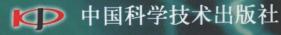
Basic and Clinical Research on Collateral Disease Theory

络病学基础与临床研究(2)

吴以岭 主编



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前言

络病学说是中医学术体系的独特组成部分,是研究络病发病特点、病机变化、临床表现、辨证论治、治疗原则及治法方药的应用理论,络病学是研究中医络病学说及其临床运用的临床学科。络病是广泛存在于多种内伤疑难杂病和外感重症中的病机状态与过程,科学界定络病概念内涵及外延、系统研究络病发生发展及辨证治疗规律、初步建立"络病证治"体系,推动络病学学科建立与发展和对于促进中医学术发展,提高现代多种难治性疾病诊疗水平,具有重要学术价值。

学科是构成科学技术体系的各分支科学,是在一定研究领域形成的专门化知识。学科是 学术的载体,是学术理论发展到一定水平的体现,通过学科的建设与发展又会推动学术理论不 断深入发展与完善。为推动建立络病学学科,国内不少专家学者开始进行络脉与络病研究。 我们坚持络病学说的理论探讨和临床应用研究,并承担了国家中医药管理局课题——络病理 论及其应用研究。该课题研究中提出络病学说理论框架"三维立体网络系统",从时空与功能 的统一性研究络脉理论,系统探讨络病发病、病机、辨证与治疗,使络病学说理论体系初步成 形。运用络病学说指导临床难治性疾病治疗,已完成国家级及省部级课题6项,研制出治疗心 脑血管病、肿瘤、外感热性病的系列自主知识产权创新中药。2004年11月国家中医药管理局 主持了该课题鉴定,与会专家认为:"该项研究按照中医学术自身发展规律对络病学说进行了 全面系统研究,初步建立'络病证治'体系,首次形成系统络病理论,为络病学学科建立奠定理 论基础,属国内外创新性科研成果。"该项研究获得2005年中华中医药学会和河北省科技进步 一等奖,已通过2006年国家科技进步二等奖答辩。在课题鉴定的基础上,我们组织国内知名 专家共同编写出版了系统论述络病理论及其临床应用的专著——《络病学》,并获得 2005 年中 华中医药学会学术著作一等奖。将《络病学》专著改编成"新世纪全国高等中医药院校创新教 材",2006年8月召开了"首届全国《络病学》教学研讨会",全国有20余所高等医药院校开设面 向本科生、硕士生和博士生的络病学课程,这为学科的建设与发展奠定了理论基础。

学术队伍建设是学科建设的重要内容。在络病理论研究过程中,形成了包括中医、西医、中西医结合、生物学、物理学等多学科专家在内的专家群体,建立了国家二级学会——中华中医药学会络病分会。络病分会成立两年来,充分发挥多学科专家群体优势,积极开展多种形式的学术交流,组织申报国家级及省部级重点课题,申办国家级继续教育项目,不仅推动了络病学学科自身发展,而且吸引更多的专家学者投入到络病理论研究。特别是随着络病学教学工作在全国高等医药院校逐步开展,络病学教育将与本科生、硕士生、博士生等高层次中医药人才培养相结合,对推动络病研究领域人才队伍建设具有重要意义。

学科建设的首要任务是明确学科发展方向,随着络病理论体系构建完成,明确其学科发展方向对于络病学学科建设具有极为重要的意义。广义的络脉包括气络与脉络两大网络系统, 气络与脉络病变成为络病学两大学科分支。对这两个学科分支及其相互影响的深入研究对提 高难治性疾病的辨证治疗水平具有重要学术价值,也将促进络病学学科的发展。在既往研究中,基于气络与神经内分泌免疫调节功能、脉络与中小血管、微血管及微循环的相关性,提出了"气络-NEI 网络"和"脉络-血管系统"概念,为进一步深入开展这两大学科分支研究启迪了新的思路。

将"络病证治"用于脉络病研究,与心脑血管病及糖尿病微血管并发症等重大疾病研究进展相结合,以络病学术创新带动络病治疗新药研发,研制出治疗心脑血管病的中药新药通心络胶囊、抗心律失常的参松养心胶囊、治疗慢性心衰的芪苈强心胶囊,形成络病学说指导心脑血管病治疗的系列药物。

络病理论代表方通心络胶囊研究以络病学说探讨心脑血管病的中医病理机制,为心脑血管病的中医治疗开辟了新的有效治疗途径,获得2000年国家科技进步二等奖。近年发表的通心络胶囊的基础与临床研究文章达700余篇,包括数十篇博士、硕士学位论文。众多研究结果显示了通心络胶囊对血管病变不同病理环节的确切防治作用:可明显改善血管内皮功能,促进人体外周血内皮祖细胞增殖、迁移和粘附;延缓动脉粥样硬化进程,抑制内膜增殖及再狭窄,稳定易损斑块;缓解血管痉挛;明显保护缺血再灌注后微血管结构和功能完整性,通过抑制细胞粘附分子的表达,减少中性粒细胞浸润和炎症反应,保护内皮细胞屏障功能完整性,减少缺血区无复流区面积,抑制心室重构,缩小心肌梗死面积,改善心功能;提高急性心肌梗死患者介入治疗后室壁运动异常节段恢复率,改善室壁节段运动指数恢复时间。采用 Meta 分析方法对国内外发表的通心络胶囊治疗心血管病的文献研究结果表明,通心络胶囊治疗心血管病具有确切疗效。

应用络病理论指导脑血管病治疗研究,提出"血管保护——脑梗死治疗的新靶点",以通心络胶囊为代表方进行的实验证实对其缺血区微血管完整性具有明显保护作用,可降低缺血区脑血管通透性,减轻脑水肿,明显增加缺血区微血管数目,并可促进缺血区治疗性毛细血管新生,使缺血区血流代偿性增加,保护缺血区脑组织;减少脑缺血自由基及兴奋性氨基酸释放,降低缺血区神经元胞浆钙离子浓度,抑制脑缺血诱导的神经元凋亡,促进脑缺血神经干细胞增殖分化,进而改善神经功能缺损。Meta分析结果表明通心络能明显提高脑血管病临床疗效,对急性脑梗死患者具有促进神经功能恢复的作用。

络病理论与通心络胶囊研究引起国外医学界关注。论文参加 2005 年欧洲与北美心脏病年会,受到国外专家好评。美国德州大学心脏病研究中心、哈佛大学医学院心脏病研究中心正在开展通心络胶囊研究,部分研究已取得显著成果。

由于运用络病理论指导血管病变治疗取得的显著成效,特别是络病理论代表方通心络胶囊对血管病变不同病理环节的确切作用,2005年由河北以岭医药研究院联合国内数所著名医学院校申报的"络病理论指导血管病变防治的基础研究"列入国家973计划项目。该研究中基于"脉络—血管系统"解剖形态同一性,提出"脉络—血管系统病"新概念。通过3000余例临床病例调查并在此基础上制定"脉络—血管系统病"诊断标准和病证结合的动物模型,逐步阐明通络方药对"脉络—血管系统病"共性病理环节及急性心肌梗死、脑梗死、糖尿病微血管病变的作用机制,最终建立对心脑血管病、糖尿病微血管病变具有普遍指导意义的"脉络—血管系统

病变学说",提高这类重大疾病的防治水平,项目研究的开展将会进一步推动络病学学科的建设。

同时我们积极开展与国外医学界的学术交流与合作。美国东卡大学医学院、北京阜外心血管病医院有关专家就络病理论及通心络胶囊科研进行了学术研讨。随着通心络胶囊作为药品在韩国注册上市,络病理论也引起韩国医学界关注。2005年中韩学术界多次举行络病学的研讨与交流,2006年9月初双方共同在韩国举行"首届中韩国际络病学研讨会"。2005年10月我们在北京召开了"首届国际络病学大会",参会的美国、韩国及国内专家500余人,反映了络病学近年蓬勃发展之势!2006年10月,将在上海召开"第二届国际络病学大会",预计参会的国内外专家达600余人,收到参会论文160余篇,内容涉及理论探讨、实验研究、临床总结等,充分显示出络病理论的学术价值。所有参会论文将被汇编成《络病学基础与临床研究》(第二辑)。值此会议召开之际,我们向关心与支持络病分会工作的国家中医药管理局、中华中医药学会领导,向承办本次会议的上海中医药学会、上海中西医结合学会、第二军医大学长征医院表示衷心感谢!向多年来一直支持络病理论研究的医学界前辈邓铁涛教授、颜德馨教授、路志正教授、焦树德教授、朱良春教授、陈灏珠院士,向亲自审阅论文的王永炎院士表示衷心的感谢!向所有致力于络病学研究的国内外专家学者表示衷心的感谢!愿我们携手并进,合力推动络病学学科的建设与发展!

2006年10月

PREFACE

The theory of collateral disease is a unique component of traditional Chinese medicine academic theoretical system. It is an applied theory for studying its characteristic when it is occurring, variety of Pathological mechanism, clinical behaviour, dialectical treatment, cure principle and recipes. The doctrine of Collateral disease is a clinical subject which researches collateral diseases of traditional Chinese medicine academic theory and its clinical subject. Collateral disease is pathological condition and process which has extensively existed in all kinds of refractory and spiny diseases. scientifically defining the conception in both extension and intension of Collateral diseases, systematically researching the law of its development and dialectical treating, constituting the dialectical system of collateral disease treatment, promoting the establishment and development of collateral disease doctrine have important academic significance in the development of academic level in TCM and in the improvement for the treating level of various modern spiny diseases.

The doctrine is a branch which constitutes the science system and is special knowledge that has formed in a certain field. The subject is the carrier of academy , and is also the reflection when science develops to a certain level. By developing the subject it makes the development and academic perfect. In order to construct the Collateral disease subject, a lots of specialists and experts start to do the research in LuoMai and collateral diseases. we insist on the theory discussion and clinical applied research. We also take in hand the subject from our national administration bureau for TCM - theory of collateral disease. This question for discussion put forward the theory frame of "Three - dimensional solid net system". This system discusses disease occurring , Pathological mechanism, dialectical treatment, makes the theoretical system of collateral disease into shape from the unity of space - time and function point. with the theory of collateral disease to treat various spiny diseases we have achieved 6 national and provincial research subjects and developed new Chinese traditional medicine which is a series of independent intellectual property creation in curing cerebral and vascular diseases , tumor and various diseases due to external heat. State Administration bureau of Traditional Chinese Medicine presided the evaluation for this subject in November, 2004. The attendance deemed that the study has done a systematic and across - the aboard research for theory of collateral diseases, established the system of dialectical Collateral Disease treating', so it has laid the theoretical foundation for collateral disease . Its innovative production is initial both at home and abroad. This research won the first class award of China Traditional Chinese Medicine Academy and Hebei province science and technology progress prize. It has also got through the defence for a second class award of national science and technology progress prize. On the basis of the evaluation,

We organized some well – known experts to issue a book – (theory of collateral disease) which refers to theory of collateral diseases and clinical applications, and won the first class award of Chinese Academy of Traditional Chinese Medicine in 2005. We tend to change (theory of collateral disease) into a book which is for college students as a text book of TCM.

More than 20 medical colleges hold this textbook for undergraduates, graduate students, and doctoral students. This paves the theoretical basis for the development and construction of the subject.

The academic group is vital to complete the construction of the subject. In the researching process of the theory of collateral disease, we have enrolled many professors and experts on traditional Chinese medicine, Western medicine, Combination of Traditional Chinese and Western Medicine, biology, physics, and so on, and build second class association in our country – collateral disease committee of Chinese Traditional Chinese Medicine Academy. For the past two years, we make use of the advantages of various experts, actively carry out all kinds of academic communications, apply for important topic research projects in the country and province, bid for national continuous education project, all of which not only promote the development of collateral disease, but also attract more experts researching in it. Especially with the development in collateral disease teaching in each school, Collateral disease education will be carried out in undergraduates, postgraduates, doctoral students and some other higher rank medical personnels of TCM, which has important sig-

nificance in the promotion of the building of talents and medical pesonnels of TCM. The first task of subject construction is having clear direction which plays an important role in the construction of collateral disease doctrine. general Collateral disease is composed of qi collaterals and channel collaterals which has become the two branches. The research on the two branches have important value in improving the treatment of extremely difficult cases and the dialectical curing level as well as the development of the subject. Based on the research on the relation among nerve incretion immunity regulation function, venation small vessels, micrangium and microcirculation, the notion of "qi collaterals – NEI network "and" mailuo – blood vascular system " was put forward, opening a new idea in deep researching for the two branches.

Linking Collateral Disease dialectical treating and some severe disease curing research such as cardio – cerebral vascular diseases or diabetes mellitus, and the Collateral Disease academic innovation advanced new medicine, so we have developed a series of new medicines such as Tong Xin Luo Capsule, anti – arrhythmia cordis – – Can Song Yang Xin Capsule, OinLiOiangXin Capsule for curing chronic heart failure and so on.

Under the guidance of collateral disease theory, in the research of Pathological mechanism of cardio - cerebral vascular diseases Tong Xin Luo capsule was successfully made, the Tong Xin Luo Capsule has pioneered an effective way for traditional Chinese medicine curing of cardio - cerebral vascular diseases and was awarded the second class National Science and Technology Progress Prize in 2000. There are about 700 articles which are about the Tong Xin Luo Capsule foundation and clinical research, including tens of thesis by doctoral and master degree graduates. Many research results reveal that this medicine has prevention and cure function to Vascular Lesion in different phases: improving Vascular endothelium function obviously and make peri - endothelia cells proliferate, transplant and adhere; delaying atherosclerosis process and inhibiting endometrial hyperplasia, stabilizing damageable plaque; relieving vascular spasm; conspicuously protecting micrangium intactness at late period of reperfusion, inhibiting expression of adhesive molecule to decrease soakage and inflammation of neutral cell, protecting shielding function of endothelium cell integrity, reducing ischemic area, restraining ventricular restructuring, shrinking myocardial infarction area, improving heart function, enhancing myocardial function for infarction sufferers who have ventricular niche movement abnormity section renew rata after medium treating, improving ameliorate niche section movement index healing time. The references which are about Tong Xin Luo Capsule curing Vascular Lesion by Meta analysis method reveal that Tong Xin Luo Capsules have evident effect. By using the research of theory of collateral disease to instruct treating Vascular Lesion , we bring forward "protect blood vessel - the new therapeutics". The experiments by using Tong Xin Luo Capsules confirm that this method protects micrangium at ischemic area, reduces micrangium at ischemic area penetrating, alleviates encephaledema, increases the number of micrangium at ischemic area, also can rebirth capillary vessels and enhance blood stream reimbursive in ischemic area., decrease encephalon ischemic free radical and release excitability aminophenol, reduce the thickness of calcium hydronium in nerve fiber in ischemic area, depress nerve fiber decease lured by ischemia, promote the proliferation of neuron stem cells improve the function of the nervous system. Meta analysis reveals that Tong Xin Luo capsules can effectively increase clinical treatment result, promote the restoration of the function in the nervous system for patients with acute cerebral infarction.

The research of theory of collateral disease and Tong Xin Luo Capsules have aroused attention at home and abroad. Overseas experts gave the paper highly praise in Europe and North America Cardiopathy Conference. Heart Disease Research Center at university of Texas, Harvard Heart Disease Research Center are ResearchingTong Xin Luo Capsule and some have had achievement.

Because of evident effect using the theory of collateral disease in treating Vascular Lesion, especially the representative recipes - - Tong Xin Luo Capsules have evident effect, "theory of collateral instructing disease vascular lesion" which was referred by HeBei YiLing Medicinal Academy with a few famous Academy of Medicinal Science has been listed in "national 973 programs". This research was based on identity of Collateral - vascular system, and "Collateral - vascular system illness as a new conception was put forward. The diagnosis criterion and animal model of "Collateral - vascular system illness", which was summed up by more than 3000 clinical cases, illuminate that drugs removing obstruction in

the channels have effect to acute cerebral infarction, diabetes mellitus micrangium Lesion, and at last come into being the theory of Collateral – vascular system illness which have common direction signification to above diseases. This work has improved prevention and cure level to those diseases, and it will make the subject perfect.

At the same time, we communicate and cooperate with overseas medicine circle. The experts of east carolina university in America and cardiovascular institution of FuWai hospital in BeiJing have done some research, held celebrated seminars about Collateral Disease and Tong Xin Luo Capsule. Theory of collateral disease has aroused attention in Korea along with Tong Xin Luo Capsule came into Korea market.

China – Korea Academia held conference talking about theory of collateral disease many times in 2005. when we held "the first international collateral disease conference" in BeiJing. More than 500 specialists who come from America Korea and China pointed out that theory of collateral disease would be more and more flourishing. There will be 600 experts in the seconds "international collateral disease conference". We have received more than 600 papers which include theory discussion, experiment study and clinical summing – up. Achievements all above show the academic value in theoretical system. All the paper will be compiled to clinical research to collateral diseases (the secondary edition). We should be thankful to State Administration bureau of Traditional Chinese Medicine, Chinese Academy of Combination of Traditional Chinese Medicine with western medicine, ShangHai Academy of Combination of Traditional Chinese Medicine of the integration of Traditional Chinese and Western Medicine, Second Military Medical University. We should express our appreciation to those noble – minded predecessors in medicine field. They are professors Tietao Deng, De. xin Yan, Zhizeng Lu, Shude Jiao Liangchun Zhu, and academician ZC. We should be thankful to Yongyan Wang who had checked and approved our paper. Finally we should express our appreciation to experts and scholars and those who dedicate themselves to theory of collateral disease. In a word we should hold together to accelerate construction and progress of the subject.

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