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The Effect of *T’ai Chi* Exercise on Immunity and Infections: A Systematic Review of Controlled Trials

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Abstract

**Purpose:** The aim of this review is to summarize and assess critically clinical trial evidence of the effect of *t’ai chi* (TC) exercise on immunity and TC efficacy for treating infectious diseases.

**Methods:** Fourteen databases were searched from their respective inceptions through January 2011. No language restrictions were imposed. Quality and validity of the included clinical trials were evaluated using standard scales.

**Results:** Sixteen (16) studies, including 7 randomized controlled trials, 4 controlled clinical trials, and 5 retrospective case-control studies, met the inclusion criteria for this review. One (1) study examined clinical symptoms, 3 studies tested functional measures of immunity (antigen-induced immunity), and the other studies tested enumerative parameters of immunity, such as lymphocytes, immunoglobulins, complements, natural-killer cells, and myeloid dendritic cells. Overall, these studies suggested favorable effects of TC exercise.

**Conclusions:** TC exercise appears to improve both cell-mediated immunity and antibody response in immune system, but it remains debatable whether or not the changes in immune parameters are sufficient to provide protection from infections.

Introduction

*T’ai Chi* (TC), a complementary and alternative modality of Traditional Chinese Medicine, combines characteristics of physical exercise and meditative practice. TC is popularly practiced by a large number of people in Chinese communities to improve physical fitness and overall well-being. The gentle movements and postures of the exercise coordinated with breathing patterns and meditation are designed to achieve a harmonious flow of energy (*qi*) in the body. TC exercise is equivalent to moderate-intensity aerobic exercise. However, TC is not only an exercise but also a mind-body intervention. Its beneficial effects on health have been documented in increasing number of studies.1–3 It is hypothesized that TC as a modality of mind-body intervention with a moderate intensity of physical exercise may improve immune functions of human body.4,5 However, few reviews have examined the scientific evidence of the effect of TC on immunity. It is well-known that infections, such as influenza and herpes zoster, are associated with human body’s immunity, and that individuals who have substantial declines in immune function are at increased risk for contracting a number of infectious diseases. Epidemiologic studies have suggested that moderate exercise training is associated with reduction in the incidence of upper respiratory tract infection (URTI), whereas endurance athletes are at increased risk for URTIs during periods of heavy training.5,6 It is still unclear whether or not TC exercise may reduce the incidence or the severity of infectious diseases. Thus, this systematic review aims to summarize and evaluate critically clinical trial evidence of the effectiveness of TC exercise for improving immune functions and its efficacy for treating infectious diseases.

**Methods**

**Data sources**

The following electronic databases were searched from their respective inceptions through January 2011: PubMed/MEDLINE; CENTRAL; CINAHL; EMBASE; AMED; Qigong and Energy Medicine Database; SPORTDiscus Database; China Journals Full-text Database-Medicine/Hygiene Series, China Proceedings of Conference Full-text Database, Chinese Master Theses Full-text Database, China Doctor Dissertations.
Full-text Database, Electronic Theses and Dissertation System (Taiwan), Taiwan Electronic Periodical Services, and Index to Taiwan Periodical Literature System. The search terms used for this review included: tai chi, taichi, tai ji, taiji, tajiquan, shadowboxing, influenza, infection, infectious, inflammation, inflammatory, immune, immunity, immunological, lymphocyte, and antibody. Both traditional and simplified Chinese translations of these terms were used in Chinese databases. Reference lists of all included studies, existing reviews, and other archives of the located publications were hand-searched for further relevant articles.

**Study selection**

All controlled clinical trials (CCTs) were included if they examined the effects of TC exercise on the parameters of immunity or for treatment of various infectious diseases. Given the limited number of prospective clinical trials in the field, retrospective case-control studies (RCSs) were also included to provide alternative evidence, but uncontrolled observational studies were excluded, because of their susceptibility to bias and lack of significant evidence. Case reports and qualitative studies were also excluded for lack of significant evidence. To assess the effect of TC on immunity, studies on any subject were included, but studies among athletes were excluded because of their high intensity of physical exercise. To evaluate the effect of TC on infections, any study about TC concerning the incidence or severity of infectious diseases were included. For all included studies, primary data from the original sources were reviewed and analyzed.

To assess the effects of TC exercise on improvement of immunity and TC’s effectiveness for treating infectious diseases, such outcome measures as physical symptoms relevant to infections and biomedical indicators of immunity were considered. Generally, an individual’s immune status can be assessed using either enumerative or functional measures. Functional measures assess how well a specific immune process works (e.g., how effectively natural-killer [NK] cells destroy laboratory-grown tumor cells or how much lymphocytes divide following stimulation with a mitogen [a substance that induces mitosis or cell division]). In clinical practice, one of the most valid and commonly used functional measures of immunity is used to assess people’s immune responses to antigens that people are highly sensitive to. This technique involves placing a small piece of antigen directly underneath the skin, a procedure that causes a local inflammatory response consisting of induration (a swollen round bump) and erythema (redness around the bump), and measuring the magnitude of this response immediately or 24–48 hours later, depending on the antigen that is used. Enumerative measures involve counting different immune-system components (or biomarkers), including white blood–cell populations (granulocytes, monocytes, lymphocytes, NK cells, B-lymphocytes, T-lymphocytes, helper T-lymphocytes, and suppressor/cytotoxic T-lymphocytes), antibody populations in the blood (immunoglobulin [Ig]A, IgG, and IgM) and in saliva (secretory IgA), and antibodies to specific pathogens. The current review focused on the number and the percentage of white blood cells, mainly T-lymphocytes, and levels of serum IgG and complements in peripheral blood, because these biomarkers are commonly used parameters in clinical practice and in the field of exercise research. Psychosocial outcomes, such as quality of life and psychologic well-being, were not considered because it is difficult to attribute effects on such outcomes to the change of immunity.

**Data extraction and assessment**

For each included study, data were extracted by 1 main researcher and then verified by another researcher. All discrepancies were resolved by discussion. The strength of the evidence was evaluated for all the included studies using the Oxford Centre for Evidence-based Medicine Levels of Evidence.8 These criteria are applied to grade the methodological rigor of studies from level 1 or grade A (systematic review of RCTs, 1a; individual RCT with narrow confidence interval, 1b) to level 5 or grade D (expert opinion). The quality and validity of the included RCTs were also evaluated using the Jadad scale,9 which is based on three criteria: (1) description of randomization and allocation concealment; (2) double-blinding; and (3) withdrawals or dropouts (the score ranges from 0 to 5). This is a standard scale used in systematic reviews of RCTs. Given that it was difficult to blind patients to TC, only assessor blinding was evaluated. The risk of bias in the included trials was assessed using the framework for methodological quality recommended by Juni et al.10 According to this framework, biases fall into four categories: (1) selection bias (biased allocation to comparison groups); (2) performance bias (unequal provision of care apart from intervention under evaluation); (3) detection bias (biased assessment of outcomes); and (4) and attrition bias (biased occurrence of loss to follow-up).

**Results**

**Study description**

The database searches identified 87 potentially relevant articles (Fig. 1). Of them, 51 articles were excluded because they were not clinical trials or not related to infection or immunity. Full reports of 36 studies were acquired, and 20 were also excluded because they were uncontrolled observational studies (10), studies with unparallel controls (2), duplicate publications (3), studies using athletes (2), and studies with other outcome measures (3). Sixteen studies published between 1988 and 2010, including 7 RCTs11–17 4 CCTs,18–21 and 5 RCSs,22–26 met this review’s inclusion criteria. These studies were conducted in the United States,11,14,16,19 Taiwan,22 and mainland China.12,13,15,17,18,20,21,23,26 Seven (7) studies13,14,16,19,22,24,25 were published in English, 8 studies12,13,15,17,18,21,23,26 were published in Chinese, and the remaining 1 study20 was a proceeding.

Of the 16 included studies, 4 used samples of young college students11,12,18,20 1 study used a sample of persons with HIV infection,16 and the other studies used samples of middle-age or older healthy adults. Four (4) studies11,12,17,18 focused solely on females and 2 studies20,22 focused solely on males. Sample sizes in the included studies ranged from 16 to 252. A sample shared by 2 studies11,12 was considered to be one sample. In total, these studies covered 939 subjects, including 577 subjects in the TC exercise groups and 362 subjects in the control groups. Characteristics of the included RCTs and non-RCTs (CCTs and RCSs) are presented in Tables 1 and 2, respectively.
Effect of TC intervention

One CCT\textsuperscript{18} on clinical symptoms suggested a favorable effect of TC exercise. Three (3) studies, including 2 RCTs\textsuperscript{13,14} and 1 CCT,\textsuperscript{14} examined the effects of TC on antigen-induced, virus-specific cell-mediated immunity and antibody response in the human immune system. The results of these studies indicated that TC exercise could augment immune responses to virus and influenza vaccines.

Six (6) studies, including 2 RCTs,\textsuperscript{15,16} 1 CCT,\textsuperscript{21} and 3 RCSs,\textsuperscript{23,25,26} examined number and/or percentage of T-lymphocytes. One (1) RCT\textsuperscript{15} on older adults suggested that the number of CD4\textsuperscript{+} and the ratio of CD4\textsuperscript{+}/CD8\textsuperscript{+} increased significantly after 8 weeks of TC practice. Another RCT\textsuperscript{16} on persons with HIV suggested that lymphocyte proliferation function was augmented significantly at the 6-month follow-up visit after 10 weeks of TC exercise. One (1) CCT\textsuperscript{21} and 3 RCSs,\textsuperscript{23,25,26} suggested that the number of lymphocytes, mainly CD4\textsuperscript{+}, and the ratio of CD4\textsuperscript{+}/CD8\textsuperscript{+} were significantly higher in TC groups, compared to control groups. One (1) RCS\textsuperscript{24} examined the number and percentage of B-lymphocytes and suggested that the number of ZC-rosette-forming cells (B-lymphocytes) was lower in a TC group at resting status but increased significantly after 20 minutes of exercise, compared to controls.

One (1) RCT\textsuperscript{22} indicated that number of NK cells increased significantly after 6 months of TC practice, but 1 RCS\textsuperscript{23} suggested that the percentage of NK cells decreased significantly in a TC group after 25 minutes of exercise. One RCS\textsuperscript{25} examined the effect of TC on circulating myeloid dendritic cells (the potent antigen-presenting cells linking innate and adoptive immunity) and indicated a favorable effect of TC exercise.

Five (5) studies, including 2 RCTs and 3 CCTs, examined the concentration of Igs in peripheral blood. One (1) RCT\textsuperscript{15} and 3 CCTs\textsuperscript{16,20,21} suggested favorable effects of TC exercise on IgG and IgA, while another RCT\textsuperscript{12} only suggested a favorable effect on IgG. One (1) RCT\textsuperscript{11} examined concentrations of complements (C3, C4), and the results indicated that concentrations of C3 and C4 increased significantly after 12 weeks of TC exercise. One (1) CCT\textsuperscript{20} also suggested a favorable effect of TC exercise on C3.

Study quality

Jadad scores for the included RCTs ranged from 1 to 4, with a value of 3 or above only for 2 studies. Levels of evidence for the included studies were ranked as "A" for 3 studies, "B" for 12 studies, and "C" for 1 study.

Discussion

This review aimed to assess the efficacy and the effectiveness of TC exercise for treating infectious diseases. Following a comprehensive search of existing literature, it was found that clinical trials of TC in patients with infectious diseases were very limited. Only one study\textsuperscript{18} examined clinical symptoms relevant to infections among female students. The results indicated that the durations of URTIs shortened significantly after 6 months of TC exercise. Instead, many studies in the field examined the effectiveness of TC exercise on improvement of human immune function. Overall, this review demonstrated that the available evidence suggested favorable effects of TC exercise both for increasing effective components of the immune system and for improving immune function, as indicated by functional measures of immunity.

The risk of bias for the studies examined in this review was assessed, based on the descriptions of randomization, allocation concealment, blinding, and withdrawals.\textsuperscript{10} A high risk of bias might have existed in some of the included trials, which might have led to false-positive results. Of the 7 included RCTs, only 3\textsuperscript{13,14,16} described method of randomization and allocation concealment; 2 RCTs\textsuperscript{13,16} adopted assessor blinding, and 4 RCTs\textsuperscript{13–16} reported details about dropouts and withdrawals and adopted intention-to-treat
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<td>Wait-list</td>
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<td>1 B (3b)</td>
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<td>Huang et al., 2006&lt;sup&gt;12&lt;/sup&gt;</td>
<td>Female students from a nursing school (NR)</td>
<td>TG: 10; TG: 10; CG: 10</td>
<td>TC exercise (style: NR): TG: 1 time per week; TG: 2 time per week; TG: 3 time per week (45 minutes each time)</td>
<td>Wait-list</td>
<td>12 weeks</td>
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<td>Irwin et al., 2007&lt;sup&gt;13&lt;/sup&gt;</td>
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<td>3 A (1b)</td>
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<td>Liu, 2006&lt;sup&gt;15&lt;/sup&gt;</td>
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<td>TG: 10; CG: 10</td>
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<td>8 weeks</td>
<td>1. T-lymphocytes 2. Serum IgG, IgM, &amp; IgA</td>
<td>VZV-specific CMI increased 50% from baseline to 1-week postintervention in TCC group ($p &lt; 0.05$) but unchanged in CG. 1. Significant increase in expression of CD4+ ($p &lt; 0.05$) &amp; CD4+ /CD8+ ratio ($p &lt; 0.01$) &amp; decrease in expression of CD8+ ($p &lt; 0.05$) in TG at postintervention; no significant change in T-lymphocytes in CG. 2. Concentrations of IgG &amp; IgA increased significantly ($p &lt; 0.05$) in TG at postintervention; no significant change for these variables in CG</td>
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RCTs, randomized controlled trials; NR, not reported; TC, t’ai chi; TG, t’ai chi group; CG, control group; Ig, immunoglobulin; TCC, T’ai Chi Chih; VZV, varicella zoster virus; CMI, cell-mediated immunity; HIV, human immunodeficiency virus; NKC, natural-killer cells, NK, natural killer; IL, interleukin.
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<th>Studies author, year &amp; reference</th>
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<td>1. Duration of URTI became shorter significantly at postintervention in participants in TG 1 &amp; TG 2, compared to CG (p&lt;0.05) 2. Levels of IgA and IgG increased significantly at postintervention, compared to CG (p&lt;0.05)</td>
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<td>Yang et al., 2008&lt;sup&gt;59&lt;/sup&gt;</td>
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<td>Compared with baseline measures, all measures in TG group increased significantly; in qigong group, only levels of serum complement C3 &amp; saliva lysozyme increased significantly</td>
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<td>Liu and Zhang, 2002&lt;sup&gt;23&lt;/sup&gt;</td>
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<td>Sun et al., 1990&lt;sup&gt;24&lt;/sup&gt;</td>
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<td>B (3b)</td>
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<tr>
<td>Sun et al., 1989&lt;sup&gt;25&lt;/sup&gt;</td>
<td>Healthy aging subjects (NR)</td>
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<td>88-style t'ai chi chuan Practice for 2–10 years</td>
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<td>Total number of T-lymphocytes &amp; number of active T-lymphocytes increased significantly in TG, compared to CG controls (p&lt;0.01)</td>
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<td>Zhu &amp; Sun, 1998&lt;sup&gt;26&lt;/sup&gt;</td>
<td>Middle-age &amp; older healthy adults (NR)</td>
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<td>88-style t'ai chi chuan Practice for average of 7 years</td>
<td>Routine activities</td>
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<td>1. The numbers and the percentages of LC and E-RFC were higher in TG at resting status, compared to CG (p&lt;0.01) 2. After 20 minutes of exercise, number &amp; percentage of E-RFC &amp; Y-RFC increased significantly in TG, compared to CG (p&lt;0.01)</td>
<td>B (3b)</td>
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CCTs, controlled clinical trials; RCSs, retrospective case-control studies; TG, t'ai chi group; CG, control group; URTI, upper–respiratory tract infection; Ig, immunoglobulin; TC, t'ai chi; NR, not reported; ZC-RFL, ZC-rosette-forming lymphocytes; E-RFL, E-rosette-forming lymphocytes; WBC, white blood cells; LC, lymphocytes; E-RFC, E-rossette-forming cells; Y-RFC, Y-rossette-forming cells; NK, natural killer; NKC, natural-killer cell.
sympathetic–adrenal medullary (SAM) axis. These factors induce changes in cellular trafficking, lymphocyte proliferation, and antibody production. As a consequence of muscular contraction and catecholamine-induced immediate changes, for instance, cellular components of the immune system may be mobilized to the blood. Moreover, TC exercise may lead to an increased oxygen supply and alterations in metabolism and metabolic factors, such as plasma glutamine and plasma glucose, which also contribute to exercise-associated changes in immune function. An additional possibility is that immune enhancement is mediated, in part, by improvements of psychosocial factors that are promoted by TC as a mind–body intervention. It has been suggested that psychologic stress can affect immunity through the hypothalamic–pituitary–adrenal (HPA) axis. Stress-induced activation of the HPA axis results in the release of neuroendocrine hormones, such as adrenocorticotropin, from the anterior pituitary gland. Adrenocorticotropin then circulates through the bloodstream to the adrenal glands, where the hormone induces release of glucocorticoids, which can bind receptors at the cell surfaces of lymphoid and myeloid cells. TC as a form of mind–body intervention may buffer the effects of stress on plasma glucocorticoids and, thus, induce alterations in immune function.

It should be noted that the immune system is a complex system, and both functional and enumerative measures of immunity provide rough estimates of specific processes rather than global indications of the immune system's capacity to resist infectious disease. First, the normal functioning range is very broad for most immune measures, and it is still unclear whether or not the magnitude of changes induced by exercise are sufficient to move outside of the normal ranges. Even if these changes were sufficient, it is not clear whether or not the alterations would persist for a sufficient duration of time to alter risk for infectious disease. Second, most of the included studies were conducted using healthy adults, and the clinical implications of the changes in these parameters in healthy people are unclear. Changes in cell number may just reflect changes in the dynamics of lymphocyte migration and recirculation, or shifts in plasma volume, rather than absolute changes in total cell numbers. In addition, absolute changes in cell number will not necessarily result in a significant change in the capacity of the immune system to make an effective response to antigenic challenge. Thus, it would be inappropriate to conclude that TC-induced changes in any specific immune parameter signal a state of “immune enhancement” or resistance to infections.

This review may have had some limitations. Similar to any systematic review, one major limitation was the potential incompleteness of the evidence reviewed. The aim was to identify all controlled trials in this area in a large number of databases with no restrictions on publication language. The current authors are confident that the search strategy used for this review had located all relevant data; however, a degree of uncertainty remains. Moreover, selective publishing and reporting can be major causes of bias in the included studies. In addition, it was not possible to perform meta-analyses because of the heterogeneity of study designs and outcome measures in the included studies.

Conclusions

The available evidence suggest that TC exercise may improve both cell-mediated immunity and antibody response in immune system, but it remains debatable whether or not the exercise-induced alterations in immune function are sufficient to alter human body defense, disease susceptibility, and severity. Because of methodological flaws in existing studies, further vigorously designed large-scale placebo-controlled, randomized trials are needed. Future studies should also test the efficacy of TC exercise for reducing the incidence or the severity of infectious disease.

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