Epidemic obesity and type 2 diabetes in Asia

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The proportions of people with type 2 diabetes and obesity have increased throughout Asia, and the rate of increase shows no sign of slowing. People in Asia tend to develop diabetes with a lesser degree of obesity at younger ages, suffer longer with complications of diabetes, and die sooner than people in other regions. Childhood obesity has increased substantially and the prevalence of type 2 diabetes has now reached epidemic levels in Asia. The health consequences of this epidemic threaten to overwhelm health-care systems in the region. Urgent action is needed, and advocacy for lifestyle changes is the first step. Countries should review and implement interventions, and take a comprehensive and integrated public-health approach. At the level of primary prevention, such programmes can be linked to other non-communicable disease prevention programmes that target lifestyle-related issues. The cost of inaction is clear and unacceptable.

Introduction

The proportions of people with type 2 diabetes and obesity have increased throughout Asia, and the rates of increase show no signs of slowing. The International Diabetes Federation estimates that in 2003, 194 million people had diabetes, and that by 2025, 333 million people will have this disease.1 Some argue that Asia is emerging as the epicentre of this epidemic.2 This region contains some of the most populous countries in the world, and has undergone pronounced demographic, epidemiologic, and socioeconomic change in recent decades. Within the region is the world's most populous country, China, which contains 20% of the world's population, followed by India, with a population of more than one billion. India and China have the greatest numbers of people with diabetes, and are likely to remain in this position in 2025, by which time they could each have 20 million affected individuals.³ The economic burden from type 2 diabetes associated with obesity has also probably been underestimated in this region, which has delayed appropriate prevention and management strategies by national governments and regional agencies. We review the specific clinical characteristics of obesity and type 2 diabetes in Asia, and discuss the effects of environmental changes, such as urbanisation, in the distinct ethnic groups in the region.

Search strategy and selection criteria

The data in this review article are based on MEDLINE and PubMed searches with the search term "diabetes mellitus", "obesity", or both, in combination with the keywords "prevalence", "Asia", and selected Asian countries. A search of MEDLINE and PubMed was also done with search terms "diabetes", "Asian", or both, together with the terms " β -cell defect", "lean body mass", "abdominal adipose tissue", "nutrition transit", and "sedentary lifestyle".

We selected papers that were published between Jan 1970 and July 2006, but also included frequently referenced and highly regarded older papers. Some review articles and book chapters were included because they provide comprehensive overviews that are beyond the scope of this article.

Clinical characteristics of the epidemics

The increase in type 2 diabetes in Asia differs from that reported in other parts of the world: it has developed in a much shorter time, in a younger age group, and in people with much lower body-mass index (BMI).45 Substantial differences in the prevalence of obesity and type 2 diabetes have been noted between the different countries in Asia, and between different locations in the same countries. Evidence also shows that the epidemics of obesity and type 2 diabetes can happen at different stages of urbanisation.6 Although environmental factors such as urbanisation are important, they cannot account for all the characteristics of the epidemic in Asia, and genetic influences probably have an important role. The pronounced differences in the Asian population include the high proportion of body fat and prominent abdominal obesity in Asian people

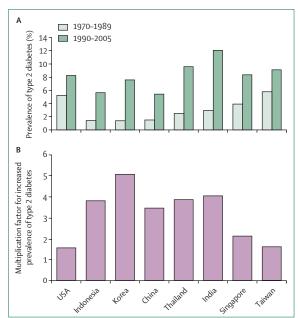


Figure 1: Comparison of prevalence rates of diabetes in selected countries between 1970–1989 and 1990–2005

(A) Prevalence of diabetes in selected nations. (B) Multiplication factor for change in prevalence of diabetes in selected nations between 1970–1989 and 1990–2005.

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	Year	Sample	Age distribution	Method of diagnosis	Classification of diabetes	Adjustment for age
USA ¹³	1976-80	Nationwide	20-74	FBS	‡§	US 2000 census population
	1999-2000	Nationwide	20-74	FBS	‡ §	US 2000 census population
Indonesia ²⁵	1982	Jakarta	Not known	Not known	Not known	Not adjusted
	1995	Jakarta	Not known	Not known	Not known	Not adjusted
Korea	1971 ²³	Rural	≥30	UG+50 g OGTT¶	UG	Not adjusted
	2001 ²⁴	Nationwide	>20	FBS	‡§	2001 standard Korean population
China	198628	Da Qing	25-74	OGTT	*§	Segi standard world population ³⁸
	200029	Nationwide	35-74	FBS	‡§	Not adjusted
Thailand	197126	Urban	Not known	Not known	Not known	Not adjusted
	200027	Nationwide	≥35	FBS	‡§	Not adjusted
India	1979³°	Urban	Not known	Not known	Not known	Not adjusted
	200021	National urban study (six major city)	≥20	OGTT	†§	Urban population of India using the 1991 census population
Singapore	198431	Nationwide	18-69	OGTT	†§	1990 Singapore resident population
	199232	Nationwide	18-69	OGTT	†§	1990 Singapore resident population
Taiwan	1985³³	Taipei city	≥40	FBS or 2 h PG	*§	Segi standard world population ³⁸
	1996³⁴	Tainan city	≥20	OGTT	*\$	Segi standard world population ³⁸

UG=urinary glucose. FBS=fasting blood sugar. PG=plasma glucose. OGTT=oral glucose tolerance test; *WHO, 1985 classification: 35 fasting plasma glucose (FPG) ≥7.8 mmol/L or $2 \text{ h PG} \ge 11.1 \text{ mmol/L}$. 1 mmol/L 1 mmol/L

Table 1: Description of survey data used for figure 1

compared with those of European origin with similar BMI values.^{7,8} These characteristics mean that Asian people have a higher predisposition to insulin resistance at a lesser degree of obesity than people of European descent.^{7,8} Another factor that predisposes Asian people to type 2 diabetes is the pronounced dysfunction in early insulin secretion that has been reported in various populations in Asia.⁹⁻¹²

In the USA, the prevalence of type 2 diabetes has doubled—from 4% to 8% during the past 40 years.13 But although this is dramatic, increases in newly developed and developing countries in Asia have been even greater.1,14 In Chinese adults, the prevalence tripled between 1980 and 1996, from about 1.0% to 3.2%. 15,16 The data from the Indian subcontinent are equally disconcerting,17-22 and the same trend can be seen in other countries in the region. For example, the prevalence rates of type 2 diabetes in Korea, Indonesia, and Thailand have also increased three-fold to five-fold during the past 30 years.23-27 Thus, although the prevalence of type 2 diabetes in Asian countries is currently similar to (or only slightly higher than) that in the USA, the rate at which diabetes has increased during the past three decades (figure 1; table 1) and the likelihood that it will continue to increase at this rate, provide substantial grounds for concern.

In developed countries in which most people are of European descent, diabetes affects mainly those who are older than 65 years.³⁹ But in developing countries, most people with diabetes are aged between 45 and

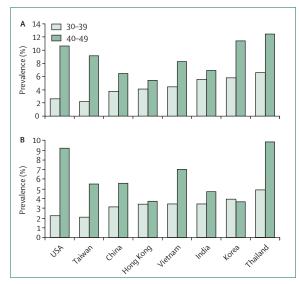


Figure 2: Comparison of prevalence of diabetes between 30 and 40 year-old age-groups in male and female populations
(A) Male. (B) Female.

64 years.³⁹ The prevalence of type 2 diabetes in those aged between 30 and 50 years in developing countries is also high in comparison with other countries. 24,34,40,41 Figure 2 shows the prevalence of diabetes in the 30–39 and 40-49 year age-groups in Asian populations. The prevalence of diabetes in people aged 30-39 years in Asia is generally high-eg, 5.6% of Korean men aged 30-39 years have diabetes (figure 2, table 2). From 1980 to 1996, the prevalence of obesity in children and young adolescents in Taiwan grew steadily, from 4% to 12%.45 Moreover, in China, the proportion of children aged 7 to 18 years who were obese and overweight increased 28-fold between 1985 and 2000.46 The age at which type 2 diabetes develops has also decreased, and the prevalence of the disease in children and adolescents has risen.47 Cases of type 2 diabetes now greatly outnumber cases of type 1 diabetes in children and adolescents.³⁹ Type 2 diabetes mellitus in children has been examined in more detail in Japan than elsewhere in Asia. About 80% of Japanese children with diabetes have the type 2 form. 48 The incidence rate of type 2 diabetes for 1981 to 1990 was reported to be 4.1 per 100 000 person-years, which was about twice as high as the rate of type 1 diabetes in Japanese children.49 Although the prevalence of type 2 diabetes in adolescents in Hong Kong is not as high as in Japan, it is becoming more common.50 Thus, a problem that currently affects only a minority of youth worldwide is threatening the majority in Asia.39

The onset of type 2 diabetes in younger age-groups is likely to result in major economic burdens for countries in Asia due to premature ill health and death. Already, the proportions of patients in Asia who need haemodialysis because of diabetes are similar to those in developed countries. Furthermore, most patients in this region who have end-stage renal disease have type 2 diabetes (figure 3). WHO argues that chronic diseases are the major cause of death in almost all countries, including those in Asia. Only 30% of deaths from chronic disease happen in upper middle-income and high-income countries, whereas 70% of such deaths happen in low-income and lower middle-income countries, such as China, India, Pakistan, Cambodia, and Vietnam.

Evidence from several national health surveys in Asia suggests that the prevalence of overweight and obese individuals has increased, but that it varies between countries (figure 4; table 3). $^{63-65}$ The prevalence of adult obesity (BMI $\geq 30~kg/m^2$) in most Asian countries is quite low compared with developed countries such as the USA. $^{53-55,57,58,60-62}$ The highest rate of obesity in Asia is in Thailand, where $6\cdot 8\%$ of adults are reported to be obese. 52 The lowest obesity rates in the region are in the less developed parts of Asia: $2\cdot 2\%$ in India and $3\cdot 3\%$ in the Philippines. $^{54.55}$ The prevalence of overweight individuals (who have a BMI of more than 25 kg/m² but less than $30~kg/m^2$) ranges from $10\cdot 0\%$ in India to $28\cdot 3\%$ in

	Year	Sample	Age groups	Method of diagnosis	Classification of diabetes
USA ⁴²	1999-2000	Nationwide	20-39 and 40-59	FBS	‡ §
Taiwan ³⁴	1996	Tainan City	30-39 and 40-49	OGTT	*§
China ²⁹	2000-01	Nationwide	35-44 and 45-54	FBS	‡§
Hong Kong⁴⁰	1990	Hospital data	30-39 and 40-49	OGTT	*§
Vietnam ⁴³	2001	Urban	35-44 and 45-54	FBS	‡ §
India ⁴⁴	1999-2002	Nationwide	30-39 and 40-49	OGTT	†§
Korea ²⁴	2001	Nationwide	30-39 and 40-49	FBS	‡ §
Thailand ²⁷	2000	Nationwide	34-44 and 45-54	FBS	‡ §

FBS=fasting blood sugar. OGTT=oral glucose tolerance test. *WHO, 1985 classification: s fasting plasma glucose (FPG) ≥7.8 mmol/L or 2 h PG ≥11.1 mmol/L. † WHO, 1999 classification: FPG ≥7.0 mmol/L or 2 h PG ≥11.1 mmol/L. † American Diabetes Association, 1997 classification: FPG ≥7.0 mmol/L. \$Previously diagnosed diabetes.

Table 2: Survey data used for figure 2

Thailand. $^{53-55,57,58,60-62}$ Since the prevalence of diabetes ranges from $5\cdot 1\%$ to $12\cdot 1\%$, this feature does not seem to correspond with the prevalence of obesity. $^{13\cdot21,2429,32,34,39,56,59}$ For example, the prevalence of obesity in India is the lowest of this group, at $2\cdot 2\%$, but the prevalence of diabetes is the highest, at 12%.

Evidence from some prospective studies in Asia suggests that obesity is directly related to the incidence of diseases such as hypertension, type 2 diabetes, and hypercholesterolaemia. ^{64,66,67} Such diseases have been reported

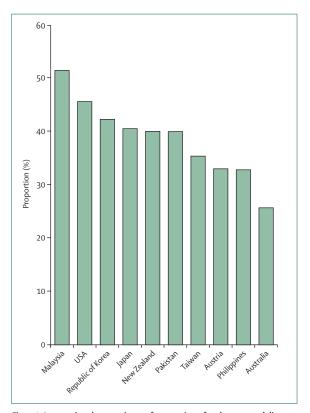


Figure 3: International comparisons of proportion of end stage renal disease patients with diabetes, 2003

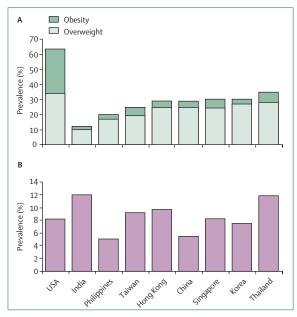


Figure 4: International comparison of prevalence of adult obesity and diabetes

(A) Proportion of overweight and obese adults. (B) Prevalence of diabetes.

	Survey year	Prevalence of overweight adults (%)*	Prevalence of obese adults (%)†	Prevalence of diabetes (%)	
USA	1999-2000 ⁵³	34.0	30-0		
	1999-2000 ¹³			8-2	
India	1998-9954	10.0	2.2		
	200021			12-1	
Philippines	199855	16.9	3.3		
	199656			5.1	
Taiwan	1993-9657	21.1	4.0		
	1996³⁴			9-2	
Hong Kong	1996-9758	25.1	3.8		
	1995-9659			9.8	
China	1999-200051	25.0	4.0		
	2000-200129			5.5	
Singapore	199860	24.4	6.0		
	199232			8-4	
Korea	200161	27-4	3.2		
	200124			7.6	
Thailand	199862	28-3	6.8		
	199539			11.9	
*25 kg/m²≤BMI<30 kg/m². †BMI≥30 kg/m².					
Table 3: Survey data used for figure 4					

in individuals with lower BMI (≥25 kg/m²) than people with the same diseases in developed countries. ^{64,66,67} Thus, in general, although Asians have lower BMI levels than people of European descent, they have a higher prevalence of type 2 diabetes (figure 4). However, waist circumference or waist-to-hip ratio might be more appropriate indices of obesity for Asian people. ⁶⁸

Environmental factors and genetic predisposition

If environmental factors have a major role in triggering development of diabetes, one would expect lower prevalence of diabetes in rural areas, where people follow a traditional lifestyle. This urban–rural difference has been reported in several countries. ^{17,18,27,29,44,69} In India, two populations with different socioeconomic status showed wide differences in the prevalence of diabetes: 8 · 2% in the urban group and 2 · 4% in the rural group, ⁷⁰ and much the same trends have been reported in studies from the Philippines ⁷¹ and Cambodia. ⁶ However, the prevalence of diabetes in the rural population in Thailand ²⁶ and Korea ⁷² was not low compared with the urban population (figure 5). These results suggest that the rate of diabetes might increase in rural communities as they become urbanised.

Rapid economic developments have improved the availability of nutrients, together with socioeconomic and health conditions, in many countries.73 These improvements have led to lower morbidity and mortality and to a pronounced decrease in nutritional deficiencies. However, high nutrient availability, and specifically an energy dense diet, can predispose people to both obesity and type 2 diabetes.74,75 Unlike the gradual transition in nutrient availability that happened in the USA and most European countries, this change has happened rapidly in many lower-income countries. In Asia, economic factors have had especially apparent effects on the nutrition transition.76,77 For example, a rapid shift in dietary structures was reported during Japan's accelerated economic growth from 1950 to 1970.78 An even more rapid shift in diet can be seen in China, especially in urban residents.79-83 Obesity has increased concurrently with these nutritional transitions in most Asian countries.84 South Korea experienced earlier economic change than did most Asian countries and its gross national product increased more than 17 times between 1962 and 1996.85 During this period, the proportion of plant-food in the diet decreased from 97% in 1969 to 79% in 1995.86 By contrast, the animal-food intake increased seven times during that period. Carbohydrate intake

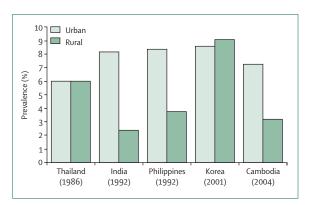


Figure 5: Comparison of the prevalence of diabetes in urban and rural areas

decreased gradually after 1940, from 81% of total energy intake to 64% in 1995. Total protein intake remained nearly constant throughout that period, whereas fat-derived energy intake increased gradually, from $6\cdot2\%$ to $18\cdot8\%$. Similar data have been reported from Japan. Thus, for 5 year-old boys in the Tokyo metropolitan area, the intake of fat as a proportion of total energy intake grew from $12\cdot6\%$ in 1952 to $33\cdot2\%$ in 1994. From 1994.

The behavioural patterns of the very young have been rapidly altered. Fast foods are readily available and lifestyles have become more sedentary due to internet communications, computer games, televised entertainment, academic study and private lessons, poor urban planning, and transportation by motor car.88 The health consequences threaten to become the epidemic of this millennium.88 Furthermore, with rapid shifts in income, changes in occupation linked with reduced physical activity have happened in most developing countries, including Asian countries. Thus, in South Korea in 1960, the urban population was only 27.7%; by 1996 it was 82.3%.86 A shift from energy-intensive occupations in the rural primary product sectors of agriculture, forestry, and fisheries to occupations in services and manufacturing led to the growth in South Korea's gross national product.

In people of European origin, a BMI of 30 correlates with about 25% body fat in men and 30% body fat in women.89 However, for the same age, sex, and BMI, African Americans have a lower fat percentage and Asians have a higher fat percentage.89 One study showed 32% body fat in Chinese females with a BMI of only 21 · 2 kg/m², and 25% body fat content in Chinese males with a BMI of 23 · 7 kg/m². BMI values ranging between 23 and 26 kg/m² have been reported consistently in studies of Chinese people with diabetes for both sexes.90 Thus, Asians have higher body fat content and are at higher risk for diabetes, high blood pressure, and heart disease than other people with the same BMI.91 This difference has been termed the Yudkin-Yajnik paradox.92 Furthermore, at an early stage in the increment of visceral fatness in some Asian people, the risk of dysglycaemia is greater than for Europeans of the same age. 93,94 Park and co-workers have also reported that Asian ethnicity is an independent variable for the determinant of visceral obesity in Asian-American populations.7 These observations prompted WHO to establish an Asia-Pacific guideline for the diagnostic criteria of obesity. WHO concluded that a substantial proportion of Asian people with BMI values lower than 25 kg/m² (and thus not classified as overweight under the existing WHO definition) are at high risk of type 2 diabetes and cardiovascular disease.95 Although understanding of the specific features of obesity in Asia is increasing, little is known about the underlying pathogenesis of increased body fat content and central obesity. Low birthweight has been associated with subsequent risks of non-communicable disease. Strong arguments suggest that the environment does not exclusively determine the health outcomes, but that the so-called thrifty genotype plays a substantial role.⁹⁶

Both a high proportion of body fat and a predominance of central obesity are associated with insulin resistance. A high proportion of Asian people have both these characteristics, and might also have pancreatic β -cell secretory defects. Several studies have reported the importance of early-phase insulin secretory defects in relation to insulin resistance in the development of glucose intolerance in Asian people. 9,10 Some investigators have reported a slight impairment in insulin secretion that begins in subjects with normal glucose tolerance. 11,12 Impaired insulin secretion might be induced by insufficient β -cell mass, by functional defects within the β cells themselves, or both. Although a good linear correlation between β-cell mass and BMI has been reported in normal and type 2 diabetic patients, 97 measurements of β-cell mass in non-obese diabetes patients were lower than those in other patients and were not related to the duration of diabetes or to the glycosylated haemoglobin (HbA_{1c}) levels of patients.⁹⁷ These results suggest that maximal β-cell mass and the regenerative capacity of β cells of an individual patient's response to insulin resistance could be established at an early stage of life, either by the intrauterine environment or by genetically determined factors, or both. Other investigators have reported similar findings.98,99

Leiter has clearly summarised factors that can induce β-cell apoptosis in diabetic patients: so-called glucolipotoxicity (toxic effects of high concentrations of glucose and free fatty acids); low-grade chronic inflammation and amylin deposition with fibrotic islet destruction;100-102 and a reduction of β -cell mass. 99,103,104 Pancreatic stellate cells and angiotensin receptors exist in the islets105 and long-term treatment with the angiotensin converting enzyme (ACE) inhibitor, ramipril, in an animal model of type 2 diabetes showed much improved glucose tolerance tests and reduced islet fibrosis. 106,107 Significant activation of pancreatic stellate cells in the islets of diabetic patients has also been noted. All these results suggest that islet fibrosis and β -cell loss advance over time, and that protection of islet fibrosis might be a strategy to halt the progression of type 2 diabetes.

Interventions and strategies for prevention

The global emergence of obesity and diabetes is an economic issue as much as a health issue. 108 Asia faces especially serious difficulties. Studies such as the Diabetes Prevention Programme, 109 Da-qing study, 110 Finnish Diabetes Prevention Study, 111 Japan lifestyle study, 112 and Indian Diabetes Prevention Programme 113 have shown that lifestyle changes and some medications are effective in prevention of type 2 diabetes in individuals at risk, such as those with impaired glucose tolerance. In particular, lifestyle modification is likely to affect the morbidity and mortality of diabetes, and should be recommended for all people at high risk.

Public health strategies aimed at prevention of weight gain and obesity will probably be more cost effective than treatment of consequences such as diabetes. Therefore, WHO has emphasised that the prevention of obesity and type 2 diabetes should be regarded as of high priority.⁷³

In conclusion, people in Asia develop diabetes at a lower degree of obesity and at younger ages, suffer longer with chronic diabetic complications, and die sooner than those in developed countries. More children are becoming overweight and obese at the same time as Asia undergoes rapid urbanisation, and the incidence of type 2 diabetes reaches epidemic levels in the region. Asian people have different associations between BMI, percentage of body fat and health risks compared with European people. Thus, people in Asia have a strong genetic susceptibility to type 2 diabetes, characterised by early β-cell failure and prominent central obesity. Preventive action should begin urgently, and lifestyle changes such as weight control and exercise are the first step. However, lifestyle is notoriously resistant to change, and public health measures that have been taken so far have been indecisive and insufficiently systematic. Improvement of public health remains an urgent need, since the looming epidemic of diabetes and its complications threatens to drain finite health care resources. Therefore, strong public actions, supported by well-targeted government policies and very clear action plans, will be crucial. As the late Jong-Wook Lee, former Director-General of the WHO, commented, "Until recently, the impact and profile of chronic disease has generally been insufficiently appreciated. It is vital that countries review and implement the interventions described, taking a comprehensive and integrated public health approach. The cost of inaction is clear and unacceptable. Through investing in vigorous and well targeted prevention and control now, there is a real opportunity to make significant progress and improve the lives of populations across the globe."52

Contributors

K H Yoon conceived of the article and suggested its structure. J H Lee, Y H Choi and J H Cho gathered and reviewed literature about the epidemiological issue of obesity and type 2 diabetes, and drafted the manuscript. J W Kim and S H Ko gathered and reviewed literature on the manifestations of type 2 diabetes, and contributed to the manuscript, which was edited by P Zimmet, H Y Son and K H Yoon. The final version was approved by all authors.

Conflict of interest statement

We declare that we have no conflict of interest.

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References

- King H, Aubert RE, Herman WH. Global burden of diabetes, 1995–2025: prevalence, numerical estimates, and projections. Diabetes Care 1998; 21: 1414–31.
- Sicree R, Shaw JE, Zimmet PZ. The global burden of diabetes. In: Gan D, ed. Diabetes Atlas, 2nd edn. Brussels: International Diabetes Federation, 2003: 15–71.

- 3 Roglic G, King H. Diabetes mellitus in Asia. Hong Kong Med J 2000; 6: 10–11
- Ko GT, Chan JC, Cockram CS, Woo J. Prediction of hypertension, diabetes, dyslipidaemia or albuminuria using simple anthropometric indexes in Hong Kong Chinese. Int J Obes Relat Metab Disord 1999; 23: 1136–42.
- 5 He J, Klag MJ, Whelton PK, Chen JY, Qian MC, He GQ. Body mass and blood pressure in a lean population in southwestern China. Am J Epidemiol 1994; 139: 380–89.
- 6 King H, Keuky L, Seng S, Khun T, Roglic G, Pinget M. Diabetes and associated disorders in Cambodia: two epidemiological surveys. *Lancet* 2005; 366: 1633–39.
- 7 Park YW, Allison DB, Heymsfield SB, Gallagher D. Larger amounts of visceral adipose tissue in Asian Americans. *Obesity Res* 2001; 9: 381–87.
- 8 He Q, Horlick M, Thornton J, et al. Sex and race differences in fat distribution among Asian, African-American, and Caucasian prepubertal children. J Clin Endocrinol Metab 2002; 87: 2164–70.
- 9 Chen KW, Boyko EJ, Bergstrom RW, et al. Earlier appearance of impaired insulin secretion than of visceral adiposity in the pathogenesis of NIDDM. 5-Year follow-up of initially nondiabetic Japanese-American men. *Diabetes Care* 1995; 18: 747–53.
- Matsumoto K, Miyake S, Yano M, et al. Glucose tolerance, insulin secretion, and insulin sensitivity in nonobese and obese Japanese subjects. *Diabetes Care* 1997; 20: 1562–68.
- 11 Fukushima M, Suzuki H, Seino Y. Insulin secretion capacity in the development from normal glucose tolerance to type 2 diabetes. *Diabetes Res Clin Pract* 2004; 66 (suppl 1): 37–43.
- 12 Rattarasarn C, Soonthornpan S, Leelawattana R, Setasuban W. Decreased insulin secretion but not insulin sensitivity in normal glucose tolerant Thai subjects. *Diabetes Care* 2006; 29: 742–43.
- 13 Gregg EW, Cadwell BL, Cheng YJ, et al. Trends in the prevalence and ratio of diagnosed to undiagnosed diabetes according to obesity levels in the US. *Diabetes Care* 2004; 27: 2806–12.
- 14 King H, Rewers M. Global estimates for prevalence of diabetes mellitus and impaired glucose tolerance in adults. WHO Ad Hoc Diabetes Reporting Group. *Diabetes Care* 1993; 16: 157–77.
- 15 National Diabetes Co-operative Study Group. A mass survey of diabetes mellitus in a population of 300 000 in 14 provinces and municipalities in China. *Chinese J Intern Med* 1981; 20: 678–83.
- 16 Xiang HD, Liu CQ, Wu W, et al. An epidemiological study on diabetes mellitus 1995–1996, in China. Chinese J Diabetes 1998; 6: 121, 22
- 17 Qiao Q, Hu G, Tuomilehto J, et al. Age- and Sex-Specific Prevalence of Diabetes and Impaired Glucose Regulation in 11 Asian Cohorts. *Diabetes Care* 2003; 26 (6): 1770–80.
- 18 Sadikot SM, Nigam A, Das S, et al. Comparing the ADA 1997 and the WHO 1999 criteria: Prevalence of Diabetes in India Study. Diabetes Res Clin Pract 2004: 66: 309–15.
- 19 Ramachandran A, Snehalatha C, Baskar ADS, et al. Temporal changes in prevalence of diabetes and impaired glucose tolerance associated with lifestyle transition occurring in the rural population in India. *Diabetologia* 2004; 47: 860–65.
- 20 Ramachandran A, Snehalatha C, Vijay V. Temporal changes in prevalence of type 2 diabetes and impaired glucose tolerance in urban southern India. *Diabetes Res Clin Pract* 2002; 58: 55–60.
- 21 Ramachandran A, Snehalatha C, Kapur A, et al. High prevalence of diabetes and impaired glucose tolerance in India: National Urban Diabetes Survey. *Diabetologia* 2001; 44: 1094–101.
- 22 Patandin S, Bots ML, Abel R, Valkenburg HA. Impaired glucose tolerance and diabetes mellitus in a rural population in South India. Diabetes Res Clin Pract 1994; 24: 47–53.
- 23 Kim KS, Choi CH, Lee DY, Jin KE. Epidemiological study on diabetes mellitus among rural Korean. J Kor Diabetes Assoc 1972; 1: 17–24.
- 24 Kim SM, Lee JS, Lee J, et al. Prevalence of diabetes and impaired fasting glucose in Korea: Korean national health and nutrition survey 2001. *Diabetes Care* 2006; 29: 226–31.
- 25 Sutanegara D, Budhiarta AAG. The epidemiology and management of diabetes mellitus in Indonesia. *Diabetes Res Clin Pract* 2000; 50 (suppl 2): 9–16.
- Khalid BA, Usha R, Ng ML, Norella Kong CT, Tariq AR. Prevalence of diabetes, hypertension and renal disease amongst railway workers in Malaysia. *Med J Malaysia* 1990; 45: 8–13.

- 27 Aekplakorn W, Stolk RP, Neal B, et al. The prevalence and management of diabetes in Thai adults: The international collaborative study of cardiovascular disease in Asia. *Diabetes Care* 2003; 26: 2758–63.
- 28 Pan XR, Hu YH, Li GW, Liu PA, Bennett PH, Howard BV. Impaired glucose tolerance and its relationship to ECG-indicated coronary heart disease and risk factors among Chinese. Da Qing IGT and diabetes study. *Diabetes Care* 1993; 16: 150–56.
- 29 Gu D, Reynolds K, Duan X, et al. Prevalence of diabetes and impaired fasting glucose in the Chinese adult population: International Collaborative Study of Cardiovascular Disease in Asia (InterASIA). Diabetologia 2003; 46: 1190–98.
- Chau I, Khalid B. A survey of diabetes mellitus and its complications in the General Hospital, Kuala Lumpur. J ASEAN Fed Endoc Soc 1987; 6: 74–81
- 31 Cheah JS, Yeo PP, Thai AC, et al. Epidemiology of diabetes mellitus in Singapore: comparison with other ASEAN countries. Ann Acad Med Singapore 1985; 14: 232–39.
- 32 Tan CE, Emmanuel SC, Tan BY, Jacob E. Prevalence of diabetes and ethnic differences in cardiovascular risk factors. The 1992 Singapore National Health Survey. *Diabetes Care* 1999; 22: 241–47.
- 33 Chang C-J, Lu F-H, Yang Y-C, et al. Epidemiologic study of type 2 diabetes in Taiwan. Diabetes Res Clin Pract 2000; 50 (suppl 2): 49–59.
- 34 Feng-Hwa Lu, Yang Y-C, Wu J-S, Wu C-H, Chang C-J. A population-based study of the prevalence and associated factors of diabetes mellitus in southern Taiwan. *Diabet Med* 1998; 15: 564–72.
- 35 World Health Organization Study Group. Diabetes Mellitus: Technical Report Series 727. Geneva, Switzerland: WHO, 1985.
- 36 Alberti KGMM, Aschner P, Assal JP, et al. Definition, diagnosis and classification of diabetes mellitus and its complications. Part 1: diagnosis and classification of diabetes mellitus. Geneva, World Health Organization, 1999.
- 37 Gavin JR, Alberti KGMM, Davidson MB, et al. Report of the Expert Committee on the Diagnosis and Classification of Diabetes Mellitus. *Diabetes Care* 1997; 20: 1183–97.
- 38 Segi M. Cancer mortality for selected sites in 24 countries (1950–57). Sendai, Japan: Tohoku University Press, 1960.
- Cockram C. The epidemiology of diabetes mellitus in the Asia-Pacific region. Hong Kong Med J 2000; 6: 43–52.
- 40 Cockram CS, Woo J, Lau E, et al. The prevalence of diabetes mellitus and impaired glucose tolerance among Hong Kong Chinese adults of working age. *Diabetes Res Clin Pract* 1993; 21: 67–73.
- 41 Takahashi Y, Noda M, Tsugane S, Kuzuya T, Ito C, Kadowaki T. Prevalence of diabetes estimated by plasma glucose criteria combined with standardized measurement of HbA1c among health checkup participants on Miyako Island, Japan. *Diabetes Care* 2000; 23: 1092–96.
- 42 Centers for Disease Control and Prevention. Prevalence of diabetes and impaired fasting glucose in adults—United States, 1999–2000. MMWR Morb Mortal Wkly Rep 2003; 52: 833–37.
- 43 Duc Son LN, Kusama K, Hung NT, et al. Prevalence and risk factors for diabetes in Ho Chi Minh City, Vietnam. *Diabet Med* 2004; 21: 371–76.
- 44 Sadikot SM, Nigam A, Das S, et al. The burden of diabetes and impaired glucose tolerance in India using the WHO 1999 criteria: prevalence of diabetes in India study (PODIS). Diabetes Res Clin Pract 2004; 66: 301–07.
- 45 Chu NF. Prevalence of obesity in Taiwan. Obesity Rev 2005; 6: 271-74.
- 46 Wu Y. Overweight and obesity in China. BMJ 2006; 333: 362-63.
- 47 Alberti G, Zimmet P, Shaw J, Bloomgarden Z, Kaufman F, Silink M. Type 2 diabetes in the young: the evolving epidemic: the International Diabetes Federation consensus workshop. Diabetes Care 2004; 27: 1798–811.
- 48 Owada M, Kitagawa T. Diabetes Screening in School Health System. Chiryo 1995; 77: 3047–53.
- 49 Kitagawa T, Owada M, Urakami T, et al. Increased incidence of non-insulin dependent diabetes mellitus among Japanese schoolchildren correlates with an increased intake of animal protein and fat. Clin Pediatr 1998; 37: 111–15.
- 50 Sung RYT, Tong PCY, Yu C-W, et al. High prevalence of insulin resistance and metabolic syndrome in overweight/obese preadolescent Hong Kong Chinese children aged 9–12 years. *Diabetes Care* 2003; 26: 250–51.

- 51 United State Renal Data System. 2005 Annual data report; atlas of end-stage renal disease in the United States. Bethesda, MD, USA: National Institutes of Health, National Institute of Diabetes and Digestive and Kidney Diseases, 2006.
- 52 WHO. Preventing chronic disease: a vital investment. WHO global report. Geneva, Switzerland: World Health Organization, 2005.
- 53 Flegal KM, Carroll MD, Ogden CL, Johnson CL. Prevalence and trends in obesity among US adults, 1999–2000. JAMA 2002; 288: 1723–27.
- 54 Griffiths PL, Bentley ME. The nutrition transition is underway in India. J Nutr 2001; 131: 2692–700.
- 55 Prentice AM. The emerging epidemic of obesity in developing countries. Int J Epidemiol 2006; 35: 93–99.
- 56 Baltazar JC, Ancheta CA, Aban IB, Fernando RE, Baquilod MM. Prevalence and correlates of diabetes mellitus and impaired glucose tolerance among adults in Luzon, Philippines. Diabetes Res Clin Pract 2004; 64: 107–15.
- 57 Lin Y-C, Yen L-L, Chen S-Y, et al. Prevalence of overweight and obesity and its associated factors: findings from National Nutrition and Health Survey in Taiwan, 1993–1996. Prev Med 2003: 37: 233–41.
- 58 Ko GT, Wu MM, Tang J, Wai HP, Chan CH, Chen R. Body mass index profile in Hong Kong Chinese adults. Ann Acad Med Singapore 2001; 30: 393–96.
- 59 Janus ED, Wat NMS, Lam KSL, et al. The prevalence of diabetes, association with cardiovascular risk factors and implications of diagnostic criteria (ADA 1997 and WHO 1998) in a 1996 community-based population study in Hong Kong Chinese. *Diabet Med* 2000; 17: 741-45.
- 60 Deurenberg-Yap M, Chew SK, Lin VFP, Tan BY, van Staveren WA, Deurenberg P. Relationships between indices of obesity and its co-morbidities in multi-ethnic Singapore. *Int J Obes* 2001; 25: 1554-62.
- 61 Kim DM, Ahn CW, Nam SY. Prevalence of obesity in Korea. Obesity Rev 2005; 6: 117–21.
- 62 Aekplakorn W, Chaiyapong Y, Neal B, et al. Prevalence and determinants of overweight and obesity in Thai adults: results of the second national health examination survey. J Med Assoc Thai 2004: 87: 685–93.
- 63 Ge K. Body mass index of young Chinese adults. Asia Pac J Clin Nutr 1997; 6: 175–79.
- 64 Ko GT, Chan JC, Woo J, et al. Simple anthropometric indexes and cardiovascular risk factors in Chinese. Int J Obes Relat Metab Disord 1997; 21: 995–1001.
- 65 Yoshiike N, Matsumura Y, Zaman MM, Yamaguchi M. Descriptive epidemiology of body mass index in Japanese adults in a representative sample from the National Nutrition Survey 1990–1994. Int J Obes Relat Metab Disord 1998; 22: 684–87.
- 66 Oh SW, Shin S-A, Yun YH, Yoo T, Huh B-Y. Cut-off point of BMI and obesity-related comorbidities and mortality in middle-aged Koreans. *Obesity Res* 2004; 12: 2031–40.
- 67 Shin CS, Lee HK, Koh CS, et al. Risk factors for the development of NIDDM in Yonchon County, Korea. *Diabetes Care* 1997; 20: 1842–46.
- 68 Yusuf S. Obesity and risk of myocardial infarction: the INTERHEART study—Author's reply. Lancet 2006; 367: 1054.
- 69 Dong Y, Gao W, Nan H, et al. Prevalence of Type 2 diabetes in urban and rural Chinese populations in Qingdao, China. *Diabet Med* 2005; 22: 1427–33
- 70 Ramachandran A, Snehalatha C, Dharmaraj D, Viswanathan M. Prevalence of glucose intolerance in Asian Indians. Urban-rural difference and significance of upper body adiposity. *Diabetes Care* 1992: 15: 1348–55.
- 71 Ramachandran A, Snehalatha C, Latha E, Vijay V, Viswanathan M. Rising prevalence of NIDDM in an urban population in India. *Diabetologia* 1997; 40: 232–37.
- 72 Korean Ministry of Health and Welfare. Korean national health and nutrition survey. Republic of Korea: Ministry of Health and Welfare, 2002
- 73 WHO. Diet, nutrition and the prevention of chronic diseases. Geneva, Switzerland: World Health Organization, 2003.
- 74 Prentice AM, Jebb SA. Obesity in Britain: gluttony or sloth? BMJ 1995; 311: 437–39.

- 75 Hodge A, Dowse G, Zimmet P. Diet does not predict incidence or prevalence of non-insulin-dependent diabetes in Nauruans. Asia Pac J Clin Nutr 1993; 2: 35–42.
- 76 Popkin B. The nutrition transition and its health implications in lower-income countries. *Public Health Nutr* 1998; 1: 5–21.
- 77 Drewnowski A, BM. P. The nutrition transition: new trends in the global diet. Nutr Rev 1997; 55: 31–43.
- 78 Popkin B. Nutritional patterns and transitions. *Popul Dev Rev* 1993; 19: 138–57.
- 79 Popkin BM, Keyou G, Zhai F, Guo X, Ma H, Zohoori N. The nutrition transition in China: a cross-sectional analysis. Eur J Clin Nutr 1993; 47: 333–46.
- 80 Popkin BM. The nutrition transition in low-income countries: an emerging crisis. *Nutr Rev* 1994; 52: 285–98.
- 81 Popkin BM, Horton S, Kim S, Mahal A, Shuigao J. Trends in diet, nutritional status, and diet-related noncommunicable diseases in China and India: the economic costs of the nutrition transition. *Nutr Rev* 2001; 59: 379–90.
- 82 Du S, Lu B, Zhai F, Popkin BM. A new stage of the nutrition transition in China. *Public Health Nutr* 2002; 5: 169–74.
- 83 Popkin BM, Du S. Dynamics of the nutrition transition toward the animal foods sector in China and its implications: a worried perspective. *J Nutr* 2003; 133 (suppl 2): 3898–906.
- 84 Popkin BM, Doak CM. The obesity epidemic is a worldwide phenomenon. Nutr Rev 1998; 56: 106–14.
- 85 World Bank. World development indicators 1998. Socioeconomic Time Series Access and Retrieval System. Version 4.01. [CD-ROM] Washington, DC: World Bank, 1998.
- 86 Kim S, Moon S, Popkin BM. The nutrition transition in South Korea. Am J Clin Nutr 2000; 71: 44–53.
- 87 Yoshiike N, Matsumura Y, Iwaya M, Sugiyama M, Yamaguchi M. National nutrition survey in Japan. J Epidemiol 1996; 6 (suppl 3): 189–200
- 88 Bar-Or O, Foreyt J, Bouchard C, et al. Physical activity, genetic, and nutritional considerations in childhood weight management. Med Sci Sports Exerc 1998; 30: 2–10.
- 89 Wang J, Thornton JC, Burastero S, et al. Comparisons for body mass index and body fat percent among Puerto Ricans, blacks, whites and Asians living in the New York City area. Obesity Res 1996; 4: 377–84.
- 90 He MZ, Kung AW, Li ET. The validity of BMI as an obesity criterion in Chinese adults. HK Pract 1998; 20 (suppl 2): 6.
- 91 Araneta MRG, Wingard DL, Barrett-Connor E. Type 2 diabetes and metabolic syndrome in Filipina-American women: A high-risk nonobese population. *Diabetes Care* 2002; 25: 494–99.
- 92 Yajnik CS, Yudkin JS. The Y-Y paradox. Lancet 2004; 363: 163.
- 93 Deurenberg-Yap M, Li T, Tan WL, van Staveren WA, P. D. Validation of a semiquantitative food frequency questionnaire for estimation of intakes of energy, fats and cholesterol among Singaporeans. Asia Pac J Clin Nutr 2000; 9: 282–88.
- 94 Tam TTT, Gross R, Lukito W, Rumawas JSP. Chronic energy deficiency and relative abdominal overfatness coexist in free-living elderly individuals in Ho Chi Minh City, Vietnam. Asia Pac J Clin Nutr 1999; 8: 129–35.
- 95 World Health Organization, International Obesity Task Force. The Asian-Pacific perspective: redefining obesity and its treatment. Geneva, Switzerland: WHO Western Pacific Region, 2000.

- 96 Zimmet PZ. Globalization, coca-colonization and the chronic disease epidemic: can the Doomsday scenario be averted? J Intern Med 2000; 247: 301–10.
- 97 Yoon KH, Ko SH, Cho JH, et al. Selective beta-cell loss and alpha-cell expansion in patients with type 2 diabetes mellitus in Korea. J Clin Endocrinol Metab 2003; 88: 2300–08.
- 98 Butler AE, Janson J, Bonner-Weir S, Ritzel R, Rizza RA, Butler PC. Beta-cell deficit and increased beta-cell apoptosis in humans with type 2 diabetes. *Diabetes* 2003; 52: 102–10.
- 99 Sakuraba H, Mizukami H, Yagihashi N, Wada R, Hanyu C, Yagihashi S. Reduced beta-cell mass and expression of oxidative stress-related DNA damage in the islet of Japanese Type II diabetic patients. *Diabetologia* 2002; 45: 85–96.
- 100 Gepts W, Lecompte PM. The pancreatic islets in diabetes. Am J Med 1981; 70: 105–15.
- 101 Ken S, Nobuhisa Y, Takahashi T. Differential volumetry of A, B and D cells in the pancreatic islets of diabetic and nondiabetic subjects. *Tohoku J Exp Med* 1979; 129: 273–83.
- 102 Kloppel G, Lohr M, Habich K, Oberholzer M, Heitz PH. Islet pathology and pathogenesis of type 1 and type 2 diabetes mellitus revisited. Surv Synth Path Res 1985; 4: 110–25.
- 103 Clark A, Wells CA, Buley ID, et al. Islet amyloid, increased A-cells, reduced B-cells and exocrine fibrosis: quantitative changes in the pancreas in type 2 diabetes. *Diabetes Res* 1988; 9: 151–59.
- 104 Stefan Y, Orci L, Malaisse-Lagae F, Perrelet A, Patel Y, Unger RH. Quantitation of endocrine cell content in the pancreas of nondiabetic and diabetic humans. *Diabetes* 1982; 31: 694–700.
- 105 Ko SH, Hong OK, Kim JWK, et al. High glucose increases extracellular matrix production in pancreatic stellate cells by activating the renin-angiotensin system. *J Cell Biochem* 2006; 98: 343–55.
- 106 Ko S-H, Kwon H-S, Kim S-R, et al. Ramipril treatment suppresses islet fibrosis in Otsuka Long-Evans Tokushima fatty rats. Biochem Biophys Res Commun 2004; 316: 114–22.
- 107 Cooper ME, Tikellis C, Thomas MC. Preventing diabetes in patients with hypertension: one more reason to block the renin-angiotensin system. J Hypertens 2006; 24 (suppl 1): 57–53.
- 108 Derek Yach, David Stuckler, Brownell KD. Epidemiologic and economic consequences of the global epidemics of obesity and diabetes. *Nature Med* 2006; 12: 62–66.
- 109 Knowler WC, Barrett-Connor E, Fowler SE, et al. Reduction in the incidence of type 2 diabetes with lifestyle intervention or metformin. N Engl J Med 2002; 346: 393–403.
- 110 Pan XR, Li GW, Hu YH, et al. Effects of diet and exercise in preventing NIDDM in people with impaired glucose tolerance. The Da Qing IGT and Diabetes Study. *Diabetes Care* 1997; 20: 537–44.
- 111 Tuomilehto J, Lindstrom J, Eriksson JG, et al. Prevention of type 2 diabetes mellitus by changes in lifestyle among subjects with impaired glucose tolerance. N Engl J Med 2001; 344: 1343–50.
- 112 Kosaka K, Noda M, Kuzuya T. Prevention of type 2 diabetes by lifestyle intervention: a Japanese trial in IGT males. Diabetes Res Clin Pract 2005; 67: 152–62.
- 113 Ramachandran AA, Snehalatha CC, Mary SS, Mukesh BB, Bhaskar AAD, Vijay VV. The Indian Diabetes Prevention Programme shows that lifestyle modification and metformin prevent type 2 diabetes in Asian Indian subjects with impaired glucose tolerance (IDPP-1). Diabetologia 2006; 49: 289–97.