## WORLD JOURNAL OF PHARMACEUTICAL RESEARCH

SJIF Impact Factor 8.084

Volume 12, Issue 7, 1246-1261.

**Review Article** 

ISSN 2277-7105

# A REVIEW: FINDING A LINK BETWEEN HYPERTENSION AND **DIABETES**

Nidhi  $Zar^1$ , Shivam Choudghal $*^2$  and Rajat Sharma $^3$ 

<sup>1</sup>Clinical Pharmacist, Department of Clinical Pharmacology, Shri Mata Vaishno Devi Narayana Super-Speciality Hospital, Katra, J&K (182320).

<sup>2</sup>Research Scholar, Department of Pharmacy Practice, ISF College of Pharmacy, Moga, Punjab (142001).

<sup>3</sup>Research Scholar, Department of Pharmacy Practice, Chandigarh Group of Colleges, Landran, Mohali, Punjab (184144).

Article Received on 20 March 2023,

Revised on 10 March 2023, Accepted on 30 April 2023,

DOI: 10.20959/wjpr20237-28081

### \*Corresponding Author Shivam Choudghal

Research Scholar. Department of Pharmacy Practice, ISF College of Pharmacy, Moga, Punjab (142001).

#### **ABSTRACT**

Diabetes and Hypertension commonly occur together. There is considerable overlap between diabetes and hypertension in etiology and disease mechanisms. Obesity, inflammation, oxidative stress, and insulin resistance are thought to be the common pathways. Most patients with type 2 diabetes are insulin resistant, and about half of those with essential hypertension are insulin resistant. Therefore, insulin resistance is an important common link between diabetes and hypertension. Recent advances in the understanding of these pathways have provided new insights and perspectives. Physical activity plays an important protective role in the two diseases. By knowing the common causes and disease mechanisms allows us more effective and proactive

approach in the prevention and treatment of the two diseases. Hypertension in the diabetic individual markedly increases the risk of cardiac disease, peripheral vascular disease, stroke, retinopathy, and nephropathy. Diabetic nephropathy is an important factor involved in the development of hypertension in diabetics, particularly type I patients. Increased sodium may also play a role in the pathogenesis of blood pressure in diabetics. There is increasing evidence that insulin resistance/hyperinsulinemia may play a key role in the pathogenesis of hypertension in both subtle and overt abnormalities of carbohydrate metabolism. Population studies suggest that elevated insulin levels, which often occurs in type II diabetes mellitus, is an independent risk factor for cardiovascular disease. Other cardiovascular risk factors in

diabetic individuals include abnormalities of lipid metabolism, platelet function, and clotting factors. The goal of antihypertensive therapy in the patient with coexistent diabetes is to reduce the inordinate cardiovascular risk as well as lowering blood pressure.

**KEYWORDS:** Diabetes, Hypertension, Obesity, Metabolic syndrome, Metabolic pathway, Insulin resistance, peripheral vascular disease, stroke, retinopathy, and nephropathy, insulin resistance/ hyperinsulinemia.

#### INTRODUCTION

Hypertension and diabetes are two of the main hazard factors for atherosclerosis and its entanglements which incorporates heart assaults and strokes. There is a covering among diabetes and hypertension, reflecting significant cover in their etiology and malady components. Hypertension, or hypertension, is a condition that is found in individuals with sort 2 diabetes. It's obscure for what reason there's such a noteworthy connection between the two maladies. It's accepted that the accompanying adds to the two conditions: corpulence. an eating routine high in fat and sodium. constant aggravation, inertia. In the Hong Kong Cardiovascular Hazard Factor Commonness Study, just 42% diabetic patients had typical pulse and just 56% with hypertension had ordinary glucose resilience. In the US populace, hypertension happens in around 30% of sort 1 diabetic patients and in half to 80% of patients with sort 2 diabetes mellitus. A planned report in the US revealed that type 2 diabetes mellitus was practically 2.5 occasions as prone to create in patients with hypertension as in patients with typical circulatory strain. Diabetes and hypertension are found in a similar individual more regularly than would happen by some coincidence, though the cover among deglycation and raised circulatory strain is much more extensive than that among diabetes and hypertension. This recommends either shared hereditary or ecological factors in the etiology.

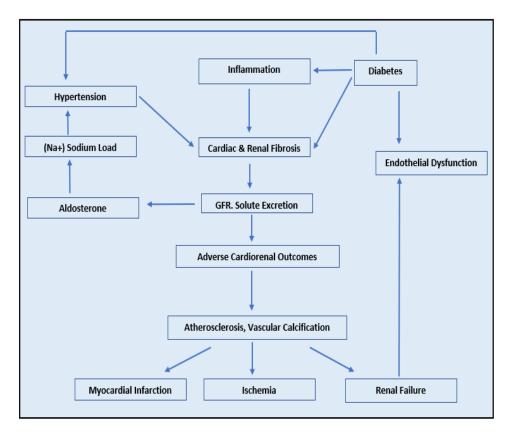


Figure 1.

#### **ETIOLOGY**

### Hereditary qualities

Genome sweeps including a huge number of topics and controls have uncovered numerous qualities with little impacts, instead of few qualities with huge impacts expected initially. Hereditary variations in the quality encoding angiotensinogen, adrenomedullin, apolipoprotein, and  $\alpha$ -adducin have been accounted for to go with regular conditions, for example, diabetes, hypertension, deglycation, or metabolic disorder. In investigations of single nucleotide polymorphisms (SNPs), SNPs that anticipate the improvement of diabetes were found additionally to foresee the advancement of hypertension. In genome examines in Hong Kong Chinese people, the district related with diabetes was likewise related with the metabolic disorder, which incorporates hypertension as a constituent. An ongoing report at Columbia College on physical quality transformation and erasure recommended that huge numbers of normal SNPs are included.

#### **Obesity**

Corpulence, a worldwide medical issue, has been distinguished as the most significant hazard factor for hypertension and diabetes. Stout people have an essentially higher danger of hypertension and type 2 diabetes. Investigations of heftiness in Western nations where there is a high commonness have prompted a more noteworthy comprehension of the wonder of hazard factor grouping and of the pathophysiologic joins among hypertension, corpulence, diabetes. Weight is commonly considered as the joined consequence of brokenness of encouraging focus in the mind, lopsidedness in vitality admission and consumption, and hereditary varieties. Corpulence is to a great extent controlled by qualities; roughly half to 90% of the variety in weight is the aftereffect of hereditary inclination as indicated by twin examinations.

The corpulent (ob.) quality that was found in 1950 was the primary quality distinguished to be identified with the beginning of weight. From that point on, specialists have looked to recognize the hereditary components of weight notwithstanding concentrating metabolic physiology. Genome-wide affiliation studies have uncovered various qualities affecting the helplessness to weight. The FTO quality, advancing weight and indulging, was one of the key corpulence weakness qualities. Together with the GNDPA 2 quality, they foresee diligent focal heftiness in the Chinese populace. Other likely diabetes-related qualities incorporate BCDIN3D/FAIM2, SH2B1, and KCTD15 just as CRTC3, which has been appeared to hinder the speed of fat oxidation.

It isn't astounding to find that diabetes and heftiness share some regular powerlessness qualities. As heftiness is a typical factor in the etiology of hypertension and diabetes, we would anticipate that hypertension, diabetes, and stoutness share normal pathophysiologic pathways as well as regular defenselessness qualities. The pestilence like ascent in the commonness of weight establishes an undoubted and genuine worldwide medical issue. Critically, hypertension and diabetes are habitually connected with weight and, together, establish a noteworthy weight as far as patients' bleakness and raising social insurance costs. At the point when considered in seclusion, stoutness, hypertension, and diabetes are altogether connected with expanded danger of the advancement of cardiovascular and renal confusions; be that as it may, the conjunction of this triumvirate creates a generous rise in ailment hazard. The main thrusts connecting corpulence, hypertension, and diabetes stay to be explained due, to some extent, to the unpredictable and multifactorial nature of the conditions that include blends of ecological, hereditary, way of life, and social confounders. Furthermore, it is perceived that neuroendocrine instruments, including insulin obstruction,

thoughtful apprehensive enactment, and incitement of the renin-angiotensin-aldosterone framework (RAAS), are likewise included.

This exceptional issue on hypertension and type 2 diabetes identified with weight incorporates a few epidemiological investigations concentrating on the predominance of metabolic disorder, type 2 diabetes, and hypertension. New information exuding from Peru, Ethiopia, Sudan, Egypt, and Nepal records the commonness of cardiometabolic ailment in these nations as being like that detailed in westernized nations, for example, the USA, Canada, Australia, and European nations just as Japan. Critically, a rising assortment of information, for example, that introduced by Teacher C. Brufani et al., features the significance of weight in kids. They announced the commitment of birth weight to focal fat station and insulin affectability in metabolic disorder in corpulent Italian youngsters. The test will be in planning and executing successful methodologies to capture and switch this example.

The solid linkage among hypertension and type 2 diabetes was investigated by Educator E. Senior member which gives knowledge into the instruments included. Dr. S. Horita et al. checked on the commitment of the kidneys, particularly renal sodium transport, to the advancement of insulin opposition and hypertension in weight. Diabetic patients and fat people every now and again present with various circadian examples of circulatory strain contrasted with nondiabetic or nonobese subjects. A nondipping example is normal in stout hypertensive patients. In diabetic patients, mobile circulatory strain checking gives a progressively strong measure in anticipating future cardiovascular occasions than facility pulse. Dr. C. Anigbogu et al. given proof that in rodents the circadian mood of circulatory strain and pulse changes with movement of diabetes. Teacher K. Eguchi checked on late epidemiological examinations in diabetes and stoutness utilizing mobile circulatory strain observing. Taken together, these perceptions exhibit the significance of walking pulse observing.

The main line of treatment for the treatment of sort 2 diabetes and corpulence related hypertension is weight reduction with way of life adjustments, for example, diet and exercise. Nonpharmacological medicines were laid out by Dr. J. Pappachan et al. Another article by S. Fellow et al. shown that video gaming gave some advantage in starting way of life alterations that supported in weight reduction. Patients with diabetes and hypertension often present with atherogenic illnesses and dyslipidemia. Dr. E. Spirits Villegas and partners gave an audit

showing that statins are extremely powerful in treating dyslipidemia and lessening cardiovascular hazard.

This unique issue secured a wide scope of materials with an attention on sort 2 diabetes and hypertension. Articles included the study of disease transmission, physiology, and medications. In synopsis, this issue showed that (I) stomach heftiness is identified with the high pervasiveness of hypertension and type 2 diabetes paying little heed to ethnicity, (ii) insulin opposition is a noteworthy system connecting the beginning and improvement of hypertension and type 2 diabetes, and (iii) weight reduction with eating regimen and exercise is a significant viewpoint in treating hypertension in sort 2 diabetes and helps in expanding the viability of antihypertensive meds. Further examinations on instruments and hereditary qualities are required to create proper and powerful restorative regimens to avert and restrain corpulence related diseases, for example, hypertension and type 2 diabetes. Early mediation is imperative, given rising proof of end-organ brokenness in youthful overweight or large people.

#### **Inflammation and Oxidative stress**

A second-rate incendiary procedure happens in both diabetes and hypertension. Indeed, even constant periodontitis is a dormant factor in the advancement of diabetes, hypertension, cardiovascular maladies, and the metabolic disorder. Somehow or another, diabetes and hypertension could be considered as ceaseless fiery infections.

Incendiary markers (e.g., C-responsive protein (CRP)) are expanded in patients with diabetes, hypertension, and the metabolic disorder, and anticipate the advancement of these illnesses. The neighborhood renin-angiotensin-aldosterone framework (RAAS) assumes a significant job in vascular pathophysiology. Angiotensin-changing over catalyst (Expert) is communicated in the shoulder of coronary supply route plaques. Angiotensin II (Ang II) is to an enormous degree in charge of activating vascular irritation and actuating oxidative pressure. It invigorates NADH/NADPH oxidase, and enacts Rho/Rho kinase, protein kinase C (PKC), and mitogen-initiated protein kinase (MAPK). Additionally, Ang II down-manages proinflammatory interpretation factors, for example, atomic factor-κB (NF-κB), bringing about the age and emission of receptive oxygen species (ROS), incendiary cytokines (e.g., interleukin-6 [IL-6]), chemokines, and attachment particles. These activities lead to endothelial brokenness and vascular damage.

Quality administrative system examination has uncovered oxidative worry as a key fundamental sub-atomic component in diabetes and hypertension. The oxidative pressure intervened guideline course is the normal robotic connection among the pathogenesis of diabetes, hypertension, and other related incendiary maladies.

Peroxisome proliferator-enacted receptor (PPAR) activators lower circulatory strain, prompt positive consequences for the heart, and enhance endothelial brokenness through cell reinforcement, calming, antiproliferative, antihypertrophic, and antifibrotic impacts, Ang II down-controls the mRNA and protein of PPAR-α and PPAR-γ, bringing about the decrease of PPAR mitigating limit and initiation of aggravation. PPAR-α and PPAR-γ activators have been exhibited to apply cardiovascular defensive impacts free of their metabolic activities. Be that as it may, ongoing investigations with double PPAR activators have cast questions on their clinical viability in cardiovascular anticipation contrasted and the first PPAR activators right now advertised.

Conventional pharmacologic methodologies, for example, statins, ACE inhibitors, and Ang II receptor blockers (ARBs), which lessen cardiovascular occasions in randomized clinical preliminaries, likewise diminish vascular aggravation in patients with diabetes and hypertension. Enhancement of way of life (e.g., weight reduction, exercise, Mediterranean-style diet) additionally has the impact of diminishing vascular irritation.

#### **Insulin Resistance**

Insulin is a pleiotropic hormone that assumes a crucial job in the improvement of hypertension, diabetes, and the metabolic disorder. The primary metabolic activities of insulin are to invigorate glucose take-up in skeletal muscle and heart and to smother the generation of glucose and exceptionally low-thickness lipoprotein (VLDL) in the liver. Under fasting conditions, insulin emission is smothered, prompting expanded glucose blend in the liver and kidneys (gluconeogenesis) and expanded transformation of glycogen to glucose in the liver (glycogenolysis). After a supper, insulin is discharged from pancreatic  $\beta$ -cells and restrains gluconeogenesis and glycogenolysis. Insulin invigorates the thoughtful sensory system (SNS) to build cardiovascular yield and the conveyance and use of glucose in the fringe tissues. Other metabolic impacts of insulin incorporate restraint of glucose discharge from the liver, hindrance of the arrival of free unsaturated fats (FFAs) from fat tissue, and incitement of the procedure by which amino acids are consolidated into protein.

Insulin opposition, a condition wherein abandons in the activity of insulin are with the end goal that typical degrees of insulin don't trigger the sign for glucose retention, signifies a weakened reaction to insulin in skeletal muscle, liver, fat, and cardiovascular tissue. Insulin obstruction emerges because of different hereditary, procured, and ecological components, including corpulence. Expanded RAAS exercises may likewise cause insulin opposition through the incitement of Ang II type 1 receptors, which trigger expanded generation of responsive oxygen species (ROS) in adipocytes, skeletal muscle, and cardiovascular tissue of corpulent people. FFAs are accepted to prompt insulin opposition and increment the degree of oxidative pressure, bringing about endothelial brokenness and atherogenesis.

Insulin obstruction is related with hindered insulin flagging, impeded fibrinolysis, and irritation. Rising proof proposes that insulin obstruction may result from variations from the norm in key atoms of the insulin-flagging pathways, including overexpression of phosphatases and downregulation or potentially actuation of protein kinase falls, prompting irregularities in the articulation and activity of different cytokines, development variables, and peptides, and overproduction of VLDL. Insulin opposition may likewise bring about hindered fibrinolysis, which is portrayed by hypercoagulability and height of fibrinogen and plasminogen activator inhibitor (PAI)- 1. PAI-1 movement is raised in a wide assortment of insulin opposition patients. Indeed, even in patients with ordinary glucose resistance, raised degrees of fasting insulin are related with hindered fibrinolysis. Hence, insulin obstruction is a prothrombotic state portrayed by a rise of PAI-1 and fibrinogen levels, prompting expanded danger of cardiovascular occasions. Insulin obstruction might be a consequence of an overproduction of proinflammatory cytokines (e.g., IL-6, tumor rot factor (TNF), and CRP) and an overall insufficiency of mitigating cytokines (e.g., adiponectin) delivered from fat tissues because of heftiness.

Insulin-intervened glucose take-up by muscle changes more than six-crease in obviously sound people, with around half of the inconstancy in insulin activity being hereditarily decided and the other half coming about because of contrasts in the level of adiposity and physical wellness. Most patients with sort 2 diabetes are insulin safe, and about portion of those with basic hypertension are insulin safe. In this way, insulin opposition is a significant regular connection among diabetes and hypertension.

#### **Mental Pressure and Thoughtful Sensory system**

Stressors are inherent or extraneous boosts prompting unsettling influences in physiology and brain research and may undermine wellbeing. Contrasted and physical stressors, present day stressors emerging from mental risk (e.g., work pressure, abusive behavior at home, and cataclysmic events) are increasingly continued. Perpetual mental pressure, coming about because of the cutting-edge way of life, is much of the time related with physiologic and mental aggravations, and may by implication lead to diabetes and hypertension.

Albeit epidemiologic examinations have exhibited that psychological pressure is related with hypertension, cardiovascular infection, weight, and the metabolic disorder (which incorporates diabetes as a part). The impact of mental weight all in all body isn't totally comprehended. Creature analyses instructed us that the components incorporate renal thoughtful nerve movement (RSNA) and circulatory strain control in which baroreflex capacity is included.

In the human body, incitement of the thoughtful sensory system (SNS), brought about by unending pressure, lifts heartbeat rate and cardiovascular moment yield and initiates the RAAS, which is another significant pressor component. Expanded movement of the SNS likewise has an influence in the improvement of disabled glucose and lipid digestion. Considering the SNS and RAAS enables us to comprehend their jobs in the etiology and treatment of hypertension, metabolic disorder, and diabetes.

There is likewise a connection between mental pressure and stoutness in patients with diabetes and hypertension. A high pervasiveness of hypertension in fat subjects has been identified with psychosocial factors, including perpetual pressure. The hypothalamicpituitary–adrenal pivot was proposed as a key instrument connecting stoutness, hypertension, and ceaseless pressure. Hence, individuals ought to diminish worry to escape from the endless loop of mental pressure, weight, diabetes, and hypertension.

#### **Physical Action**

In the Da Qing Disabled Glucose Resistance and Diabetes Study, episode diabetes diminished by 46% in the activity gathering. In the nonrandomized Malmö Plausibility Concentrate in 260 moderately aged men with weakened glucose resistance, the occurrence of diabetes was half lower in the intercession bunch following 5 years. In the Finnish Diabetes Anticipation Study, subjects with an adjustment in moderate-to-fiery recreation time physical movement (LTPA) in the most elevated tertile were 49% to 65% less inclined to create diabetes than those in the least tertials. In the Coronary Conduit Hazard Improvement in Youthful Grown-ups contemplate (CARDIA) with more than 15 years of development, there was a huge 17% decrease of danger of episode hypertension for each 300-practice unit increase in normal physical action. In the Atherosclerosis Hazard in Networks (ARIC) consider, the most elevated quartile of recreation movement (principally cycling and strolling) had 34% lower chances of creating hypertension more than 6 years contrasted with the least dynamic. Accordingly, physical action diminishes the danger of creating diabetes and hypertension. The system includes changes in body weight and glucose resilience, just as different elements.

The impact of corpulence weakness qualities on the beginning of heftiness is affected by physical movement in the person. The genotypic impact of FTO is more articulated in inert than dynamic people. The previous is bound to convey chance alleles, for example, rs9939609. All things considered, people meeting the day-by-day physical action proposals may defeat the impact of FTO genotype on weight related infections, for example, diabetes, hypertension, and the metabolic disorder.

The potential advantages of physical action in the counteractive action and treatment of diabetes and hypertension are very much perceived however normal physical action is troublesome and here and there difficult to do, all things considered. General wellbeing endeavors should by the by still expect to raise open mindfulness and encourage normal physical action to avoid against diabetes, hypertension, and other related sicknesses.

#### **CONCLUSION**

Diabetes and hypertension share regular pathways, for example, SNS, RAAS, oxidative pressure, adipokines, insulin opposition, and PPARs. These pathways communicate and impact one another and may even reason an endless loop. Hypertension and diabetes are both final products of the metabolic disorder. They may, in this manner, create consistently in a similar person. Focal weight is the reason for the metabolic disorder. Just "ORLISTAT" is presently accessible for the long-haul treatment of stoutness. Accordingly, advancement of way of life remains the foundation in the anticipation and treatment of diabetes and hypertension.

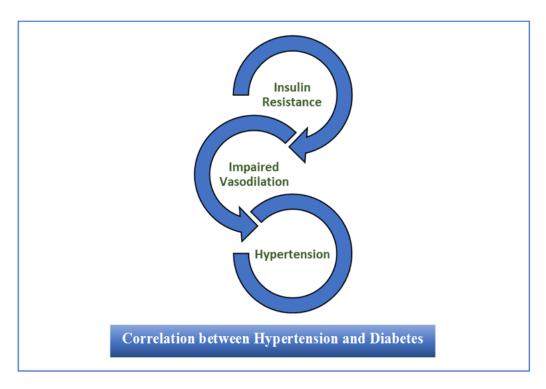


Figure 2:

#### ACKNOWLEDGEMENT

Authors would like to thank all the staff members, Especially **Dr Shilpa Pandita** (Deputy Medical Superintendent) at "Shri Mata Vaishno Devi Narayana Superspeciality Hospital," Kakryal-Katra, Jammu, Jammu and Kashmir for guiding us and to get this project done.

#### **REFERENCES**

- 1. Cheung BM. The hypertension-diabetes continuum. J Cardiovasc Pharmacol, 2010; 55: 333–9. This is a brief review of the overlap between hypertension and type 2 diabetes that proposes there is a spectrum ranging from hypertension without deglycation to type 2 diabetes without elevated blood pressure.
- 2. Landsberg L, Molitch M. Diabetes and hypertension: pathogenesis, prevention and treatment. Clin Exp Hypertens, 2004; 26: 621–628. doi: 10.1081/CEH-200031945.
- 3. Gress TW, Nieto FJ, Shahar E, et al. Hypertension and antihypertensive therapy as risk factors for type 2 diabetes mellitus. Atherosclerosis Risk in Communities Study. N Engl J Med, 2000; 342: 905–912. doi: 10.1056/NEJM200003303421301.
- 4. Cheung BM, Wat NM, Tso AW, et al. Association between raised blood pressure and dysglycemia in Hong Kong Chinese. Diabetes Care, 2008; 31: 1889–1891. doi: 10.2337/dc08-0405.

- 5. Zeggini E, Scott LJ, Saxena R, et al. Meta-analysis of genome-wide association data and large-scale replication identifies additional susceptibility loci for type 2 diabetes. Nat Genet, 2008; 40: 638–645. doi: 10.1038/ng.120.
- 6. Sober S, Org E, Kepp K, et al. Targeting 160 candidate genes for blood pressure regulation with a genome-wide genotyping array. PLoS One, 2009; 4: e6034. doi: 10.1371/journal.pone.0006034.
- 7. Ong KL, Tso AW, Leung RY, et al. A genetic variant in the gene encoding adrenomedullin predicts the development of dysglycemia over 6.4 years in Chinese. Clin Chim Acta, 2011; 412: 353–357. doi: 10.1016/j.cca.2010.11.007.
- 8. Ong KL, Jiang CQ, Liu B, et al. Association of a genetic variant in the apolipoprotein A5 gene with the metabolic syndrome in Chinese. Clin Endocrinol (Oxf), 2011; 74: 206–213. doi: 10.1111/j.1365-2265.2010.03899. x.
- 9. Cheung CY, Tso AW, Cheung BM, et al. Genetic variants associated with persistent central obesity and the metabolic syndrome in a 12-year longitudinal study. Eur J Endocrinol, 2011; 164: 381–388. doi: 10.1530/EJE-10-0902.
- 10. Ong KL, Li M, Tso AW, et al. Association of genetic variants in the adiponectin gene with adiponectin level and hypertension in Hong Kong Chinese. Eur J Endocrinol, 2010; 163: 251–257. doi: 10.1530/EJE-10-0251.
- 11. Ong KL, Leung RY, Wong LY, et al. Association of a polymorphism in the lipin 1 gene with systolic blood pressure in men. Am J Hypertens, 2008; 21: 539–545. doi: 10.1038/ajh.2008.21.
- 12. Ong KL, Wong LY, Man YB, et al. Haplotypes in the urotensin II gene and urotensin II receptor gene are associated with insulin resistance and impaired glucose tolerance. Peptides, 2006; 27: 1659–1667. doi: 10.1016/j.peptides.2006.02.008.
- 13. Chow WS, Cheung BM, Tso AW, et al. Hypoadiponectinemia as a predictor for the development of hypertension: a 5-year prospective study. Hypertension, 2007; 49: 1455–1461. doi: 10.1161/HYPERTENSIONAHA.107.086835.
- 14. Tso AW, Sham PC, Wat NM, et al. Polymorphisms of the gene encoding adiponectin and glycaemic outcome of Chinese subjects with impaired glucose tolerance: a 5-year follow-up study. Diabetologia, 2006; 49: 1806–1815. doi: 10.1007/s00125-006-0324-2.
- 15. Ng MC, So WY, Lam VK, et al. Genome-wide scan for metabolic syndrome and related quantitative traits in Hong Kong Chinese and confirmation of a susceptibility locus on chromosome 1q21-q25. Diabetes, 2004; 53: 2676–2683. doi: 10.2337/diabetes.53.10.2676.

- 16. Ng MC, So WY, Cox NJ, et al. Genome-wide scan for type 2 diabetes loci in Hong Kong Chinese and confirmation of a susceptibility locus on chromosome 1q21-q25. Diabetes, 2004; 53: 1609–1613. doi: 10.2337/diabetes.53.6.1609.
- 17. Ross KA. Evidence for somatic gene conversion and deletion in bipolar disorder, Crohn's disease, coronary artery disease, hypertension, rheumatoid arthritis, type-1 diabetes, and type-2 diabetes. BMC Med, 2011; 9: 12. This article showed that somatic gene conversion could be a significant causative factor in seven diseases, including hypertension, type 1 diabetes, and type 2 diabetes. There are some common SNPs identified in some of the seven diseases.
- 18. Moore TR. Fetal exposure to gestational diabetes contributes to subsequent adult metabolic syndrome. Am J Obstet Gynecol, 2010; 202: 643–649. doi: 10.1016/j.ajog.2010.02.059.
- 19. Xita N, Tsatsoulis A. Fetal origins of the metabolic syndrome. Ann N Y Acad Sci., 2010; 1205: 148–55. This article argued that consequences of fetal adaptive responses might be evident later in life rather than at birth. Risk factors in pregnancy might predispose the fetus to hypertension, diabetes, or the metabolic syndrome in adulthood.
- 20. Guerrero-Romero F, Aradillas-Garcia C, Simental-Mendia LE, et al. Birth weight, family history of diabetes, and metabolic syndrome in children and adolescents. J Pediatr, 2010; 156: 719–23, 723 e1.
- 21. Sowers JR. Insulin resistance and hypertension. Am J Physiol Heart Circ Physiol, 2004; 286: H1597–1602. doi: 10.1152/ajpheart.00026.2004.
- 22. Davy KP, Hall JE. Obesity and hypertension: two epidemics or one? Am J Physiol Regul Integr Comp Physiol, 2004; 286: R803–813. doi: 10.1152/ajpregu.00707.2003.
- 23. He YH, Jiang GX, Yang Y, et al. Obesity and its associations with hypertension and type 2 diabetes among Chinese adults age 40 years and over. Nutrition, 2009; 25: 1143–9. This was a cross-sectional study of over 5000 people in the community in Shanghai showing that obesity was associated with a higher risk of both hypertension and type 2 diabetes.
- 24. Loos RJ, Bouchard C. Obesity-is it a genetic disorder? J Intern Med, 2003; 254: 401-425. doi: 10.1046/j.1365-2796.2003.01242. x.
- 25. Maes HH, Neale MC, Eaves LJ. Genetic and environmental factors in relative body 1997; 27: weight and human adiposity. Behav Genet, 325–351. doi: 10.1023/A:1025635913927.
- 26. Ingalls AM, Dickie MM, Snell GD. Obese, a new mutation in the house mouse. J Hered, 1950; 41: 317–318.

- 27. Saunders CL, Chiodini BD, Sham P, et al. Meta-analysis of genome-wide linkage studies in BMI and obesity. Obesity (Silver Spring), 2007; 15: 2263–2275. doi: 10.1038/oby.2007.269.
- 28. Wang K, Li WD, Zhang CK, et al. A genome-wide association study on obesity and obesity-related traits. PLoS One, 2011; 6: e18939. doi: 10.1371/journal.pone.0018939.
- 29. Cheung CY, Tso AW, Cheung BM, et al. Obesity susceptibility genetic variants identified from recent genome-wide association studies: implications in a chinese population. J Clin Endocrinol Metab, 2010; 95: 1395–1403. doi: 10.1210/jc.2009-1465.
- 30. Wang T, Huang Y, Xiao XH, et al. The association between common genetic variation in the FTO gene and metabolic syndrome in Han Chinese. Chin Med J (Engl), 2010; 123: 1852–1858.
- 31. Wing MR, Ziegler JM, Langefeld CD, et al. Analysis of FTO gene variants with obesity and glucose homeostasis measures in the multiethnic Insulin Resistance Atherosclerosis Study cohort. Int J Obes (Lond), 2010.
- 32. Ng MC, Tam CH, So WY, et al. Implication of genetic variants near NEGR1, SEC16B, TMEM18, ETV5/DGKG, GNPDA2, LIN7C/BDNF, MTCH2, BCDIN3D/FAIM2, SH2B1, FTO, MC4R, and KCTD15 with obesity and type 2 diabetes in 7705 Chinese. J Clin Endocrinol Metab, 2010; 95: 2418–2425. doi: 10.1210/jc.2009-2077.
- 33. Song Y, Altarejos J, Goodarzi MO, et al. CRTC3 links catecholamine signalling to energy balance. Nature, 2010; 468: 933–939. doi: 10.1038/nature09564.
- 34. Savoia C, Schiffrin EL. Inflammation in hypertension. Curr Opin Nephrol Hypertens, 2006; 15: 152–158.
- 35. Stehouwer CD, Gall MA, Twisk JW, et al. Increased urinary albumin excretion, endothelial dysfunction, and chronic low-grade inflammation in type 2 diabetes: progressive, interrelated, and independently associated with risk of death. Diabetes, 2002; 51: 1157–1165. doi: 10.2337/diabetes.51.4.1157.
- 36. Ross R. Atherosclerosis–an inflammatory disease. N Engl J Med, 1999; 340: 115–126. doi: 10.1056/NEJM199901143400207.
- 37. Tracy RP. Emerging relationships of inflammation, cardiovascular disease and chronic diseases of aging. Int J Obes Relat Metab Disord, 2003; 27(Suppl 3): S29–34. doi: 10.1038/sj.ijo.0802497.
- 38. Tracy RP. Inflammation, the metabolic syndrome and cardiovascular risk. Int J Clin Pract Suppl. 2003; 134: 10–17.

- 39. Karnoutsos K, Papastergiou P, Stefanidis S, et al. Periodontitis as a risk factor for cardiovascular disease: the role of anti-phosphorylcholine and anti-cardiolipin antibodies. Hippokratia, 2008; 12: 144–149.
- 40. Persson GR, Persson RE. Cardiovascular disease and periodontitis: an update on the associations and risk. J Clin Periodontol, 2008; 35: 362–379. doi: 10.1111/j.1600-051X.2008.01281. x.
- 41. Watanabe K, Petro BJ, Shlimon AE, et al. Effect of periodontitis on insulin resistance and the onset of type 2 diabetes mellitus in Zucker diabetic fatty rats. J Periodontol, 2008; 79: 1208–1216. doi: 10.1902/jop.2008.070605.
- 42. Nesbitt MJ, Reynolds MA, Shiau H, et al. Association of periodontitis and metabolic syndrome in the Baltimore Longitudinal Study of Aging. Aging Clin Exp Res., 2010; 22: 238–242.
- 43. Tsioufis C, Kasiakogias A, Thomopoulos C, et al. Periodontitis and blood pressure: the concept of dental hypertension. Atherosclerosis, 2011; 219: 1–9.
- 44. Page RC. The pathobiology of periodontal diseases may affect systemic diseases: inversion of a paradigm. Ann Periodontol, 1998; 3: 108–120. doi: 10.1902/annals.1998.3.1.108.
- 45. Hung HC, Willett W, Merchant A, et al. Oral health and peripheral arterial disease. Circulation, 2003; 107: 1152–1157. doi: 10.1161/01.CIR.0000051456. 68470.C8.
- 46. Blake GJ, Rifai N, Buring JE, et al. Blood pressure, C-reactive protein, and risk of future cardiovascular events. Circulation, 2003; 108: 2993–2999. doi: 10.1161/01.CIR.0000104566. 10178.AF.
- 47. Blake GJ, Ridker PM. Novel clinical markers of vascular wall inflammation. Circ Res., 2001; 89: 763–771. doi: 10.1161/hh2101.099270.
- 48. Sesso HD, Buring JE, Rifai N, et al. C-reactive protein and the risk of developing hypertension. JAMA, 2003; 290: 2945–2951. doi: 10.1001/jama.290.22.2945.
- 49. Savoia C, Schiffrin EL. Vascular inflammation in hypertension and diabetes: molecular mechanisms and therapeutic interventions. Clin Sci., 2007; 112: 375–384. doi: 10.1042/CS20060247.
- 50. Griendling KK, Minieri CA, Ollerenshaw JD, et al. Angiotensin II stimulates NADH and NADPH oxidase activity in cultured vascular smooth muscle cells. Circ Res., 1994; 74: 1141–1148.

- 51. Yamakawa T, Tanaka S, Numaguchi K, et al. Involvement of Rho-kinase in angiotensin II-induced hypertrophy of rat vascular smooth muscle cells. Hypertension, 2000; 35: 313–318.
- 52. Taubman MB, Berk BC, Izumo S, et al. Angiotensin II induces c-fos mRNA in aortic smooth muscle. Role of Ca2+ mobilization and protein kinase C activation. J Biol Chem. 1989; 264: 526–530.
- 53. Shreya Jain, Henrita Boro, Drishti Sharma, Gurpreet Singh Multani, Sudhanshu Bansal, Saksham kumar, & Shivam Choudghal. Assessment of quality of life in diabetes and hypertensive patients attending tertiary care hospitals in (Jammu), (Jammu and Kashmir). *International Journal of Research in Pharmacology* & *Pharmacotherapeutics*, 2021; *10*(3): 236-243. https://ijrpp.com/ijrpp/article/view/397.