

## Hui-Chun Wang, Ph.D.

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### PERSONAL

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**Tel:** +886 (07) 312-1101 ext. 6921

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### EDUCATION

*Aug. 1999 to May. 2006*

Graduate Institute of Life Sciences, National Defense Medical Center, Taipei, Taiwan

Ph.D. degree

Graduate study with Professor Dr. C-Y Shen

*“Ataxia telangiectasia mutated and checkpoint kinase 2 regulate BRCA1 to promote the fidelity of DNA end-joining”*

*Aug. 1997 to Jun. 1999*

Graduate Institute of Physiology, National Defense Medical Center, Taipei, Taiwan

M.S. degree

Graduate study with Professor Dr. C-J Tseng

*“Role of oxidative stress in amphetamine-induced neurotoxicity of rat striatum”*

### WORKING EXPERIENCE

*Aug 2018 to present*

**Director**

**Graduate Institute of Natural Products, Kaohsiung Medical University**

*Aug. 2017 to present*

**Professor**

**Graduate Institute of Natural Products, Kaohsiung Medical University**

*Aug. 2013 to Jul.2017*

**Associate Professor**

**Graduate Institute of Natural Products, Kaohsiung Medical University**

*Aug. 2008 to Jul.2013*

**Assistant Professor**

**Graduate Institute of Natural Products, Kaohsiung Medical University**

*Aug. 2006 to Aug. 2008*

**Concurrently Assistant Professor**

**School of public health, National Defense Medical Center**

*Jun. 2006 to Aug.2008*

**Postdoctoral Fellow**

**Institute of Biomedical Science, Academia Sinica**

### AWARDS

AACR-ITO EN, Ltd. Scholar-in-Training Award, USA

Merit Mentors Award, KMU

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Outstanding Research Award, KMU

The Faculty Teaching Award, KMU

Award of Patent Approval, KMU

### **TEACHING COURSES**

1. Special Topics in Development of Natural Products
2. Special Topics on Target-based Research of Natural Products
3. Special Topics on Natural Product Biochemistry
4. Advanced Topics on Biotechnology and Botanical New Drug Research
5. Advanced Topics on Cancer Biology

### **MEMBERSHIP**

1. Member of the Chinese Natural Medicine Society of Taiwan since 2008
2. Active member of the American Association for Cancer Research since 2008

### **TECHNICAL SPECIALTY**

1. Molecular techniques: DNA cloning, genotyping by sequencing, RFLP and TaqMan probe assays, RNAi, NGS, CRISPR
2. Cell culture-based techniques: DNA transfection by methods of liposome and electroporation, RNA interference, flow cytometry, immunofluorescence staining and laser confocal microscopy, survival assays.
3. Gene expressions: RT-PCR, Northern blot, Western blot, reporter assay
4. Expression and isolation of recombinant proteins in bacterial and insect systems
5. Protein-protein interaction assays: immunoprecipitation, pull-down assay
6. Protein-DNA interaction assays: chromatin immunoprecipitation assay, gel mobility shift assay
7. Others: SAS program for statistical analysis, DNA repair assays, DNA methylation assay, protein kinase assay, apoptosis staining

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### PUBLICATIONS

1. **Wang HC**, Chou WC, Shieh SY, Shen CY\*. Ataxia telangiectasia mutated and checkpoint kinase 2 regulate BRCA1 to promote the fidelity of DNA end-joining. *Cancer Res.* 2006;66:1391-400.
2. Genome-wide association study identifies novel breast cancer susceptibility loci. *Nature* 2007;447:1087-93. Easton DF *et al.* (**Wang HC** is listed as the 27th among 105 authors).
3. Hsu HM, **Wang HC**, Chen ST, Hsu GC, Shen CY, Yu JC. Breast cancer risk is associated with the genes encoding the DNA double-strand break repair Mre11/Rad50/Nbs1 complex. *Cancer Epidemiol Biomarkers Prev.* 2007;16:2024-32.
4. Chou WC, **Wang HC**, Wong FH, Ding SL, Wu PE, Shieh SY\*, Shen CY\*. Chk2-dependent phosphorylation of XRCC1 in the DNA damage response promotes base excision repair. *EMBO J.* 2008; 27:3140-1450. (the 5th. *TienTe Lee* Biomedical Foundation Best Thesis Award, 2009.)
5. Udler MS *et al.* (**Wang HC** is listed as the 25th among 35 authors). FGFR2 variants and breast cancer risk: fine-scale mapping using African American studies and analysis of chromatin conformation. *Hum Mol Genet.* 2009; 18:1692-1703.
6. Gaudet MM *et al.* (**Wang HC** is listed as the 96th among 119 authors). Five polymorphisms and breast cancer risk: results from the Breast Cancer Association Consortium. *Cancer Epidemiol Biomarkers Prev.* 2009; 18:1610-1616.
7. Milne R. *et al.* (**Wang HC** is listed as the 86th among 95 authors). Risk of estrogen receptor-positive and -negative breast cancer and single-nucleotide polymorphism 2q35-rs13387042. *J Natl Cancer Inst.* 2009; 101:1012-1018.
8. Lee CL, Huang CH, **Wang HC**, Chuang DW, Wu MJ, Wang SY, Hwang TL, Wu CC, Chen YL, Chang FR\*, Wu YC\*. First total synthesis of antrocamphin A and its analogs as anti-inflammatory and anti-platelet aggregation agents. *Org Biomol Chem.* 2011; 9:70-73.
9. Lai WC, **Wang HC**, Chen GY, Yang JC, Korinek M, Hsieh CJ, Nozaki H, Hayashi KI, Wu CC\*, Wu YC, Chang FR\*. Using the pER8:GUS Reporter System to Screen for Phytoestrogens from *Caesalpinia sappan*. *J Nat Prod.* 2011;74:1698-1706.
10. Chung YM, **Wang HC**, El-Shazly M, Leu YL, Cheng MC, Lee CL, Chang FR\*, Wu YC\*. Antioxidant and Tyrosinase Inhibitory Constituents from the Desugared Sugarcane Extract, a By-Product of Sugar Production. *J Agric Food Chem.* 2011; 59:9219-9225.
11. Hunyadi A, Chuang DW, Danko B, Chiang MY, Lee CL, **Wang HC**, Wu CC, Chang FR\*, Wu YC\*. Direct semi-synthesis of the anticancer lead-drug Protoapigenone from Apigenin, and synthesis of

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- further new cytotoxic protoflavone derivatives. PLoS One 2011; 6:e23922.
12. Liao WT, Huang TS, Chiu CC, Pan JL, Liang SS, Chen BH, Chen SH, Liu PL, **Wang HC**, Wen ZH, Wang HM\*, Hsiao SW\*. Biological properties of acidic cosmetic water from seawater. Int J Mol Sci. 2012;13:5952-5971.
  13. **Wang HC**, Tsai YL, Wu YC, Chang FR, Liu MH, Chen WY, Wu CC\*. Withanolides-induced breast cancer cell death is correlated with their ability to inhibit heat protein 90. PLoS One. 2012;7(5):e37764
  14. **Wang HC\***, Lee AY, Chou WC, Wu CC, Tseng CN, Liu KY, Lin WL, Chang FR, Chuang DW, Hunyadi A, Wu YC\*. Inhibition of ATR-dependent signaling by protoapigenone and its derivative sensitizes cancer cells to interstrand cross-link-generating agents *in vitro* and *in vivo*. Mol Cancer Ther. 2012 11 (7):1443-1453
  15. Danko B, Martins A, Chuang DW, **Wang HC**, Amaral L, Molnár J, Chang FR, Wu YC\*, and Hunyadi A\*. *In vitro* cytotoxic activity of novel protoflavone analogs – selectivity towards a multidrug resistant cancer cell line. Anticancer Res. 2012; 32:2863-2869. {SCI} (IF=1.725 Ranking/ONCOLOGY: 136/196=69.39%)
  16. Chen WC, Wang SY, Chiu CC, Tseng CK, **Wang HC\***, and Lee JC\*. Lucidone suppresses hepatitis C virus replication by Nrf2-mediated heme oxygenase-1 induction. Antimicrobial Agents and Chemotherapy 2013; 57(3):1180-91.
  17. Tseng CN\*, Chang HW, Stocker J, **Wang HC**, Lu CC, Wu CH, Yang JG, Cho CL, Huang HW\*. A Method to Identify RNA A-to-I Editing Targets Using I-Specific Cleavage and Exon Array Analysis. Mol Cell Probes. 2013 Feb;27(1):38-45.
  18. **Wang HC\***, Wu CC, Cheng TS, Kuo CY, Tsai YC, Chiang SY, Wong TS, Wu YC, Chang FR\*. Active constituents from *Liriope platyphylla* root against cancer growth *in vitro*. Evid-Based Compl Alt. 2013; 2013: 857929.
  19. **Wang HC**, Chu FH, Chien SC, Liao JW, Hsieh HW, Li WH, Lin CC, Shaw JF, Kuo YH\*, Wang SY\*. Establishment of metabolite profile for an *Antrodia cinnamomea* health food product and investigation of its chemoprevention activity. J Agric Food Chem. 2013; 61: 8556–8564
  20. **Wang HC**, Tseng YH, Wu HR, Chu FH, Yueh-Hsiung Kuo YH\* and Wang SY\*. Anti-proliferation Effect on Human Breast Cancer Cells *via* Inhibition of pRb Phosphorylation by Taiwanin E Isolated from *Eleutherococcustrifoliatus*. Nat Prod Commun. 2014; 9: 1303-1306.
  21. Kuo CY, **Wang HC**, Kung PH, Lu CY, Liao CY, Wu MT, Wu CC\*. Identification of CalDAG-GEFI as an intracellular target for the vicinal dithiol binding agent phenylarsine oxide in human platelets. Thromb Haemost. 2014; 111:892-901

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22. Ting YC, Ko HH, **Wang HC**, Peng CF, Chang HS, Hsieh PC, Chen IS. Biological evaluation of secondary metabolites from the roots of *Myrica adenophora*. *Phytochemistry*. 2014; 103:89-98.
23. Chang HW, **Wang HC**, Chen CY, Hung TW, Hou MF, Yuan SS, Huang CJ, Tseng CN. 5-azacytidine induces anoikis, inhibits mammosphere formation and reduces metalloproteinase 9 activity in MCF-7 human breast cancer cells. **(co-first author)** *Molecules*. 2014;19:3149-59.
24. Cheng YB, Chien YT, Lee JC, Tseng CK, **Wang HC**, Lo IW, Wu YH, Wang SY, Wu YC\*, Chang FR\*. Limonoids from the seeds of *Swietenia macrophylla* with inhibitory activity against dengue virus 2. *J Nat Prod*. 2014;77:2367-74. {SCI} [IF=3.947 (2013) Rank/PLANT SCIENCES: 21/199 =10.6%]
25. Hou YL, Chang HS, **Wang HC**, Wang SY, Chen TY, Lin CH, Chen IS\*. Sassafrandainol: A new neolignan and anti-inflammatory constituents from the stem of *Sassafras randaiense*. *Nat Prod Res*. 2015;29:827-32. {SCI} [IF=0.919 (2014) Rank/CHEMISTRY, APPLIED: 46/72 =63.9%]
26. Liao CY, Lee CL, **Wang HC**, Liang SS, Kung PH, Wu YC, Chang FR, Wu CC\*. CLL2-1, a chemical derivative of orchid 1,4-phenanthrenequinones, inhibits human platelet aggregation through thiol modification of calcium-diacylglycerol guanine nucleotide exchange factor-I (CalDAG-GEFI). *Free Radic Biol Med*. 2015; 78:101-10.
27. **Wang HC\***, Chang FR, Huang TJ, Kuo CY, Tsai YC, Wu CC\*. (-)-Liriopein B Suppresses Breast Cancer Progression via Inhibition of Multiple Kinases. *Chem Res Toxicol*. 2015;28:897-906.
28. Kuo CY, Chou WC, Wu CC, Wong TS, Kakadiya R, Lee TC, Su TS\*, **Wang HC\***. Repairing of N-mustard derivative BO-1055 induced DNA damage requires multiple DNA repair mechanisms. *Oncotarget* 2015;6:25770-83 {SCI} [IF=6.359 (2014) Rank/ONCOLOGY: 21/211 =10.0%]
29. Stanković T, Dankó B, Martins A, Dragoj M, Stojković S, Isaković A, **Wang HC**, Wu YC, Hunyadi A, Pešić M. Lower antioxidative capacity of multidrug-resistant cancer cells confers collateral sensitivity to protoflavone derivatives. *Cancer Chemother Pharmacol*. 2015;76:555-65.
30. Kuo CY, Zupkó I, Chang FR, Hunyadi A, Wu CC, Weng TS, **Wang HC\***. Dietary flavonoid derivatives enhance chemotherapeutic effect by inhibiting the DNA damage response pathway. *Toxicol Appl Pharmacol*. 2016; 311:99-105. 2016; 311:99-105 {SCI} [IF=3.585 (2018) Rank/TOXICOLOGY: 21/99=22.6%]
31. Huang CY, Chang CW, Tseng YJ, Lee J, Sung PJ, Su JH, Hwang TL, Dai CF, **Wang HC**, Sheu JH\*. Bioactive Steroids from the Formosan Soft Coral *Umbellulifera petasites*. *Mar Drugs*. 2016;14. pii: E180.
32. Chang HS, Lin CH, Chen YS, **Wang HC**, Chan HY, Hsieh SY, Wu HC, Cheng MJ\*, Yuan GF,

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- Lin SY, Lin JY, and Chen IS\*. Secondary metabolites of the endophytic fungus *Lachnum abnorme* from *Ardisia cornudentata*. Int. J. Mol. Sci. 2016;17. pii: E1512.
33. Bózsity N, Minorics R, Szabó J, Mernyák E, Schneider G, Wölfling J, **Wang HC**, Wu CC, Ocsovszki I, Zupkó I\*. Mechanism of antiproliferative action of a new D-secoestrone-triazole derivative in cervical cancer cells and its effect on cancer cell motility. J Steroid Biochem Mol Biol. 2017;165(Pt B):247-257.
34. Chao CH, Wu CY, Huang CY, **Wang HC**, Dai CF, Wu YC, Sheu JH\*. Cubitanoids and Cembranoids from the Soft Coral *Sinularia nanolobata*. Mar Drugs. 2016;14. pii: E150. [IF=3.345 (2015) Rank/ CHEMISTRY, MEDICINAL: 13/59=22.0%]
35. Cheng AN, Fan CC, Lo YK, Kuo CL, **Wang HC**, Lien IH, Lin SY, Chen CH, Jiang SS, Chang IS, Juan HF, Lyu PC, Lee AY\*. Cdc7-Dbf4-mediated phosphorylation of HSP90-S164 stabilizes HSP90-HCLK2-MRN complex to enhance ATR/ATM signaling that overcomes replication stress in cancer. Sci Rep. 2017;7:7024.
36. Cheng YB, Liu FJ, Wang CH, Hwang TL, Tsai YF, Yen CH, **Wang HC**, Tseng YH, Chien CT, Chen YA, Chang FR, Wu YC. Bioactive Triterpenoids from the Leaves and Twigs of *Lithocarpus litseifolius* and *L. corneus*. Planta Med. 2018;84:49-58.
37. Sinka I, Kiss A, Mernyák E, Wölfling J, Schneider G, Ocsovszki I, Kuo CY, **Wang HC**, Zupkó I. Antiproliferative and antimetastatic properties of 3-benzyloxy-16-hydroxymethylene-estradiol analogs against breast cancer cell lines. Eur J Pharm Sci. 2018;123:362-370.
38. Ötvös SB, Vágvolgyi M, Girst G, Kuo CY, **Wang HC**, Fülöp F, Hunyadi A\*. Synthesis of Nontoxic Protoflavone Derivatives through Selective Continuous-Flow Hydrogenation of the Flavonoid B-Ring. Chempluschem. 2018;83:72-76.
39. Fási L, Di Meo F, Kuo CY, Stojkovic Buric S, Martins A, Kúsz N, Béni Z, Dékány M, Balogh GT, Pesic M, **Wang HC**, Trouillas P, Hunyadi A. Antioxidant-inspired drug discovery: antitumor metabolite is formed in situ from a hydroxycinnamic acid derivative upon free radical scavenging. J Med Chem. 2019;62:1657-1668
40. **Wang HC**, Hua HH, Chang FR, Tsai JY, Kuo CY, Wu YC, Wu CC\*. Different effects of 4 $\beta$ -hydroxywithanolide E and withaferin A, two withanolides from Solanaceae plants, on the Akt signaling pathway in human breast cancer cells. Phytomedicine 2019;53:213-222. {SCI} [IF=4.180 (2018) Rank/INTEGRATIVE & COMPLEMENTARY MEDICINE: 1/27=3.70%]
41. Kuo CY, Schelz Z, Tóth B, Vasas A, Ocsovszki I, Chang FR, Hohmann J, Zupkó I, **Wang HC**\*. Investigation of natural phenanthrenes and the antiproliferative potential of juncusol in cervical cancer cell lines. Phytomedicine. 2019; 58:152770. {SCI} [IF=4.180 (2018) Rank/INTEGRATIVE

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& COMPLEMENTARY MEDICINE: 1/27=3.70%)

42. Kuo CY, Weng TS, Kumar KJS, Tseng YH, Tung TW, Wang SY, **Wang HC\***. Ethanol Extracts of Dietary Herb, *Alpinia nantoensis*, Exhibit Anticancer Potential in Human Breast Cancer Cells. *Integr Cancer Ther.* 2019; 18:1534735419866924. {SCI} [IF=2.634 (2018) Rank/INTEGRATIVE & COMPLEMENTARY MEDICINE: 7/27=25.9%)
43. Chou WC, Hsiung CN, Chen WT, Tseng LM, **Wang HC**, Chu HW, Hou MF, Yu JC, Shen CY. A functional variant near XCL1 gene improves breast cancer survival via promoting cancer immunity. *Int J Cancer.* 2020; 146:2182-2193.
44. Latif AD, Jernei T, Podolski-Renić A, Kuo CY, Vágvölgyi M, Girst G, Zupkó I, Develi S, Ulukaya E, **Wang HC**, Pešić M, Csámpai A, Attila Hunyadi A. Protoflavone-Chalcone Hybrids Exhibit Enhanced Antitumor Action through Modulating Redox Balance, Depolarizing the Mitochondrial Membrane, and Inhibiting ATR-Dependent Signaling. *Antioxidants (Basel).* 2020;9:519.