂水淵、王順成。1998. 甲氧基護谷與 2,4-地除草劑誘發大鼠子代外觀、細部骨骼及內臟異常性評估。中華民國獸醫學會會誌 24 (1) : 29-41.
8. 呂水淵、王順成。結構相似之免賴得與甲基多保淨農藥誘發大鼠畸形性之比較。台灣畜牧獸醫學會會報 67 (3-4) : 161-171, 1997.
9. 呂水淵、王順成。農藥免賴得致胚胎畸形及其測試方法之建立。中華民國獸醫學會會誌 25 (1) : 69-76, 1999
10. 呂水淵、王順成。三種農用藥劑對小鼠精子活力、體外受精率及胚胎發育之影響。中華民國獸醫學會會誌, 26(1): 24-35, 2000.
11. Lu S. Y., Liao J. W., Kuo M. L., Wang S. C., Hwang J. S., Ueng T. H. (2004) Endocrine-disrupting activity in carbendazim-induced reproductive and developmental toxicity in rats. *Journal of Toxicology and Environmental Health Part A*, **67**, 1501-1515.
12. Lu S. Y., Liao J. W., Kuo M. L., Hwang J. S., and Ueng T. H. (2006) Antagonistic and synergistic effects of carbendazim and flutamide exposures *in utero* on reproductive and developmental toxicity in rats. *Journal of Food and Drug Analysis* **14**, 120-132
13. Chan P. K., Lu S. Y., Liao J. W., Wei C. F., Tsai Y. Y., and Ueng T. H. (2006) Induction and inhibition of cytochrome P450-dependent monooxygenases of rats by fungicide bitertanol. *Food and Chemical Toxicology* **44(12)**, 2047-2057.

Keynote Speech

14. Jing-Ying Huang, Jiunn-Wang Liao, Yi-Chun Liu, Shui-Yuan Lu, Chen-Ping Chou, Wei-Hung Chan, Shee-Uan Chen, and Tzoo-Huei Ueng. (2008) Motorcycle exhaust induces reproductive toxicity and testicular interleukin-6 in male rats. *Toxicological Sciences* **103**(1), 137-148.
15. Chen-Ping Chou, Shui-Yuan Lu, Tzoo-Huei Ueng. 2008. Modulation of serum concentrations and hepatic metabolism of 17 β -estradiol and testosterone by amitraz in rats. *Arch Toxicol* DOI 10.1007/s00204-008-0288-8
16. 呂水淵、張瓊瑋、許雅惠、袁朝云、陳敏貞、陳憬輝、馬瑞悌、游碧琦。2010。蔬果中高殘留檢出率農藥對大鼠出生前發育之毒理影響評估。植保會刊 52(4) :99-116
17. 呂水淵、陳敏貞、袁朝云、許雅惠、張瓊瑋、陳憬輝、馬瑞悌、游碧琦。2010。十三種美國環保署待評估農藥其他成分之致變異性探討。植保會刊 52: 43-68
18. Ya-Hui Hsu, Chiung-Wei Chang, Min-Chen Chen, Chao-Yun Yuan, Jing-Huei Chen, Jui-Ti Ma, Tzoo-Huei Ueng, Shui-Yuan Lu. 2011 Carbendazim-induced androgen receptor expression antagonized by flutamide in male rats. *Journal of Food and Drug Analysis* 19(4): 418-428.
19. Chin-Fang Huang, Yu-Shun Lin, Zong-Cian Chiang, Shui-Yuan Lu, Yueh-Hsinung Kuo, Sunny Li-Yun Chang, Pei-Min Chao. 2014. Oxidized components of frying oil ingested during pregnancy disturb vitamin A metabolism and are potentially teratogenic in mice. *Journal of Nutritional Biochemistry* 25:549-556
20. 呂水淵。2015。應用基準劑量技術評估4種疑似內分泌干擾農藥之風險。104年農藥內分泌干擾作用專題研討會論文：54-66
21. Shui-Yuan Lu, Min-Chen Chen, Chao-Yun Yuan, Ya-Hui Hsu and Wei-Ren Tsai . 2015. Detecting Benomyl and Its Metabolite Carbendazim Inducing Androgenic Activity in Rats by Using Uterotrophic and Hershberger Assays. *Taiwan Journal of Agricultural Chemistry and Food Science* 53(6): 235-250 (灣農業化學與食品科學期刊)
22. C. M. Chang, Y. H. Ou, T. C. Liu, S.Y. Lu and M. K. Wang. 2016. A quantitative structure-activity relationship approach for assessing toxicity of mixture of organic compounds. SAR and QSAR in Environmental Research. <http://dx.doi.org/10.1080/1062936X.2016.1207204>
23. 呂水淵、牟為謙、陳敏貞、蔡韃任。2016。應用基準劑量推估31種農藥每日可接受攝食量做為健康風險評估之可行性探討。台灣農藥科學 (Taiwan Pesticide Science) 1: 24-49
24. Wei-Chun Chou, Wei-Ren Tsai, Hsiu-Hui Chang, Shui-Yuan Lu, King-Fu Lin, Pinpin Lin. 2018. Prioritization of pesticides in crops with a semi-quantitative risk ranking method for Taiwan postmarket monitoring program. *Journal of Food and Drug Analysis*. <http://creativecommons.org/licenses/by-nc-nd/4.0/>
25. Shui-Yuan Lu and Wei-Ren Tsai. 2018. Androgen receptor plays a vital role in benomyl- or carbendazim-induced reproductive and developmental toxicity and endocrine disrupting activity in rats. Book "Endocrine Disruptors" published by IntechOpen Limited. Registered in England and Wales No. 11086078. London, United Kingdom. ISBN 978-1-78984-151-0.
26. Shui-Yuan Lu, Pinpin Lin, Wei-Ren Tsai, Chen-Yi Weng. 2018. The pragmatic strategy to detect endocrine disrupting activity of xenobiotics in food. Book "Medicinal Chemistry" published by IntechOpen Limited. Registered in England and Wales No. 11086078. London, United Kingdom. ISBN 978-1-78985-173-1.
27. Shui-Yuan Lu, Pinpin Lin, Wei-Ren Tsai, Chen-Yi Weng. 2018. Deltamethrin alters thyroid hormones and delays pubertal development in male and female rats. Book "Comparative Endocrinology of Animals" published by IntechOpen Limited. Registered in England and Wales No. 11086078. London, United Kingdom. ISBN

Keynote Speech

- 978-1-83880-396-4.
28. Shui-Yuan Lu and Wei-Ren Tsai. 2018. Endocrine disrupting investigations of six pesticides with estrogen receptor binding assays and uterotrophic effects in rats. *Taiwan Pesticide Science* 5: 31-51
 29. Yu-Shun Lin, Shui-Yuan Lu, Hai-Ping Wu, Chi-Fen Chang, Yung-Tsung Chiu, Hui-Ting Yang, Pei-Min Chao. 2019. Is frying oil a dietary source of an endocrine disruptor? Anti-estrogenic effects of polar compounds from frying oil in rats. *Ecotoxicology and Environmental Safety* 169: 18-27
 30. 廖俊麟、林春蘭、呂水淵、蔡建任。2018。臺灣常用農藥之慢性危害風險排序。臺灣農藥科學 5: 53-73
 31. Shui-Yuan Lu, Min-Chen Chen, Jing-Chun Liao, and Wei-Ren Tsai. 2019. Tebuconazole decrease 17β -estradiol and aromatase activity in pubertal developmental rats. *Taiwan Pesticide Science* 7: 81-108
 32. 呂水淵。2019。淺談歐盟關於農藥累積(cumulative)殘留暴露對人體毒性評估進展。台灣風險分析學會會刊
 33. Shui-Yuan Lu and Wei-Ren Tsai. 2020. Benomyl- or carbendazim-induced androgen receptor disrupting might lead to spinal and bulbar muscular atrophy in multi-generations of rats. *Open Access Journal of Pharmaceutical Research* 4(1): 000198
 34. Hai-Ping Wu , Yu-Shun Lin , Chi-Fen Chang , Shui-Yuan Lu , Pei-Min Chao. 2020. Dietary exposure to oxidized frying oil from fetus to adulthood suppresses male reproductive development by altering testicular cholesterol and testosterone homeostasis in Sprague Dawley rats. *The Journal of Nutrition* 00: 1-9
 35. Shui-Yuan Lu and Wei-Ren Tsai. 2020. Antiandrogen flutamide might protect multi-generations of rats from androgen-dependent toxicity in spinal and bulbar muscular atrophy induced by benomyl or carbendazim. *EC Pharmacology and Toxicology* 8.6 (2020): 94-97.
 36. Shui-Yuan Lu, ¹Min-Chen Chen, ¹Jing-Chun Liao, ¹Wan-Hsin Chen, ¹Wei-Ren Tsai. 2021. Prochloraz decreased serum concentration of E2, testosterone, T3, T4 and aromatase activity in pubertal developmental rats. *Taiwan Pesticide Science*

Keynote Speech

Abstract :

Based on guidance for the identification of endocrine disruptors in the context of Regulations (EU) No 528/2012 and (EC) No 1107/2009 with Level 2 to Level 5 and Office of Chemical Safety and Pollution Prevention (OCSPP) Series 890 - Endocrine Disruptor Screening Program Test Guidelines we carried out to investigate the effect of some pesticides on reproductive and developmental toxicity and endocrine disrupting activity in rats. These studies included Level 2- In vitro assays providing data about selected endocrine mechanism(s)/ pathways(s), Level 3- In vivo assays providing data about selected endocrine mechanism(s)/ pathway(s), Level 4- In vivo assays providing data on adverse effects on endocrine relevant endpoints and Level 5- In vivo assays providing more comprehensive data on adverse effects on endocrine relevant endpoints over more extensive parts of the life cycle of the organism. Due to the specificity of mode of action positive reaction of reproductive and developmental toxicity and endocrine disrupting activity was mainly in pesticide classification of benzimidazole, imidazole, triazole, pyrethroids, dicarboximides and aryloxyphenoxypropionate. So far most of studies of endocrine disrupting activity based on Level 2-5 and OCSPP Series 890 were focused on estrogen (E), androgen (A) and thyroid (T) receptor-related. Among these studies of Level 2 to Level 5 developmental neurotoxicity study is studied relative scarce. We found out that one of benzimidazole pesticide might induce developmental neurotoxicity through androgen receptor in rats. The developmental neurotoxicity induced by one of benzimidazole seems to be more serious than that of chlorpyrifos, which was banned by USA with acetylcholinesterase inhibition. Based on connecting previous study and our finding we inferred estrogen and androgen receptor disrupting pesticides might induce developmental neurotoxicity in mammals. Facing to the endocrine disrupting pesticides we will take appropriate management with highly hazard pesticides (HHPs) to evaluate hazard identification and make risk reduction in Taiwan. More completely and efficiently study and management should be done in reproductive and developmental toxicity and endocrine disrupting activity in Taiwan.