

Review Article

Health Risks of Obesity

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Abstract

Obesity epidemic is taking the globe in its stride with the metamorphosed lifestyle. Obesity prevalence is constantly increasing with over a third of population being overweight or obese in India. It contributes in a big way to pathogenesis of chronic non-communicable diseases. Interventions to reduce burden of obesity, partly depend on detecting and understanding the risks complicating obesity. This article summarizes the need for focused clinical skills on obesity related health risks and vigorous validation of benefits possible through interventions toward weight management in patients.

Key Words: Obesity, Risks of Obesity, Metabolic Syndrome

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Introduction

World Health Organization has enabled the classification of individuals as overweight or obese by using a measure of Body Mass Index. Body Mass Index is calculated by dividing body weight in kilogram by square of height in meters. This is essentially the surrogate measure of body fat for specified categories of sex and age¹. A value equal to or more than 30kg/m², defines state of obesity². Different thresholds are defined by anthropometry (BMI) for overweight and obesity is stated for different ethnicity³. Prevalence of obesity increases rapidly in India. Large surveys conducted in different parts, time to time, indicate steady rising trends across all sections of population. In 2004, sample surveyed indicated 20.8% male and 32.3% female population overweight or obese. In 2007, urban males 32.4% and females 41.4% were seen as overweight or obese. Lately, in 2012, 46.6% of urban women above 35 year age and 23.7% of such rural women are overweight. Childhood and pre-pubertal obesity is also fast increasing, particularly in northern part⁴.

The distribution pattern of central obesity is, indicative of excess visceral fat. It exhibits stronger association with cardiovascular diseases than subcutaneous fat deposited around hip. Central obesity is reflected in waist circumference and the waist/hip circumference ratio. The anatomic location profoundly influences biology of adipose tissue. Increased visceral fat associates metabolic dysfunction and cardio-metabolic risk while subcutaneous fat may be protective. The two adipose tissues have embryologically different origins⁵. Visceral adipose tissue and its resident macrophages produce high quantities of proinflammatory cytokines, viz, TNF-alpha, IL-6, and low amounts of protective adiponectin. These changes in cytokines induce insulin resistance, endothelial dysfunction and consequent atherosclerosis⁶.

Mortality in Obesity

Body mass index (BMI) in excess of certain threshold for specific age/sex groups, is seen to be associated with increased mortality. Six to seven year reduction of longevity was found in a prospective observation cohort⁷. 3457 participants of Framingham Heart study were inducted at 30-49 year age. Substrata were made by sex and smoking status. Life expectancy and probability of death before 70 year age was analyzed using life table. Mortality rates specific for categories of different BMI (normal, overweight and obese at baseline/induction) and age groups were analyzed. Overweight and obesity caused large reductions in life expectancy. The follow up period was 42 years from 1948 to 1990. BMI at ages 30 to 49 years predicted mortality after 50 to 69 years, even upon adjustment for BMI at 50-69 year age⁸.

High waist (central) fat distribution increases while high fat distributed to hip decreases risk of getting myocardial infarction. It was a standardized case control study of acute myocardial infarction with 14037 controls and 12461 cases from multiple countries and ethnicities. The waist to hip ratio showed a graded and highly significant association with risk of myocardial infarction. BMI showed modest association, which disappeared when adjustments were made for waist hip ratio and other risk factors⁹.

Obesity and Diabetes

Type 2 Diabetes mellitus is the major most prevalent chronic non communicable inflammatory disease, arising as sequel to metabolic syndrome driven by insulin resistance. Overweight and obesity is found to be the main predictor of type 2 diabetes mellitus^{10,11}. During 16 year follow up of 84941 healthy female nurses, from 1980 to 1996, 3300 new cases of type 2 diabetes were detected. Overweight or obesity, as per

BMI defined cutoff constituted single most important predictor of diabetes development¹⁰. Data from cohort of 51529 US male health professionals in age range 40-75 years in 1986 was analyzed following 5 year follow up, in 1992. Persons developing NIDDM numbered 272. Strong positive association of obesity defined by BMI and risk of diabetes development was demonstrated¹¹.

Mechanistic links of obesity to diabetes development are hypothesized with partial supportive observations. Genetic and epigenetic predisposition for preferential deposition of fat in abdominal depots is proposed. This process may be added by stress induced hypothalamo-pituitary-adrenal axis activation. The overflow of uncontained fat from omental and mesenteric tissue probably invades organs as liver, skeletal muscles, inflicting metabolic dysfunction. Direct effect of omental and mesenteric adipose tissue depots may be exerted on insulin resistance, lipoprotein metabolism and blood pressure. Metabolic products of omental and mesenteric adipose tissue depots are released in to the portal vein, which provides direct delivery to liver. Lipolysis of omental and mesenteric adipose tissue triacyl-glycerol releases free fatty acids that can induce hepatic insulin resistance and provide substrate for lipoprotein synthesis and neutral lipid storage in hepatocytes. In addition, specific proteins and hormones produced by omental and mesenteric adipose tissue, such as inflammatory adipokines, angiotensinogen and cortisol (generated by local activity of 11 beta hydroxysteroid dehydrogenase), can also contribute to cardio-metabolic disorder¹².

Hypertension and Obesity

Obesity increases risk of developing hypertension^{13,14}. In a large study, 14924 adult participants were grouped by BMI cutoffs and waist circumference as per NIH criteria. Relative risks of hypertension, dyslipidaemia and metabolic syndrome were higher in overweight and obese. These risks correlated positively with waist circumference. Importantly, risk level associating a given waist circumference was unaffected by differences in BMI, emphasizing major significance of the parameter¹⁵.

Obesity and Heart Disease

Number of studies have established regular relation of obesity to increased risk of coronary artery disease^{14,16}. The obese are also twice as likely to suffer heart failure as the non obese¹⁷. Paradoxically, the survival rate of obese heart failure cases is better than the non obese¹⁸.

Obesity and Cerebrovascular Disease

Increased BMI associates increased risk of ischemic stroke. Hemorrhagic stroke risk appears to concentrate at either extremes of the BMI¹⁹. Lean body weight and lower cholesterol level have been linked to increased hemorrhagic stroke risk by a prospective cohort of 21414 male physicians²⁰. During 12.5 years of follow-up, 747 strokes (631 ischemic, 104 hemorrhagic, and 12 undefined) occurred. Compared with participants with BMIs less than 23, those with BMIs of 30 or greater had an adjusted relative risk of 2.00 (95% confidence

interval [CI], 1.48-2.71) for total stroke, 1.95 (95% CI, 1.39-2.72) for ischemic stroke, and 2.25 (95% CI, 1.01-5.01) for hemorrhagic stroke. When BMI was evaluated as a continuous variable, each unit increase of BMI was associated with a significant 6% increase in the adjusted relative risks of total (95% CI, 4%-8%), ischemic (95% CI, 3%-8%), and hemorrhagic stroke (95% CI, 1%-12%). Additional adjustment for hypertension, diabetes mellitus, and hypercholesterolemia slightly attenuated the risks for total and ischemic (relative risk 4%; 95% CI, 2%-7%), but not hemorrhagic, stroke. Another study was conducted in Korea, on 2,712 persons (904 cases, 904 hospital controls, and 904 community controls)²¹. Obese men ($25.0 \leq \text{BMI} < 30.0 \text{ kg/m}^2$) had an odds ratios (OR) of 1.39 (95% CI 1.03 to 1.87) a hemorrhagic stroke, compared to men with a normal BMI (18.5 to 24.9 kg/m^2). Conversely, women with lower BMI had a higher risk of having hemorrhagic stroke. About a three-fold increase was observed in the risk of intracerebral hemorrhage (ICH) in the highly obese group obesity was identified as one of the risk factors in hemorrhagic stroke, in particular ICH. Conversely, in women, a lean body weight increases the risk of hemorrhagic stroke. Consequently, managing one's weight is essential to reduce the risks of hemorrhagic stroke. Central (truncal) obesity increases stroke related mortality, however²².

Role of Obesity in Metabolic Syndrome

Metabolic syndrome is defined by presence of any three the following five features: viz. 1) waist circumference above 40 inches in male and 35 inches in female; 2) triglyceride level above 150 mg/dl; 3) High Density Cholesterol (HDL) level below 40mg/dl in male or 50mg/dl in female; 4) Blood pressure above 130/85 mmHg; and 5) Fasting blood glucose level above 100mg/dl²³. The altered lipid and glucose metabolism appears to be consequent to central obesity and insulin resistance.

Obesity and Pulmonary Abnormality

Obesity and accompanied increased neck circumference is found to strongly link to the obstructive sleep apnea²⁴. Increased fat tissue may impinge upon airway lumen increasing collapsibility²⁵. Obesity also increases asthma risk and majority of asthmatic emergency cases are found to be obese or overweight²⁶. Obesity asthma linkage may involve decrease in functional and tidal volume, chronic state of sub-acute systemic inflammation including increased production of mediators and the increased hyper responsiveness of airway²⁷.

Obesity and Gastrointestinal Disorders

Obesity is significantly associated to gastro esophageal reflux disease and its late sequel as erosive esophagitis, esophageal cancer etc²⁸. Obese men and women both suffer increased incidence of gall stones^{29,30}.

Obesity and Reproduction Disorders

Obesity and insulin resistance typically associate the polycystic ovarian disease that exhibits anovulation and hyper-androgenism. Weight loss and therapeutic remediation of insulin resistance can reverse anovulation and aberrant hormonal profile in these

patients³¹. Obesity negatively impacts male sexual function and fertility, with recovery benefit of weight reduction³². Gestational diabetes is likewise more prevalent in the obese with predictable diverse complications for mother and baby³³.

Obesity and Late Age Health

Osteoarthritis ensues upon obesity³⁴. Besides the extra strain on weight bearing joints, there is evidence of systemic dysregulation of adipokines, to explain increased osteoarthritis in the obese people³⁵. Increased incidence of certain cancers in the obese is reported, viz. cancers of gall bladder, esophagus, thyroid, kidney, uterus, colon, breast etc³⁶. A lean body weight is known to reduce both incidence and mortality of the cancers^{37,38}. Cancers of reproductive and secondary sex organs in females may draw boost from high estrogen synthesis from increased fat³⁹. The evidences are most consistent for endometrial cancer, breast cancer among the postmenopausal women, and renal cell cancer. The molecular mechanisms linking obesity to increased cancer incidence have been poorly understood.

Obesity & Health Related Quality of Life

Several neuropsychiatric maladies like somatization, hypochondriasis, phobias, and obsessive compulsive disorders have shown amelioration in obese patients after undergoing bariatric surgery⁴⁰. Many surgical procedures in the obese carry poor prognoses due to associated ill health. Obesity evidently would increase immobility and access to proper health care.

Conclusion

Thus, given the magnitude of complications related to obesity, addressing this linkage and revelation of diseases linked to obesity constitute, vital facets of clinical competence and quality care.

Authors declare no conflict of interest.

References

- 1) Pasco JA, Nicholson GC, Brennan SL, Kolowicz MA. Prevalence of obesity and the relationship between the body mass index and body fat: cross sectional population based data. *PLoS One* 2012; 7(2):e29580.
- 2) WHO Obesity: Preventing and managing the global epidemic. Report of WHO Consultation on Obesity 1998, Geneva.
- 3) WHO Technical Report Series 894. Obesity: Preventing and managing the global epidemic. Report of WHO Consultation Geneva 3-5 June 1997. 2000 WHO Press.
- 4) Pandey AK, Pandey G, Pandey SS, Pandey BL. Human biology of diet and lifestyle linked chronic inflammatory non communicable disease epidemic-a review. *Human Biology Review* 2014; 3:25-42.
- 5) Chau Y, Bandiera R, Serrels A, Martinez-Estrada OM, Qing W, Lee M et.al. Visceral and subcutaneous fat have different origins and evidence supports a mesothelial source. *Nature Cell Biology* 2014; 16:367-375.
- 6) Pandit A, Pandey AK. Atherosclerosis: Current Perspective. *Apollo Medicine* 2015 (in press)
- 7) Peeters A, Barendregt JJ, Willekens F, Mackenbach JP, Al Mamun A, Bonneux L. Obesity in adulthood and its consequences for life expectancy: a life table analysis. *Ann Intern Med* 2003; 138: 24-32.
- 8) Dawber TR, Meadows G F, Moore F E Jr. Epidemiological approaches to heart disease : The Framingham study. *Am J Pub Health.* 1951;41:279-281
- 9) Yusuf S, Hawken S, Ounpuu S, Bautista L, Franzosi MG, Commerford P et.al. Obesity and the risk of myocardial infarction in 27000 participants from 52 countries:a case control study *Lancet* 2005; 366: 1640-1649
- 10) Frank B, Hu FB, Manson JE, Stampfer MJ, Colditz G. Diet Lifestyle and the risk of type 2 diabetes in women *N.Engl J Med.* 2001; 345: 790-797
- 11) Chan JM, Rimm EB, Colditz GA, Stampfer MJ, Willet MC. Obesity and fat distribution and weight gain as risk factors for clinical diabetes in men *Diabetes Care* 1994 ;17: 961-969
- 12) Klein S, Allison DB, Heymsfield SE, Kelley DE, Leibel RL et al. Waist circumference and cardio-metabolic risk:a consensus statement from Shaping Americas Health. Association for Weight Management and Obesity Prevention. NAASO; The Obesity Society. The American Society for Nutrition and, The American Diabetes Association. *Am J Clin Nutr* 2007; 85:1197-1202
- 13) Huang Z, Willet WC, Manson JE, Rosner B, Stampfer MJ, Specizer FE et al. Body weight, weight change and risk for hypertension in women *Ann Intern Med.* 1998; 128: 81-88
- 14) Wilson PW, DAgostino RB, Sullivan L, Parise H, Kannel WB. Over weight and obesity as determinants of cardiovascular risk:the Framingham experience. *Arch Intern Med.* 2002; 162: 1867-1872
- 15) Jansen I, Katzmarzyk PT, Ross R. Waist circumference and the body mass index explains obesity related health risk *Am J Clin Nutr.* 2004; 79: 379-384
- 16) Willen WC, Manson JE, Stampfer MJ, Colditz GA, Rosner B, Speizer FE et al. Weight,weight change and coronary heart disease in women:Risk within the normal weight range. *JAMA* 1995; 273: 461-465
- 17) Kenchaiah S, Evans JC, Levy D, Wilsen PV, Benjamin EJ, Larson MC et al. Obesity and the risk

- of heart failure *N Engl J Med* 2002; 347: 305-313
- 18) Curtis JP, Salter JG, Wang Y, Rathore SS, Jovin IS, Jadbabaie F et al. The obesity paradox: Body mass index and outcomes in patients with heart failure *Arch Intern Med*. 2005; 165: 55-61
- 19) Song YM, Sung J, Davey Smith G, Ebrahim S. Body mass index and ischaemic and haemorrhagic stroke: a prospective study in Korean men *Stroke* 2004; 35: 831-836
- 20) Kurth J, Gaziano JM, Berger K, Kase CS, Rexrode KM, Cook NR et al. Body mass index and the risk of stroke in men *Arch Intern Med*. 2002; 162: 2557-2562
- 21) Kim SH, Lee YS, Lee SM, Yoon BM, Park BJ. Body mass index and risk of haemorrhagic stroke in Korean adults-case control study (Korean) *J Prev Med Public Health* 2007; 40: 313-320
- 22) Tanne D, Medalie JH, Goldbourt U. Body fat distribution and long term risk of stroke mortality *Stroke* 2005; 36: 1021-1025
- 23) Executive summary. Third Report of the National Cholesterol Education Programme (NCEP). Adult Treatment Panel III. *JAMA* 2001; 285: 2486-2497
- 24) Katz I, Stradling J, Slutsky AS, Zanel N, Hoffstein V. Do patients with obstructive sleep apnoea have thick necks? *Am Rev Respir Dis*. 1990; 141: 1228-1231
- 25) Schwartz AR, Gold AR, Schubert N, Stryzak A, Wisi RA, Permut S et al. Effect of weight loss on upper airway collapsibility in obstructive sleep apnoea *Am Rev Respir Dis*. 1991; 144: 494-498
- 26) Guerra S, Wright AI, Morgan WJ, Sherrill DI, Holberg CJ, Martinez FD. Persistence of asthma symptoms during adolescence: role of obesity and age at onset of puberty *Am J Respir Crit Care Med*. 2004; 170: 78-85
- 27) Shore SA, Fredberg JJ. Obesity, smooth muscle and airway hyperresponsiveness *J Allergy Clin Immunol* 2005; 115: 925-927
- 28) Hompel H, Abraham NS, El Serag HB. Metaanalysis: Obesity and the risk of gastroesophageal reflux disease and its complication *Ann Intern Med*. 2005; 143: 199-211
- 29) Stampfer MJ, Macbre KM, Colditz GA, Manson JE, Willet WC. Risk of symptomatic gall stones in women with severe obesity *Am J Clin Nutr*. 1992; 55: 652-658
- 30) Willet WC, Dietz WH, Golditz GA. Guidelines for healthy weight *N Engl J Med*. 1999; 341 :427
- 31) Ehrmann DA. Medical Progress: Polycystic ovarian syndrome *N Engl J Med*. 2005; 352: 1223-1236
- 32) Eposito K, Gugliano F, DiPalo C, Giugliano G, Mavfella R, D Adrea F et al. Effect of lifestyle changes on erectile dysfunction in obese men: a randomized controlled trial *JAMA*. 2004; 291: 2978-2984
- 33) Diel J. Maternal obesity and complications during pregnancy *J Perinat Med*. 2005; 33: 100-105
- 34) Cicuttini FM, Specter TD. Osteoarthritis in the aged: Epidemiological issues and optimal management *Drugs Ageing* 1995; 6: 409-420
- 35) Aspeden RM, Scheven BA, Hutchinson JD. Osteoarthritis as a systemic disorder including stromal cell differentiation and lipid metabolism *Lancet* 2001; 357: 1118-1120
- 36) Renchan AG, Tyson M, Egger M, Heller RF, Zwahlen M. Body mass index and the incidence of cancer: a systematic review and meta analysis of prospective observational studies *Lancet* 2008; 371: 578
- 37) Sjostrom I, Gummesson A, Sjostrom CD. Effect of bariatric surgery on cancer incidence in obese patients in Sweden: a prospective controlled observational trial *Lancet Oncology* 2009; 10: 653-662
- 38) Adams TD, Stroop AM, Gress RE. Cancer incidence and mortality after gastric bypass surgery *Obesity* 2009; 17: 796-802
- 39) Schmandt RE, Iglesias DA, Co NN, Lu KH. Understanding obesity and endometrial cancer risk: opportunities for prevention *Am J Obstet Gynaecol* 2011; 205: 518-525
- 40) Rosik CH. Psychiatric symptoms among prospective bariatric surgery patients: rates of prevalence and their relation to social desirability *Obes Surg*. 2005; 15: 677-683