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Publications

Journal Papers

1. Shu-Mei Wang, Chia-Yu Chang, Giunn-Jye Chuu & Ching-Kuo Lee. Assessing therapeutic efficacy of *Chlorella vulgaris* peptides on pathogenic bacteria associated with periodontal disease. *CyTA - Journal of Food*. 2025 Feb 本人為通訊作者
2. Wang SM, Chuu JJ, Lee CK, Chang CY. Exploring the therapeutic efficacy of *Chlorella pyrenoidosa* peptides in ameliorating Alzheimer's disease. *Heliyon*. 2025 Apr 本人為通訊作者
3. Chang, En-Su, Cheng-Yi Hsu, Jui-Chen Li, Rajni Dubey, Wei-Chih Kan, Ming-Jen Sheu, Chia-Hui Cheng, Chia-Yu Chang, Chin-Chia Hsu, and Giunn-Jye Chuu. 2025. "Therapeutic Potential of Probiotic *Lactobacillus* Spp. Fermented *Talinum Triangulare* on *Helicobacter Pylori* Associated Gastrointestinal Infectious Disorders." *CyTA - Journal of Food* 23 (1). 2025 Jan. 本人為通訊作者
4. Chiang KH, Cheng TJ, Kan WC, Wang HY, Li JC, Cai YL, Cheng CH, Liu YC, Chang CY, **Chuu JJ**. *Orthosiphon aristatus* (Blume) Miq. Extracts attenuate Alzheimer-like pathology through anti-inflammatory, anti-oxidative, and β -amyloid inhibitory activities. *J Ethnopharmacol*. 2024 Feb 10;320:117132. **Impact Factor (IF)=5.4** (2023 update), 本人為通訊作者
5. **Chou JJ**, Jeng-Wei Lu, Hung-Ju Chang, You-Hsiang Chu, Yi-Jen Peng, Yi-Jung Ho, Pei-Hung Shen, Yu-Shuan Chen, Chia-Hui Chen, Yi-Chien Liu, Chih-Chien Wang. Attenuative effects of collagen peptide from milkfish (*Chanos chanos*) scales on ovariectomy-induced osteoporosis. *Food Science & Nutrition Early View*. 2023 September 21. **Impact Factor (IF)=3.55** (2023 update), 本人為第一作者
6. Chiang KH, Cheng TJ, Kan WC, Wang HY, Li JC, Cai YL, Cheng CH, Liu YC, Chang CY, **Chuu JJ**. *Orthosiphon aristatus* (Blume) Miq. Extracts attenuate Alzheimer-like pathology through anti-inflammatory, anti-oxidative, and β -amyloid inhibitory activities. *Journal of Ethnopharmacology*. 2023 September 11;117132. **Impact Factor (IF)=5.4** (2023 update), 本人為通訊作者
7. Kuang-Hsing Chiang, Yi-Chun Chiu, Noi Yar, Yu-Chun Chen, Chia-Hui Cheng, Yi-Chien Liu, Chia-Yu Chang and **Chuu JJ**. Renoprotective Impacts of *Inonotus obliquus* Ethanol-Ethyl Acetate Extract on Combined Streptozotocin and Unilateral Nephrectomy-Induced Diabetic Nephropathy in Mice. *Int J Mol Sci*. 2023 Feb 23;24(5):4443. **Impact Factor (IF)=5.6** (2023 update), 本人為通訊作者
8. Shu-Mei Wang, **Chuu JJ**, Chia-Yu Chang, Ching-Kuo Lee. Exploring the Therapeutic Efficacy of *Chlorella Pyrenoidosa* Peptides in Ameliorating Alzheimer's Disease. *Heliyon*. 2023 Apr 11;9(5):e15406. **Impact Factor (IF)=4** (2023 update).

9. Wang CC, Lu JW, Chiang KH, Cheng YS, Chu YH, Peng YJ, Cheng CH, Chang CY, **Chuu JJ**. Anti-Inflammatory and Chondro-Protective Effects of Acidic Polysaccharide from *Enteromorpha Prolifera* in Experimental Models of Osteoarthritis In-Vitro and In-Vivo. *Cartilage*. 2022 Dec;13(4):157-170. **Impact Factor (IF)=2.8** (2023 update),本人為通訊作者
10. Huang KC, Li JC, Wang SM, Cheng CH, Yeh CH, Lin LS, Chiu HY, Chang CY, **Chuu JJ**. The effects of carbon monoxide releasing molecules on paraquat-induced pulmonary interstitial inflammation and fibrosis. *Toxicology*. Available online 15 March 2021, In press, journal pre-proof, Article 152750. **(SCI) JIF: 4.5** (2023 update),本人為通訊作者
11. Chou YJ, **Chuu JJ**, Peng YJ, Cheng YH, Chang CH, Chang CM, Liu HW. The potent anti-inflammatory effect of Guilu Erxian Glue extracts remedy joint pain and ameliorate the progression of osteoarthritis in mice. *J Orthop Surg Res*. 2018 Oct 19;13(1):259. **(SCI) JIF:2.677** (2023 update)
12. Yu SH, Chen,ST, Li WS, Dubey NK, Chen WH, **Chuu JJ**, Leu SJ, Deng WP. Corrigendum to Hypoglycemic Activity through a Novel Combination of Fruiting Body and Mycelia of *Cordyceps militaris* in High-Fat Diet-Induced Type 2 Diabetes Mellitus Mice. *J Diabetes Res*. 2017 Mar 16. **(SCI) JIF:4.061** (2023 update)
13. Chou YJ, Kan WC, Chang CM, Peng YJ, Wang HY, Yu WC, Cheng YH, Jhang YR, Liu HW, **Chuu JJ**. Renal Protective Effects of Low Molecular Weight of *Inonotus obliquus* Polysaccharide (LIOP) on HFD/STZ-Induced Nephropathy in Mice. *Int J Mol Sci*. 2016 Sep 13;17(9). **(SCI) JIF:6.028** (2023 update), 本人為通訊作者
14. Yu SH, Chen SY, Li WS, Dubey NK, Chen WH, **Chuu JJ**, Leu SJ, Deng WP. Hypoglycemic Activity through a Novel Combination of Fruiting Body and Mycelia of *Cordyceps militaris* in High-Fat Diet-Induced Type 2 Diabetes Mellitus Mice. *J Diabetes Res*. 2015;2015:723190. **(SCI) JIF:4.061** (2023 update)
15. Chang CY, Guo HR, Tsai WC, Yang KL, Lin LC, Cheng TJ, **Chuu JJ**. Subchronic arsenic exposure induces anxiety like behaviors in normal mice and enhances depression like behaviors in the chemically induced mouse model of depression. *BioMed Research International*. 2015;2015:159015. **(SCI) JIF:3.246** (2023 update), 本人為通訊作者
16. Wang HY, Kan WC, Cheng TJ, Yu SH, Chang LH, **Chuu JJ**. Differential anti-diabetic effects and mechanism of action of charantin-rich extract of Taiwanese *Momordica charantia* between type 1 and type 2 diabetic mice. *Food and chemical toxicology*. 2014; 69:347-56. **(SCI) JIF: 5.572** (2023 update), 本人為通訊作者
17. Liao YW, Chen CR, **Chuu JJ**, Huang HC, Hsu JL, Huang TC, Kuo YH, Chang CI. Cucurbitane Triterpenoids from the Fruit Pulp of *Momordica charantia* and Their Cytotoxic Activity. *Journal Of The Chinese Chemical Society*. 2013;60(5):526-30. **(SCI) JIF:1.753** (2023 update)
18. Chen YP, Dai ZH, Liu PC, **Chuu JJ**, Lee KY, Lee SL, Chen YJ. Effects of nanogold on the alleviation of carbon tetrachloride-induced hepatic injury in rats. *The Chinese Journal of Physiology*. 2012;55(5):331-36. **(SCI) JIF: 1.568** (2023 update)
19. Yin SH, Wang CC, Cheng TJ, Chang CY, Lin KC, Kan WC, Wang HY, Kao WM, Kuo YL, Chen JC, Li SL, Cheng CH, **Chuu JJ**. Room-temperature super-extraction system (RTSES) optimizes the anxiolytic- and antidepressant-like behavioural effects of traditional Xiao-Yao-San in mice. *Chinese Medical Journal*. 2012; 7(1):24. **(SCI) JIF: 6.133** (2023 update), 本人為通訊作者
20. Li WT, Hwang DR, Song JS, Chen CP, Chen TW, Lin CH, **Chuu JJ**, Lien TW, Hsu TA, Huang CL, Tseng HY, Lin CC, Lin HL, Chang CM, Chao YS, Chen CT. Synthesis and biological evaluation of 2-amino-1-thiazolyl imidazoles as orally active anticancer agents. *Investigational New Drugs*. 2012;30(1):164-75. **(SCI) JIF: 3.651** (2023 update)
21. Kan WC, Wang HY, Chien CC, Li SL, Chen YC, Chang LH, Cheng CH, Tsai WC, Hwang JC, Su SB, Huang LH, **Chuu JJ**. Effects of Extract from Solid-State Fermented *Cordyceps sinensis* on Type 2 Diabetes. Mellitus. *Evidence-Based Complementary and Alternative Medicine*. 2012;743107. **(SCI) JIF:2.65** (2023 update), 本人為通訊作者
22. Cheng TJ, **Chuu JJ**, Chang CY, Tsai WC, Chen KJ, Guo HR. Atherosclerosis induced by arsenic in drinking water in rats through altering lipid metabolism. *Toxicology and Applied Pharmacology*. 2011;256(2):146-53. **(SCI) JIF: 3.347** (2023 update)
23. Chang CY, Cheng TJ, Chang FR, Wang HY, Kan WC, Li SL, Huang LH, Chen YC, Tsai WC, Huang CH, Cheng CH, Lee GY, Shyue SW, Chen YP, Lin KC, **Chuu JJ**. Macrophage mediated anti-proliferation effects of *Anthodia camphorata* non-polysaccharide based extracts on human hepatoma cells. *Bioscience, Biotechnology, and Biochemistry*. 2011;75(4):624-32. **(SCI) JIF: 4.46** (2023 update), 本人為通訊作者

24. Hu CB, Chen CP, Yeh TK, Song JS, Chang CY, **Chuu JJ**, Tung FF, Ho PY, Chen TW, Lin CH, Wang MH, Chang KY, Huang CL, Lin HL, Li WT, Hwang DR, Chern JH, Hwang LL, Chang JY, Chao YS, Chen CT. BPR0C261 is a novel orally active antitumor agent with antimitotic and anti-angiogenic activities. *Cancer Science*. 2011;102(1):182-91. (SCI) JIF: 6.518 (2023 update)
25. Li SL, Huang CH, Lin CC, Huang ZN, Chern JH, Lien HY, Wu YY, Cheng CH, Chang CY, **Chuu JJ**. Antitumor effect of BPR-DC-2, a novel synthetic cyclic cyanoguanidine derivative, involving the inhibition of MDR-1 expression and down-regulation of p-AKT and PARP-1 in lung cancer. *Investigational New Drugs*. 2011;29(2):195-206. (SCI) JIF: 3.651 (2023 update), 本人為通訊作者
26. Li WT, Hwang DR, Song JS, Chen CP, **Chuu JJ**, Hu CB, Lin HL, Huang CL, Huang CY, Tseng HY, Lin CC, Chen TW, Lin CH, Wang HS, Shen CC, Chang CM, Chao YS, Chen CT. Synthesis and biological activities of 2-amino-1-arylidenamino imidazoles as orally active anticancer agents. *Journal of Medicinal Chemistry*. 2010;53(6):2409-17. (SCI) JIF: 8.039 (2023 update)
27. Yeh TK, Li CM, Chen CP, **Chuu JJ**, Huang CL, Wang HS, Shen CC, Lee TY, Chang CY, Chang CM, Chao YS, Lin CT, Chang JY, Chen CT. Antitumor activities and pharmacokinetics of silatecans DB-67 and DB-91. *Pharmacological Research*. 2010 ;61(2):108-15 . (SCI) JIF: 10.334 (2023 update)
28. Shun-Lai Li, Chien-Tai Huang, Hsien-Yi Wang, Wei-Chih Kan, Zih-Ning Huang, Guo-Yang Lee, Yi-Peng Chen, Hsiao-Hui Hsieh, **Chuu JJ**. The molecular mechanisms of the action of polysaccharide extract of *Phellinus linteus* in xenografted tumor histoculture and metastatic lung cancer mice model. *Annual Report of Chinese Medicine and Pharmacy*. 2009;203-42. (SCI) JIF: 2.139 (2023 update), 本人為通訊作者
29. Li SL, Huang ZN, Hsieh HH, Yu WC, Tzeng WY, Lee GY, Chen YP, Chang CY, **Chuu JJ**. The augmented anti-tumor effects of *Antrodia camphorata* co-fermented with Chinese medicinal herb in human hepatoma cells. *The American Journal of Chinese Medicine*. 2009;37(4):771-83. (SCI) JIF: 6.005 (2023 update), 本人為通訊作者
30. Lu MC, Du YC, **Chuu JJ**, Hwang SL, Hsieh PC, Hung CS, Chang FR, Wu YC. Active extracts of wild fruiting bodies of *Antrodia camphorata* (EEAC) induce leukemia HL 60 cells apoptosis partially through histone hypoacetylation and synergistically promote anticancer effect of trichostatin A. *Archives of Toxicology*. 2009 ;83(2):121-9. (SCI) JIF: 6.168 (2023 update)
31. Chang CY, Huang ZN, Yu HH, Chang LH, Li SL, Chen YP, Lee KY, **Chuu JJ**. The adjuvant effects of *Antrodia Camphorata* extracts combined with anti-tumor agents on multidrug resistant human hepatoma cells. *Journal of Ethnopharmacology*. 2008 ;118(3):387-95. (SCI) JIF: 4.36 (2023 update), 本人為通訊作者
32. **Chuu JJ**, Huang ZN, Yu HH, Chang LH, Lin-Shiau SY. Attenuation by methyl mercury and mercuric sulfide of pentobarbital induced hypnotic tolerance in mice through inhibition of ATPase activities and nitric oxide production in cerebral cortex. *Archives of Toxicology*. 2008 ;82(6):343-53. (SCI) JIF: 6.168 (2023 update), 本人為第一作者
33. **Chuu JJ**, Liu SH, Lin-Shiau SY. Differential neurotoxic effects of methylmercury and mercuric sulfide in rats. *Toxicology Letters*. 2007 ;169(2):109-20. (SCI) JIF: 4.271 (2023 update), 本人為第一作者
34. **Chuu JJ**, Liu JM, Tsou MH, Huang CL, Chen CP, Wang HS, Chen CT. Effects of paclitaxel and doxorubicin in histocultures of hepatocellular carcinomas. *Journal of Biomedical Science*. 2007;14(2):233-44. (SCI) JIF: 12.771 (2023 update), 本人為第一作者
35. Liu SH, Wang JH, **Chuu JJ**, Lin-Shiau SY. Alterations of motor nerve functions in animals exposed to motorcycle exhaust. *Journal of Toxicology and Environmental Health, Part A*. 2002 ;65(11):803-12. (SCI) JIF: 3.24 (2023 update)
36. Young YH, **Chuu JJ**, Liu SH, Lin-Shiau SY. Neurotoxic mechanism of cinnabar and mercuric sulfide on the vestibulo-ocular reflex system of guinea pigs. *Toxicology Science*. 2002 ;67(2):256-63. (SCI) JIF: 4.109 (2023 update)
37. Young YH, **Chuu JJ**, Liu SH, Lin-Shiau SY. Toxic effects of potassium bromate and thioglycolate on vestibuloocular reflex systems of Guinea pigs and humans. *Toxicology and Applied Pharmacology*. 2001;43(2):91-6. (SCI) JIF: 4.46 (2023 update)
38. **Chuu JJ**, Young YH, Liu SH, Lin-Shiau SY. Neurotoxicity of mercury sulfide in the vestibular ocular reflex system of guinea pigs. *Naunyn-Schmiedeberg's Archives of Pharmacology*. 2001;364(3):249-58. (SCI) JIF: 3.195 (2023 update), 本人為第一作者
39. **Chuu JJ**, Liu SH, Lin-Shiau SY. Effects of methyl mercury, mercuric sulfide and cinnabar on active avoidance responses, Na⁺/K⁺-ATPase activities and tissue mercury contents in rats. *Proceedings of the National Science*

Council, Republic of China - Part B. 2001;25(2):128-36. (SCI) JIF: 4.5 (2023 update), 本人為第一作者

40. **Chuu JJ**, Hsu CJ, Lin-Shiau SY. Abnormal auditory brainstem responses for mice treated with mercurial compounds: involvement of excessive nitric oxide. *Toxicology*. 2001;162(1):11-22. (SCI) JIF: 4.5 (2023 update), 本人為第一作者
41. **Chuu JJ**, Hsu CJ, Lin-Shiau SY. The detrimental effects of potassium bromate and thioglycolate on auditory brainstem response of guinea pigs. *The Chinese Journal of Physiology*. 2000;43(2):91-6. **Impact Factor (IF)=1.8** (2023 update), 本人為第一作者
42. 褚俊傑. 以靈芝多菌株固態發酵培養技術開發新穎機能食品. 食品資訊. 2019 ;289: 58-64
43. 張閔絜, 褚俊傑, 鄭嘉惠, 張禹柔, 周彥瑤. 何首烏單複方對衰老小鼠之抗老化作用北台灣中醫醫學雜誌.. 2017; 1-21.
44. 褚俊傑. 海藻在食品生技領域的應用與展望. 食品資訊. 2015; 270:14-20.
45. 褚俊傑, 鄭天浚, 張嘉祐, 林高章, 洪宸達, 楊淨婷, 鄭勝元, 蔡宛臻, 鄭嘉惠. SPF 雞胚蛋改善動脈硬化的分子作用機制. 南台科技大學學報. 2013; 38:1-15.
46. 褚俊傑. 低溫超臨界蛋黃油萃取技術. 食品資訊. 2011; 245:72-75.
47. 褚俊傑. 桑黃輔助癌症化學治療與評估改善多重藥物抗藥性表現之藥理機制探討. 中醫藥年報. 2009; 27(1):347-380.
48. 褚俊傑, 連小茵, 張嘉祐, 張良豪, 游承翰, 陳文如, 黃梓甯, 游嵩勳. 野生與人工紅景天對小白鼠抗疲勞之藥理研究. 南台科技大學學報 .2008; 32:171-190.

Projects

NSTC Projects:

Year	Project Title (number)	Period
110	Development of a Low-Protein, High-Nutrition, and Functional Food Ingredient Formula Suitable for the Dietary Needs of Patients with Chronic Kidney Disease Using Chlorella Protein Peptides and Bioactive Polysaccharides (MOST 110-2320-B-218-001 -) 2/2	2021/08/01~2022/07/31
109	Biotech Functional Ingredient Pilot-Scale Production Technology Alliance (II) (3/3) (109-2622-8-218-004-TE4)	2020/02/01~2021/04/30
108	Development of a Low-Protein, High-Nutrition, and Functional Food Ingredient Formula Suitable for the Dietary Needs of Patients with Chronic Kidney Disease Using Chlorella Protein Peptides and Bioactive Polysaccharides (MOST 108-2320-B-218-002 -) 1/2	2019/08/01~2020/10/31
108	Biotech Functional Ingredient Pilot-Scale Production Technology Alliance (II) (2/3) (108-2622-8-218-001-TE4)	2019/2/01~2020/4/30
107	Biotech Functional Ingredient Pilot-Scale Production Technology Alliance (II) (1/3) (107-2622-8-218-001-TE4)	2018/2/01~2019/1/31

Government Projects:

Year	Project Title (number)	Period
113	2024 Academia-Industry Cooperation Project - Development of a Natural Stomach Health Material Inhibiting Helicobacter pylori by Pilot-Scale Production of Astaxanthin and Chlorella Growth Factor from Chlorella and Verification of Efficacy (1700-C112005)	2023/11/01~2024/10/31
109	Development of a New Bone Health Food Ingredient Using Small Molecular Functional Collagen Peptides Extracted from Free-Range Chicken Bones (2/2) (MOST 109-2622-B-218-001 -CC1)	2020/11/01~2022/01/31
108	Development of a New Bone Health Food Ingredient Using Small Molecular Functional Collagen Peptides Extracted from Free-Range Chicken Bones (1/2) (MOST 108-2622-B-218-001 -CC1)	2019/11/01~2020/10/31
108	Development of a Functional Water-Soluble Fertilizer with Biostimulant Properties Using Algal Extract Hydrolyzed and Wall-Broken by Fungal Fermentation (17001080014-GP)	2019/01/01~2019/12/31
107	Development of a Microalgal Fermented Protein Feed Concentrate to Reduce Heat Stress and Improve Production Performance in Poultry (MOST 107-2622-B-218-001 -CC3)	2018/11/01~2019/12/31

Industry Collaboration Projects:

Year	Project Title (number)	Period
113	Preclinical Study on the Therapeutic Potential of NK Exosomes in Lupus Nephritis Mice (81001140007)	2025/01/01 ~ 2025/06/30
113	Assessment of the Therapeutic Efficacy of Compound Fermented Probiotic Metabolites Against Multidrug-Resistant Bacterial Infections in a Murine Model (81001140029)	2025/01/01 ~ 2025/06/30
113	Investigation of the Efficacy of Oral Administration of Three Kidney-Protective Products in Clearing Acrolein from Organs and Improving Renal Dysfunction in Mice with Acrolein-Induced Kidney Injury (17001130220)	2024/07/01~2025/06/30
113	Extraction, Isolation, Purification, and Identification of Five Plant Exosomes (EVs) and Evaluation of Their Specific Activities (17001130213)	2024/06/01~2025/05/31

113	Construction of a Clinical IRB Trial Enrollment Platform and Clinical Trial Data Analysis Report Contract (17001130165)	2024/06/01~2025/06/30
113	Contract for the Research and Development of Innovative Products and Brand Marketing Strategies to Improve Chronic Kidney Disease (CKD) (17001130164)	2024/06/01~2025/05/30
113	Investigation of the Renal Protective Effects of AC-134 on Chronic Kidney Disease (17001130151)	2024/05/01~2025/04/30

Patent

1. Extraction, Isolation, Purification, Identification, and Specific Activity Evaluation of Five Plant Exosomes (EVs)
(17001130213-TT)
2. Method for Preparing a Composition Inhibiting Helicobacter pylori
(17001130001-TT)
3. Preclinical Efficacy Evaluation of GV-002 in the Development of Therapeutics for Diabetic Nephropathy
(17001130069-TT)
4. Development of a Natural Stomach Health Material Inhibiting Helicobacter pylori by Pilot-Scale Production of Astaxanthin and Chlorella Growth Factor from Chlorella and Verification of Efficacy
(1700-C112005)
5. A Multifaceted Evaluation Method for a Cancer-Related Fatigue Mouse Animal Model
(17001100006-TT)
6. A Soil Conditioner Developed via Co-Fermentation of Microalgae and Soil Probiotics
(17001090001-TT)
7. Method for Preparing Low-Sodium and Low-Purine Poultry Protein Molecules
(17001070007-TT)
8. Analysis of Peptide Content Determination and Digestive Absorption Rate in Animals
(151040029-TT)
9. Development of a New Traditional Chinese Medicine Formula for the Treatment of Pulmonary Fibrosis Using a Drug Selection Strategy and Bioactivity-Guided Extraction
(MOST 103-2622-B-218-002-CC3)
10. Development of a New Functional Ingredient to Improve Renal Dysfunction Using Active Polysaccharides Produced via Co-Fermentation of Edible and Medicinal Fungi
(151030030-TT)

- 11.** Development of a New Functional Ingredient to Alleviate Chemotherapy Side Effects Using Active Polysaccharides Produced via Co-Fermentation of Edible and Medicinal Fungi
(151030031-TT)
- 12.** Effects of Adding Dunaliella to Poultry Feed on Egg Production Performance and the β -Carotene and Cholesterol Content in Egg Yolks
(151002002-2CC3)
- 13.** Efficacy Evaluation and Health Applications of Type II Collagen Derived from Poultry in Joint Health Products
(151001001-2CC3)
- 14.** Development of a New Functional Food Application for Taiwanese Oranges Using Limonene Extraction Technology
(150981011-2CC3)
Research on the Application of Citrus Limonene in Enhancing Antitumor Effects and Reducing Multidrug Resistance in Liver Cancer Chemotherapy
(150970008-TT)

Professional Certificates

- 1.** Senior Professional and Technical Examination Pharmacist Certificate, Ministry of Examination, Taiwan (Certificate No. 1276)
- 2.** Pharmacist License, Department of Health, Executive Yuan, Taiwan (1993/License No. 018508)
- 3.** 2017 Certificate of Recognition from Western Sydney University - English-Taught Course
- 4.** 2018 IACUC Committee Continuing Education Training Course Certificate
- 5.** 2019 IACUC Committee Continuing Education Training Course Certificate