Comparison of trends in the quantity and variety of Science Citation Index (SCI) literature on human pathogens between China and the United States

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Abstract The proportion of pathogenic microorganisms in the microbial world is relatively small, while their threat to human health, economic development and social stability is severe. The quantity and variation of Science Citation Index (SCI) literature related to pathogenic microorganisms may reflect the level of relevant research and the degree of attention. Here we compared trends in the quantity and variety of SCI literature relating to certain important pathogenic microorganisms published by scientists from United States and China from 1996 to 2010 by searching the Science Citation Index database. The pathogenic microorganisms in this study comprise two categories of pathogens: Bacillus anthracis, Yersinia pestis, Francisella tularensis, Ebola virus, Burkholderia pseudomallei, which belong to biodefense-associated pathogens (BDAPs) and the human immunodeficiency virus (HIV), SARS coronavirus, hepatitis B virus (HBV), Mycobacterium tuberculosis, influenza virus, which belong to the commonly encountered health-threatening pathogens. Our results showed that the United States (US) published much more SCI literature on these pathogens than China. Furthermore, literature on BDAPs published by scientists from the US has increased sharply since 2002. However, the numbers of literature relating to CEHTPs from China has demonstrated a gradual increase from 1996 to 2010. Research into pathogenic microorganisms requires three balance to be achieved: investment in BDAP and CEHTP studies; basic and applied research; a faster pace of research into pathogens and fulfilling biosafety and biosecurity requirements.

Keywords Pathogenic microorganism · SCI literature · United States · China · Variety · Quantity

Introduction

The proportion of pathogenic microorganisms in the microbial world is relatively small, while its threat to human health, economic development and social stability is severe.

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Infectious diseases are newly emergent or a recurrence of known infectious diseases, while some persistent infectious diseases are resistant to current techniques available for their control. Each year, approximately 15 million people die from infectious diseases around the world and infectious diseases are the leading cause of death among the population under the age of 50 (Morens et al. 2004; Fauci 2006a). Deliberate release of pathogenic microorganisms as biological weapons or biological terrorism agents is posing a serious threat to national security (Frischknecht 2003). The battle for control of pathogenic microorganisms is a lengthy process, requiring both detailed knowledge of the pathogens and strategies to control infection. However, pathogenic microorganisms have evolved numerous mechanisms for evasion of human defense systems.

Pathogenic microorganisms are the common enemy of humankind and many countries are increasing the level of research relating to pathogenic microorganisms, especially the United States (US) (Franco and Sell 2011; DeFrancesco 2004; Fauci 2006b) and other developed countries. China and other developing countries are following a similar trend in order to safeguard human health and to enhance national innovation, even with the significant economic restrictions compared with developed countries. The quantity and quality of scientific literature relating to pathogenic microorganisms are important indexes or parameters for evaluating the level of study and the degree of attention focused on pathogenic microorganisms. The Science Citation Index Expanded database (http://apps.isiknowledge.com) is an internationally recognized tool for academic journal evaluation. Moreover, this database facilitates statistical analysis of literature by country, year and institution.

The pathogenic microorganism included in this study were *Bacillus anthracis, Yersinia pestis, Francisella tularensis*, Ebola virus, *Burkholderia pseudomallei*, which belong to biodefense-associated pathogens (BDAPs) and human immunodeficiency virus (HIV), SARS coronavirus, hepatitis B virus, *Mycobacterium tuberculosis*, and the influenza virus, which belong to commonly encountered health-threatening pathogens (CEHTPs). This study analyzed the quantity and variety of SCI literature relating to these important pathogenic microorganisms, published by scientists from the US and China from 1996 to 2010 by searching the Science Citation Index database. This type of comparison provides useful information for the development of appropriate research guidelines relating to pathogenic microorganisms.

Materials and methods

Statistical analysis was conducted to identify trends in the quantity and variety of SCI literature relating to several BDAPs and CEHTPs published from 1996 to 2010 by scientists from the United States and China identified by searches of the Science Citation Index Expanded database (http://apps.isiknowledge.com). The search scope comprised the "title" with the English name of pathogenic microorganisms in double quotes. Searches based on two keywords connected with "or", were: "*B. anthracis*", "*Y. pestis*", "*F. tularensis*", "Ebola virus", "*B. pseudomallei*", "Human immunodeficiency virus" or "HIV", "SARS coronavirus" or "severe acute respiratory syndrome coronavirus", "Hepatitis B virus", "*M. tuberculosis*", "influenza virus". The name of pathogenic microorganisms conformed with those used by the United States Centers for Disease Control and Prevention (CDC) website (CDC 2010) and "Medical Microbiology (Second Edition)" edited by Gu Hong-xi (2009). Search results were selected in August 2011 from literature published from 1996 to 2010. The quantity of literature from the United States and China was filtered according to the "Country/Territory = (USA)" and "Country/Territory = (PEOPLES R CHINA)", respectively. In determining the search criteria, the search range was "title" instead of "theme". This method may miss some results in which the search targets do not appear in the title, although some anomalous results obtained by searches of the "theme" can be avoided. Furthermore, this study focused on comparison of literature related to pathogenic microorganisms among different years and countries, rather than different pathogenic microorganisms. Although the search criteria may not identify all the relevant literature, all publications from different countries and years have the same chance of identification and therefore, the results will not be significantly affected and missed results represent a small proportion of the data. Furthermore, this study focused on literature related to pathogenic microorganisms was selected as the search target rather than the name of the associated diseases. Data analysis did not include Variola major, an important biodefense pathogenic microorganism, due to the small number of SCI literature available.

Results

The quantity of pathogenic microorganism-related literature is large in the United States and literature related to biodefense pathogens has increased rapidly since 2002

Table 1 and Fig. 1 show that the percentages of literature on *B. anthracis*, *Y. pestis*, *F. tularensis*, Ebola virus and *B. pseudomallei* published in the United States were 67, 58, 62, 63 and 17 % respectively of the literature published worldwide for each organism individually. Publications related to BDAPs in the US sharply increased after 2002, which is in accordance with the biodefense budget increase after the 2001 "anthrax mail" event. However, the literature on *B. pseudomallei* in the United States represents a relatively low proportion (Fig. 1), maybe as this disease predominantly occurs as epidemics in Southeast Asia and Australia.

In the US, the quantity of literature relating to CEHTPs is immense (Table 2; Fig. 1). The percentages of publications on HIV, SARS coronavirus, HBV, *M. tuberculosis* and influenza virus were 48, 33, 21, 38 and 44 % respectively of the literature published worldwide for each organism individually. The quantity of HIV-related literature is growing, although this tendency is masked by the large number of publications. SARS coronavirus is a newly emerging pathogen which caused the global SARS outbreak in 2003. Relevant publications reached a peak in 2006 and decreased subsequently. Literature related to HBV has remained stable and represents a relatively low percentage of the worldwide publications, as the incidence of this is not high in the US. Literature on *M. tuberculosis* increased over the period investigated. Publications related to influenza virus were not obvious before 2005. However, the overall number of influenza virus-related literature increased rapidly due to the H5N1 influenza virus and H1N1 influenza virus global epidemic. These data show that scientific research on pathogenic microorganisms is affected by the level of attention focused on associated diseases and emergency situations.

The quantity of pathogenic microorganism-related literature is relatively small in China with an increasing trend towards investigation of certain pathogenic microorganisms

The total Chinese literature relating to pathogenic microorganisms is much lower than that of the United States (Tables 1, 2; Fig. 1). The percentage of publications in China on *B. anthracis*, *Y. pestis*, *F. tularensis*, *B. pseudomallei* were 2, 15, 1 and 2 % respectively of

Table 1	The qu ⁵	untity of bic	odefense-a:	ssociated p:	Table 1 The quantity of biodefense-associated pathogens (BDAPs) in Science Citation Index (SCI) literature from 1996 to 2010	DAPs) in {	Science Cit	ation Index	: (SCI) lite	rature from	1996 to 20	010			
Year	Bacillus	Bacillus anthracis		Yersinia pestis	pestis		Francise	Francisella tularensis	is	Ebola virus	sn		Burkhold	Burkholderia pseudomallei	ıallei
	World	United States	China	World	United States	China	World	United States	China	World	United States	China	World	United States	China
1996	12	3		24	10		19	4	1	13	7		12		1
1997	17	9		21	11		10	3		13	3		16		
1998	16	9		41	21	1	7	1		18	10		22	1	
1999	36	17	2	29	19		9	2		40	19		20	1	
2000	24	12		35	14	1	13	Ζ		26	11		29	1	
2001	30	11		30	13		11	3	1	25	17		24	3	2
2002	78	41	1	26	16		13	5		25	17		28		5
2003	78	51	1	54	21	2	30	8		39	21		33	3	3
2004	94	99	2	55	28	7	42	29		25	16		37	9	
2005	124	6L	3	63	39	6	34	16		28	18		49	7	
2006	149	113	1	62	36	7	62	45	1	34	23		56	16	
2007	150	110	2	85	53	4	96	63		51	41		56	11	
2008	158	122	3	89	60	13	84	58	2	23	16		64	19	1
2009	143	100	7	88	57	14	100	71		22	17		62	16	
2010	110	82	7	106	70	10	76	69	1	36	28		54	12	1
Total	1,219	819	24	808	468	68	624	384	9	418	264	0	562	96	13

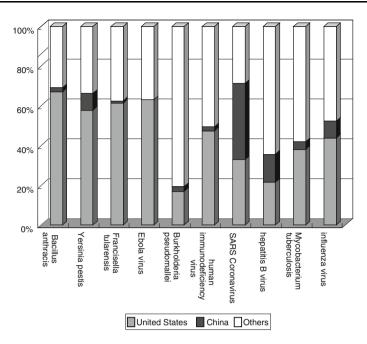


Fig. 1 Percentage of the total pathogenic microorganism-related SCI literature published in the United States and China

the literature published worldwide for each organism individually. The relatively higher number of Chinese publications related to *Y. pestis* may be related to the occurrence of natural epidemics in China.

Compared with the BDAPs, the Chinese SCI literature related to CEHTPs was much larger relative to the total publications worldwide. For example, the percentages of Chinese publications on HIV, SARS coronavirus, HBV, *M. tuberculosis* and influenza virus were 2, 38, 14, 4 and 8 % respectively of the literature published worldwide for each organism individually. There is a large gap in the quantity of HIV-related publications in China compared with the United States, although the growth rate is higher than that of the United States (Fig. 2). The quantity and trends in SARS coronavirus-related publications is similar to that of the United States (Fig. 3). However, the increasing trend is more obvious for some CEHTPs. For example, Chinese HBV-related publications have increased significantly and exceeded those of the United States since 2006 (Fig. 4). Publications related to *M. tuberculosis* and influenza virus have also increased more significantly than those of the United States.

Discussion

Factors impacting on the quantity of literature related to pathogenic microorganisms include the overall national research strength, the number of institutions and scientists engaged in such research, the amount of funding and the extent of the endemic disease. Furthermore, the amount of publications reflects the level of scientific investigations in the field and the degree of attention.

Table (2 The quan	Table 2 The quantity of commonly	ıly encountered health-threatening pathogens (CEHTPs) in Science Citation Index SCI literature from 1996 to 2010	health-thr	eatening p	athogens ((CEHTPs)) in Scienc	e Citation	ו Index SC	I literature	from 1996	to 2010		
Year	Human imr	Human immunodeficiency	virus (HIV)	SARS (SARS Coronavirus	s	Hepatiti	Hepatitis B virus		Mycobac	Mycobacterium tuberculosis	rculosis	Influenza virus	a virus	
	World	United States	China	World	United States	China	World	United States	China	W orld	United States	China	World	United States	China
1996	5,698	2,714	12				400	131	20	283	153	2	132	69	
1997	5,582	2,729	13				381	123	12	287	129		143	75	
1998	5,832	2,795	24				376	109	11	347	163		148	72	1
1999	5,422	2,645	28				420	107	16	343	139	5	156	76	2
2000	5,833	2,832	55				436	98	25	411	175	2	171	91	ю
2001	5,258	2,589	47				483	124	42	421	190	11	149	74	2
2002	5,186	2,536	57				486	136	47	479	199	Ζ	162	LL	3
2003	5,701	2,873	71	52	13	24	555	118	57	523	215	18	180	89	8
2004	6,106	3,064	95	206	2	72	479	106	61	598	236	17	179	62	13
2005	6,554	3,112	147	200	53	103	555	98	80	664	257	31	177	73	19
2006	6,752	3,311	142	153	55	54	781	126	144	629	248	27	266	127	25
2007	7,623	3,618	203	118	46	39	686	122	133	686	227	33	283	119	20
2008	7,658	3,472	249	86	27	29	727	123	133	744	253	40	377	140	51
2009	8,400	3,663	293	65	28	21	712	123	176	765	251	4	508	191	74
2010	8,838	3,928	389	49	21	13	807	136	214	66L	226	09	636	259	89
Total	9,6443	45,881	1,825	929	307	355	8,284	1,780	1,171	8,009	3,061	297	3,667	1,611	310

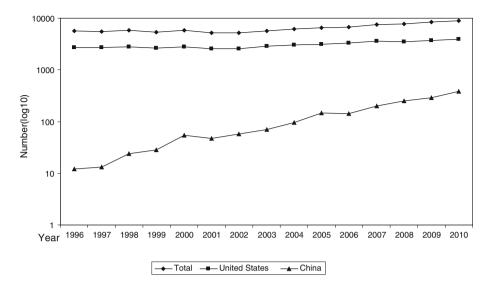


Fig. 2 Annual change in human immunodeficiency virus-related SCI literature

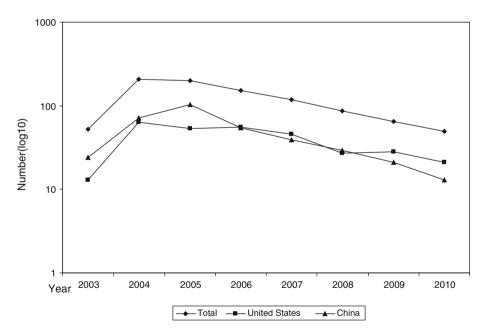


Fig. 3 Annual change in SARS coronavirus-related SCI literature

The rapid growth in publications on BDAPs in the US reflects the national interest in biodefense research. After 2001, the US budget for biological defense research was increased, reaching 66.9 billion dollars (Franco and Sell 2011) to date. Furthermore, "Project BioShield" was launched to strengthen biological defense (Gottron 2003), with a budget of 5.567 billion dollars (Franco and Sell 2011). The US has a large number of

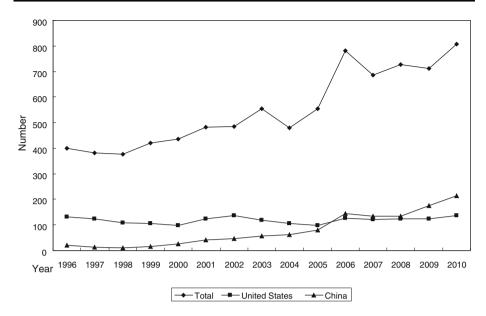


Fig. 4 Annual change in hepatitis B virus-related SCI literature

institutes engaged in biodefense research, including the CDC, the National Institute of Allergy and Infectious Diseases (NIAID) and the Army Medical Research Institute of Infectious Diseases (USAMRIID). Academic institutions such as Harvard University (Malakoff 2003) are also engaged in biodefense research. However, despite this emphasis on biodefense research in the US, studies focused on other important pathogenic microorganisms research, such as HIV and influenza virus are not neglected.

In recent years, China and other developing countries have increased their investment on science and technological innovation and carried out research into pathogenic microorganisms according to their national importance. In 2008, China initiated major national science and technology projects, such as "Prevention and Control of Major Infectious Diseases Including HIV/AIDS and Viral Hepatitis" and "Significant New Drugs Creation", which promoted basic and applied research into pathogenic microorganisms. Funding for research into pathogenic microorganisms is limited in developing countries compared with the US, and improvements in this area are essential. China has the world's largest population, diseases such as AIDS, hepatitis B and tuberculosis impact profoundly on health and social development. Therefore, increased funding to support vital research is critical. China also faces the threat of bioterrorism and emerging infectious diseases and therefore, basic research into pathogenic microorganisms and development of novel drugs and vaccines are required in response to these challenges.

Pathogenic microorganisms are a common global threat to both developed and developing countries and collaboration is vital to overcome these challenges. The study of pathogenic microorganisms requires three balances. Firstly, a balance is required between investment in research on BDAPs and CEHTPs. The United States has vigorously strengthened biodefense research and it has been suggested research into other pathogens may have suffered as a consequence (Atlas 2003; Enserink and Kaiser 2005). Secondly, basic and applied research into pathogenic microorganism should maintain a balance. Basic research provides a better understanding of the characteristics of pathogenic

bating pathogenic microorganisms. The balance between these two aspects of research is critical. For example, in China, the "973" project and National Natural Science Foundation focused on basic research, while the "863" project and the National Science and Technology major projects focused on applied research. Finally, more rapid progress in the study of pathogenic microorganisms should be balanced with meeting biosafety and biosecurity requirements. To date, global research into pathogenic microorganisms, especially biodefense, is extending and the number of laboratories and investigators engaged in studies of hazardous pathogens has increased rapidly. Although the requirement for research into pathogenic microorganisms with regard to biosafety and biosecurity are increasing (Kaiser 2007), it is important that basic investigations should not be compromised.

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