

# Varied Exercise Regimen and its Impact on Glucose Control among Type 2 Diabetes Mellitus Patients: A Systematic Review

R. Vivek, R. Kalidasan

## ABSTRACT

**Objectives:** Systematic evaluation of study varied exercise regimen impact on Hemoglobin A1C and fasting blood glucose (FG) for type 2 diabetes mellitus patients in which age ranged from 30 to 88 years. **Design:** Systematic review, qualitative aspects of analysis followed guideline as per Cochrane handbook for systematic review of intervention and Preferred Reporting Items for Systematic Review and Meta-Analysis. **Data sources:** The 23 randomized controlled trails out of 237 studies paper English language only on Web of Science and PubMed database, as where search place search term used since May 2018 and still April 2019. **Results:** The articles showed that implementation varied exercise regimen of physical exercise consisted aerobic, resistance, both combine form, walking, and endurance based as high-intensity interval training and co-intervention of low calorie diet, medicine, education counseling, and training supervision impose motivation and proper guideline could be glycosylated hemoglobin and FG reduction about type 2 diabetes mellitus patients. **Conclusion:** Varied exercise regimen of physical activity, specifically combine exercise such as endurance based aerobic and resisted exercise could be treated with pure medication to reduce glucose level in type 2 diabetes patients mellitus, managed along with lifestyle change the prolong time risk factors.

**Keywords:** Aerobic exercise, Diabetes, Diet and supervision, Resisted exercise

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## INTRODUCTION

International diabetes federations around worldwide survey exposure (Sep 2018) reported 2 in 3 people with type 2 diabetes mellitus, and have affected cardiovascular diseases risk factors.<sup>[1]</sup> Over 350 million adults are currently at high risk of developing type 2 diabetes. These affect almost 700 million people by 2045.<sup>[2]</sup> Systematic review like a review that has been acting orderly to fixed plan, approach to minimize errors and random errors which contain in methodological section. Critical evaluation in healthcare studies appraise based on inclusion and exclusion criteria and its qualitative aspects of analysis. Therefore, secondary's data are collected from reviews.<sup>[3]</sup> The complication of diabetes patients are cardiac problem, neuropathy, kidney damage, and retinopathy. Sedentary behavior is one of the most factors for developing type 2 diabetes which have major risk factors of obesity. The accumulation of fat that increased body sizes and reason to tissue and organs changing in their structure also not function normally. Type 2 diabetes is the most common type of diabetes, accounting for around 90% of all diabetes cases.<sup>[4]</sup> Physical activity is the contraction of skeletal muscle except diaphragm that requires energy to move and action, which is responsible for whole energy expenditure. In the technology world people living with too much eating, and sitting behavior such as as television watching, computer, and game using. It would be getting extra fat in human body and also physiological effect may lead chronic disease. Physical exercise is increased insulin-sensitivity and glucose uptake to show in healthy individual. The core important of exercise intervention to prevent and/or treatment, management of diabetes type 2 patients. Delivery of glucose increased and transported to myocytes, and oxidative enzyme activities. Furthermore, the capacity for glycogen synthesis increases with physical training.<sup>[5]</sup> Early intervention, ongoing support, and adherence to lifestyle changes are essential factors

Department of Physical Education, Bharathidasan University, Tiruchirappalli, Tamil Nadu, India

**Corresponding Author:** R. Vivek, Department of Physical Education, Bharathidasan University, Tiruchirappalli, Tamil Nadu, India. E-mail: viveknamakkalpd@gmail.com.

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such as exercise and diet with treating type 2 diabetes patients. The core management strategy of diabetes patients alone with physical exercise and its suggestion from health professional for glycemic control in individuals with type 2 diabetes patients. As per reported American College of Sports Medicine, American diabetes association and joint position statement that on physical exercise have been conducted weekly 150 min exercises, 3 days training program per week but not consecutive 2 days and combined each sessions follow approximately 30–60 min.<sup>[6]</sup> The meta-analysis that demonstrated a large effect on glycosylated hemoglobin (Hba1c) reduction in type 2 diabetes mellitus patients in studies of resistance exercise consisted with relatively high number of sets per bout of exercises. The study implication could be significant for establishing an ideal exercised protocol for resistance training and prescribing tailor-made exercise programs for patients' diabetes. Authors specify that the Hba1c reduction differed according to the resistance training program was consisted such as frequency, intensity or number of sets.<sup>[7]</sup> The meta-analysis found that lifestyle intervention showed significant benefit in risk factors which are

known to be associated with the development of cardiovascular disease of type 2 diabetes mellitus. Meta-analysis included 16 trails reported that HbA1c values were significantly favored the intervention groups compared with control groups thus intervention such as exercise, diet, and education.<sup>[8]</sup> Systematic literature review showed that exercise intervention (different type of exercise) in patients with type 2 diabetes mellitus resulting improved metabolic parameter as glucose control and also in molecular adaptation that enhance cardiac function in most cases.<sup>[9]</sup> The objectives of systematic review were to study the impact of varied exercise regimen such as aerobic exercise, resistance exercise and both combine form, walking, endurance based, and strength training which can control blood glucose level and HbA1c among type 2 diabetes patients.

## METHODS

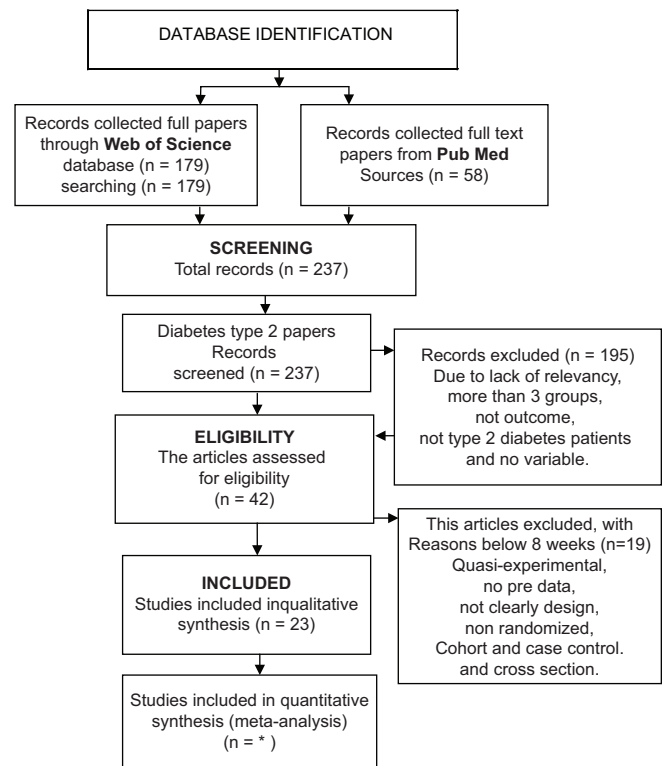
### Data Sources and Study Selection

The studies searched two database names as Web of Science and PubMed. Web of Science is one of the largest international multidisciplinary which is each subject's area covering with best journal. It was started since 1900, currently maintained by Clarivate Analytics. The database acquires world-wide leading citation in various disciplines. User obtain only online subscription basis, and freely available for research purpose in education institute of Bharathidasan University, Tamil Nadu, India. PubMed database opened 24 years ago, provide information about life science and clinical area by National Library of Medicine at United States, National Institute of Health. The sources of PubMed are easily available on open access abstract and full text paper with reference through search engine. All the information of full text paper collected largest databases.

### Search Strategy

Search term; "physical activity and diabetes mellitus not yoga," "physical exercise and DM type 2," "exercise and HbA1c," "training and glucose," "aerobic exercise and fasting glucose," "strength training and DM," "walking and FBG or fasting blood glucose (FG)," "fitness and FBS," "jogging and DM," "workout and type 2 DM," "A1c and resistance exercise," "practices and FG," and DM\* and training. Those which were used Boolean, phrases, and truncation search strategies.

The search term used that opted 179 full text papers in Web of Science in which description, 29 papers have not pre- or post-data outcome concerned to objective of systematic review and data were presents median with inter quartile, 77 have not related to age, exercises groups-quasi experimental design, study more than 3 groups, 36 have been no variables relationship of HbA1c and fasting blood glucose, 11 paper other language than English, and nine paper have non-randomized controlled trails. Eight have been cohort, case series, and case-control study. Then, nine studies finally selected. PubMed 58 full text papers follows descriptions, eight papers have not either pre- or post-data value, twenty papers not relevant to research study, 14 studies have not inclusion criteria variables, two has non-randomized controlled trails paper, and then 14 study selected meet inclusion criteria, as shown in the Figure 1. The totally setting 23 randomized controlled trails inclusion out of 237 full text papers review by



**Figure 1:** Flow chart of the result of literature search. \*implies, this is not conducted meta analysis

first author and final selected second author records have among both databases about type 2 diabetes mellitus patients and those online search started since May 2018 and still April 2019 year. Inclusion criteria randomized controlled trails exercise conducted community and/or home based and/or clinical setting in various countries. Inclusion meet 23 papers published year range 1992 to current year. This study systematic review Guideline following the Preferred Reporting Items for Systematic Review and Meta-Analysis, and International database of Prospero prospectively registered our study ID No. CRD42019155268. ([https://www.crd.york.ac.uk/prospero/display\\_record.php?ID=CRD42019155268](https://www.crd.york.ac.uk/prospero/display_record.php?ID=CRD42019155268)). Further flow chart diagram that prepared hierarchy as per the study.<sup>[10]</sup>

### Eligibility Criteria

The first important in the systematic review study decision prepares questions, that well clear idea and how to seek answers. Including eligibility criteria focused address research question on specific type of studies. This study follows as participations, interventions, comparisons, and outcomes and it could be helped pre-specified criteria.

Participants: Type 2 diabetes mellitus patients included diabetes along with obesity and/or overweight, and some diabetes mellitus patients of the participants with comorbidities. Male and female (gender) are study subjects which were ranged age 30–88 criteria selected to inclusion of the study. Intervention: Physical exercise or varied exercise regimen either particular training or combine exercise consisted such as aerobic, resistance, and both combine exercise and endurance based interval training or walking exercise, that have form two or three groups:

Study cointervention diet, and/or medication and education. Comparison: Intervention against which have been compared with no treatment group or placebo group of standard care group or conventional group or active control intervention group or shame group. These all groups' intervention variation compared to physical activity and exercise groups intervention. For example, dosage, intensity, mode, frequency and duration and acting on exercise, medicine, and diet. Outcomes: Randomized controlled trails, studies depended measuring variables of outcome as FG, and hemoglobin A1c or Hba1c which contain papers opted for inclusion of the study. Training setting conducted community center, home, and hospital. Article inclusion published only English. The exclusion criteria have non-randomized controlled trails, case series, and case-control study. Yogic exercise exclusion and exercise below 8 weeks considered exclusion as age below 29 years not considered.

## Data Collection and Analysis

### Data extraction

The data extraction by two dependent review first and second author from randomized controlled included trails and it could be reported to minimize error, information involved results of interpretation used Microsoft office-excel worksheet. The inclusion characteristics were listed such as author/year, country, study design, age in year (Mean,  $\pm$ SD), and sample size/groups male/female, Co-Intervention, Treatment, and Control/Active Control groups were documented in the Table 1. Training protocol was shown author/year, exercise, control groups, intensity, and duration which included session, repetition, set and interval then follow as outcome time point and outcome measures are presented in Table 2.

### Measure of Intervention Effect

The study pre-decided outcome measure was selected variables FG and hemoglobin A1C, to which measures of intervention mostly used enzymatic colorimetric methods, glucose oxides methods, glucometer, and chromatography. Other used a few methods measured in clinical laboratory.

### Quality Assessment of Trails

A bias is a systematic error, meaning that multiple processes making replica of the same study would lead the wrong answer on average. The validity of study might have considered two dimensions, first dimension provided whether asking appropriate research question to systematic review study and lead external validity. Second dimension, purpose of the systematic review, it correct answer its research question that is frees from bias and lead internal validity. The internal validity assessment is often referred as assessment of methodological quality. Risk of bias in randomized controlled studies was methodological quality assessed by seven domain bias such as random sequence generation (selection bias), allocation sequence concealment (selection bias), blinding of participants and personal (performance bias), blinding of outcome assessment (detection bias), incomplete outcome data (attrition bias), selecting outcome reporting (reporting bias), and other potential sources bias, as shown in Table 3. Then, categories

related judgments of each entry assigning which were the risk of bias as "low risk," "high risk," and 'unclear risk of biases. Assessing risk of bias in included randomized controlled studies, for guideline execution Cochrane hand book for systematic review of intervention in the chapter 8.<sup>[11]</sup>

## Methodics of Selection of Studies

In this research, type 2 diabetes patients with exercise intervention related full text articles collected from Web of science (179) and Pub-Med (58). Those 237 articles were screening totally in which 195 articles excluded due to lack of relevancy. After that 42 articles were assessing for eligibility in which excluded 19 articles that did not meet inclusion criteria. Remaining 23 articles selected for systematic review.

## RESULTS AND DISCUSSION

The 23 randomized controlled trails selected out of 237 full text diabetes type 2 patients. In this varied exercise regimen included studies result which would aerobic or resistance exercises and six trails both combined form, nine studies were aerobic training which is tradition or/and formal exercise including walking three studies opted. Resistance training three trails, early form of exercise keyed up with weight or without weight, resistance band used exercise. Endurance based exercise two studies of interval training. Reviewers often narrow inclusion criteria to deal with heterogeneity by including only those studies reporting a particular outcome. This was kept as guideline for this research writing.<sup>[12]</sup> In this systematic review has 23 studies but only 12 studies (29,13, 25,31,17,19,16,12,18,15,30,4) were mentioned on type 2 diabetes patients were properly diagnosed as per guideline of world health organisation and/or international diabetes federation and/or American diabetes association and other 11 studies were not mentioned clearly. Inclusion randomized controlled trails were 8 weeks and above following varied exercise regimen of training, Aerobic training<sup>[13-20,4]</sup> of nine studies were significant difference among FG and Hba1c values. Those studies only<sup>[13,16,4]</sup> were not significant that reason low intensity level trails<sup>[13,4]</sup> compare groups took low calorie but despite of low glucose level reduced among type 2 diabetes patients. The study confirms that weight reduction induced by diet that could affect glucose metabolism. Meta-analysis showed that aerobic interval training, which was at low volume and convey larger cardio-metabolic benefit than moderate intensity of continuous training of type 2 diabetes mellitus, in particular for cardio-respiratory fitness and glucose control.<sup>[21]</sup> Short time side effect could be prevented by prescribed medication but long time complication on diabetes patients delayed by regular physical activity and proper diet following that positive effect on aerobic exercise above 8 weeks.<sup>[15]</sup> Inclusion<sup>[22,23]</sup> high-intensity interval training based on endurance exercise and session per day 20 min were conducted to reduce glycemic level.<sup>[12]</sup> Unsupervised 12 weeks high-intensity interval training reduced in Hba1c of 2.8 mmol/mol (-0.20%) compared to control groups.<sup>[24]</sup> Trails of meta-analysis, a high-intensity protocol combined exercises were associated with BP reduction of type 2 diabetes mellitus and structured exercise consisted more than 150 min/week.<sup>[25]</sup> Resistance training<sup>[22,26,27]</sup> consisted such as functional and traditional exercises had no difference and not furnished session with mention already suitable glycemic level. Study 12 months progressive resistance training, power based equipment

**Table 1:** Characteristics of included study

S. No	Author/Year	Country	Study design	Age in Year (Mean ± SD)	Sample size groups (M/F)	Co-Intervention	Treatment	Control/active con
1	Rech et al., 2019 <sup>(27)</sup>	Brazil	Parallel group randomized control design	≥60 year Ex-70.5±7.4 Sham-68±6.5	38/Ex-18, Co-21 (m20, F18)	None	Resistance training; 12 weeks and 3 days per week	Active con; stretching workout once per week
2	Botton et al., 2018 <sup>(22)</sup>	Brazil	Parallel group randomized control design	60–88 year Ex-70.6±6.7 Co-68.6±7.06	26/Ex-13, Co-13	Diet	Resistance training; 12 months and 3 days per week	Stretching low intensity once per week
3	Cassidy et al., 2019 <sup>(24)</sup>	England	Parallel group randomized control design	60±2 year Ex-60±3 Co-59±3	22/Ex-11, Co-12(M17, F6)	None	High-intensity interval training; pedal cadence and band resisted exercise 12 weeks, 3 days a week	Standard care
4	Stefano et al., 2010 <sup>(28)</sup>	Italy	Parallel group randomized control design	(<60 vs. ≥60) year Ex-58.8 (8.5) Co-58.8 (8.6)	563/Ex-288, Co-275 (M329, F234)	Diet ± oral agent, life style change	Intensive exercise; 12 weeks and 3 days per week	Standard care alone Counseling
5	Rahbar et al., 2017 <sup>(17)</sup>	Iran	Parallel group randomized control design	40–60 year Ex-48.31 (5.02) Co-48.60 (4.80)	28/Ex-13, Co-15	None	Aerobic exercise; treadmill 8 weeks and 3 days per week	Normal routine activity
6	Motahari-Tabari et al., 2014 <sup>(5)</sup>	Iran	Parallel group randomized control design	30–65 year Ex-49.29 (1.12) Co-49.0 (1.60)	53/Ex-27, Co-26(F53)	Diet	Aerobic exercise; stretching and flexibility 8 weeks and 3 days per week	Normal routine activity
7	Shakil-ur-Rehman et al., 2018 <sup>(18)</sup>	Pakistan	Parallel group randomized control design	30–70 year Ex-(M) 59.12±05.78 (F) 51.31±5.78 Co-(m) 55.00±8.03 (F) 57.93±06.83	102/Ex-51, Co-51 (M55, F47)	Routine medicine and diet plan	Aerobic exercise; SSAET program 25 weeks, 3 days per week	Routine medicine and diet plan
8	Praet et al., 2008 <sup>(16)</sup>	Netherlands	Parallel group randomized control design	60±9 year Bric-61±9 Madi-59±9	37/Bric-18, Medi-19	Diet guideline	Brisk walking; 12 months, 3 days per week	Medical fitness program; 12 months, 3 session per week
9	Karstoft et al., 2013 <sup>(23)</sup>	Denmark	Three group randomized control design	lwt-57.5±2.4 Cwt-60.8±2.2 Co-57±3.0	32/lwt-12, Cit-12 Co-8(M20, F12)	Diet -CMG	IWT; Walking, Target energy expenditure, 4 months, 5 session per week CWT; Slow walking Target energy expenditure, 4 months, 5 session per week First step Program; walking, 16 weeks, 3–4 days per week	Habitual routine lifestyle Standard care
10	Tudor-Locke et al., 2004 <sup>(29)</sup>	Canada	Parallel group randomized control design	40–60 year Ex-52.8±5.7 Co-52.5±4.8	47/Ex-24, Co-23(M26, F21)	PA behaviour change, Education	Aerobic exercise; cycle ergo meter, 16 weeks, 4 session per week	Very low calorie diet
11	Snel et al., 2012 <sup>(19)</sup>	Netherlands	Parallel group randomized control design	Vex-53.0±2.5 Vlcd-56.1±2.4	27/VlcdEx-13, Vlcd-14 (M14, F13)	Diet	Aerobic exercise; treadmill 12 weeks and 4 times per week	Low intensity PA
12	Kadoglou et al., 2012 <sup>(14)</sup>	Greece	Parallel group randomized control design	50–70 year Ex-58.4±5.7 Co-62.9±4.2	53/Ex-26, Co-27(M19, F34)	Diet	Endurance type exercise program; High-intensity, 6 months, 3 session per week	Low intensity exercise
13	Hansen et al., 2009 <sup>(31)</sup>	Belgium	Parallel group randomized control design	59±8 year Hi-59±2 Li-58±1	37/ExHi-19, Li-18(M37)	None	Brisk walking; 16 weeks, 4 days per week	No exercise, maintain with usual activities
14	Kadoglou et al., 2010 <sup>(30)</sup>	Greece	Parallel group randomized control design	50–65 year Ex-56.83±6.76 Co-60.32±9.28	47/Ex-23, Co-24(M15, F32)	None		

(Contd...)

**Table 1: (Continued)**

S. No	Author/Year	Country	Study design	Age in Year (Mean ± SD)	Sample size groups (M/F)	Co-Intervention	Treatment	Control/active con
15	Kang <i>et al.</i> , 2016 <sup>[31]</sup>	South Korea	Vgroup randomized	Ex-56.0±7.4 Co-57.5±4.6	16/Ex-8, Co-8(F16)	None	Aerobic exercise; walking and resistance exercise, 12 weeks, 3 times per week	Standard care
16	De Lade <i>et al.</i> , 2016 <sup>[32]</sup>	Brazil	control design Parallel group randomized	St-57±12 At-54±9	11/ExSt-5, Aero-6(M5, F6)	None	Aerobic training; 20 weeks, 3 weekly session	Strength training given after aerobic exercise
17	Mavros <i>et al.</i> , 2013 <sup>[36]</sup>	Australia	control design Parallel group randomized	Older than 60 year	84/Ex-36, Sham-48	None	PRT; high-intensity exercise, 12 months, 3 days per week	Sham exercise; low intensity usual care
18	Anniballini <i>et al.</i> , 2017 <sup>[33]</sup>	Italy	control design Parallel group randomized	55-70 year Ex-57±9.1 Co-60±6.8	16/Ex-8, Co-8(M16)	None	Aerobic and strength exercise; 16 weeks, 2-3 times a week	Usual care
19	Taghizadeh <i>et al.</i> , 2018 <sup>[20]</sup>	Iran	control design Parallel group randomized	62.25±3.81 Yr	20/Ex-10, Co-10(F20)	None	Endurance training; treadmill, 8 weeks, 3 session per week	Not involve any kinds of exercise training
20	Liu <i>et al.</i> , 2015 <sup>[34]</sup>	China	control design Parallel group randomized	Ex-52.59±11.43 Co-51.20±11.34	42/Ex-22, Co-20(M19, F23)	None	Aerobic and resistance training; 12 weeks, 2-3 session per week	Conventional and drug therapy and psychological counseling
21	Winding <i>et al.</i> , 2018 <sup>[12]</sup>	Denmark	control design Three group randomized	Hiit-54±6Yr End-58±8Yr Co-57±7	32/Hiit-13, End-12, Co-7(M19, F13)	Diet	HIIT; cycling, 11 weeks, 3 session per week Endurance training; cycling, do same week and session	Energy intake low compare with 2 groups
22	Yang <i>et al.</i> , 2017 <sup>[35]</sup>	Canada	control design Three group randomized	52±1.2 year RT1-52.2±1.2 RT2-49.8±1.4 RT3-54.6±1.2	51/RT1-16, RT2-17, RT3-18(M30.F21)	Diet	Resistance and Aerobic training; RT2-High-intensity, 6 months, 5 days a week RT3-Endurance, low intensity, do same above	RT1-usual care
23	Vanninen <i>et al.</i> , 1992 <sup>[4]</sup>	Finland	Parallel group randomized control design	40-60 year male-53±7 Female-54±6	78/Ex-38, Co-40(M45, F33)	Diet	Aerobic exercise; walking, jogging, 12 months, 3-4 times week	Conventional treatment

Ex: Exercise group, Co: Control group, SD: Standard deviation, M/F: Male/Female, SSAET: Supervised structured aerobic exercise training, CGM: Continuous glucose monitor, IWT: Interval walking training, CWT: Continuous walking training, PRT: Progressive resistance training, PA: Physical activity, HIIT: High-intensity interval training

**Table 2:** Training protocol of included study

S. No	Author/Year	exercise	Control/active control	Supervision	Intensity	Duration/session/ Repetitions/Sets/ intervals	Outcome time point	Outcome measures
1	Rech <i>et al.</i> , 2019 <sup>[27]</sup>	RT program included Function ex and tradition ex squat and bench stepping, knee flex and extension and abdominal crunches	Stretching workout with low intensity- static move for large muscle groups	Yes	F.c- Omni Scale R.E-15 repetition maximum	12 weeks, 3 days a week, (R) F.Ex 10-15 T.Ex 10-12, (s) F.Ex and T.Ex 2-3, (l) 1-1.30 min	0 and 12 weeks	Hba1c, FG, Cholesterol
2	Botton <i>et al.</i> , 2018 <sup>[22]</sup>	RT program; strength ex on Tradition machine, free weight and F.Ex squat and steps up and down	Joint mobilization and static stretching for large muscle groups, perform low intensity	NR	F.c- Omni Scale (0 to 10)	12 weeks, 3 days a week, (R) F.Ex 10-15 T.Ex 10-12,(s) F.Ex and T.Ex 2-3, (l) 1-1.30 min	0 and 12 months	hba1c, FG, Cholesterol, muscle strength, sit-to-stand, timed up and go RTD 50&200
3	Cassidy <i>et al.</i> , 2019 <sup>[24]</sup>	High-intensity interval training; the rating exertion (RPE) increased forms 9-13 comfortable Cadence. And resistance band upper body exercise was light	Standard care and routine normal, not change their medicine, PA, diet	Yes	Borg Scale (9-17) RPE	12 weeks, 3 days a week, (l) 3 min	0 and 12 weeks	Hba1c, weight, BMI, BP, Sv, Co, Heart rate variability
4	Stefano <i>et al.</i> , 2010 <sup>[28]</sup>	Intensive ex; 150 min/Weeks 2 supervised session of progressive mixed (aerobic and resistance) training-treadmill, cycle ergo-meter	Standard care alone counseling	Yes	11.25 MET-h/ wk (vo2 max)	12 months, weekly twice, (5-8)	0 and 12 months	Hba1c, FG, Upper and lower body Strength, homa-IR, Bp, cholesterol, wc, BMI, OHAS, insulin, Antihypertensive agents, lipid
5	Rahbar <i>et al.</i> , 2017 <sup>[17]</sup>	Aerobic ex; treadmill with no slope. bruce protocol and its increased gradually	Normal routine activity	Yes	50-70% max heart rate	8 weeks, 30 min session, 3 days per week	0 and 8 weeks	Hba1c, FG, ht, wt, BMI, EF%, cholesterol, BP, micro albumin
6	Motahari-Tabari <i>et al.</i> , 2014 <sup>[15]</sup>	Aerobic ex; stretching and flexibility ex 50 min daily exercise (Brick walking 30 min)	Standard care (normal routine activity)	Yes	60% max heart rate	8 weeks, 50 min per session, 3 times a week	0 and 8 weeks	FG, Plasma insulin, hip-c, wait.c, BMI, wt
7	Shakil-ur-Rehman <i>et al.</i> , 2018 <sup>[18]</sup>	Aerobic ex; supervised structured aerobic ex treadmill training inclination (0-3) divided 5 phases with 5 week duration	Keep routine medication and diet plan	NR	Borg Scale, vo2 max by the resting heart rate	25 weeks, 20-50 min per session, 3 days a week,	0 and 25 weeks	Hba1c, FG, plasma insulin level, Insulin resistance, HDL, LDL, Interleukin-6, nitric oxide, cyclooxygenase, RPE, dyspnea level, BMI, Vo2 max
8	Praet <i>et al.</i> , 2008 <sup>[16]</sup>	Brick walking; the endurance type ex consisted of 5-6 Km/h and resistance ex, floor ex own body weight or elastic bands	Medical fitness ex; endurance type ex consisted of interval ex on home trainer and 8 difference ex target upper& lower body	NR	brick- 75 ± 5% MF-73 ± 2% max heart rate	12 months, 60 min per session, 3 days a weekly	0 and 12 months	Hba1c, FG, BMI, HOMA, resting heart rate, Bp

(Contd...)

Table 2: (Continued)

S. No	Author/Year	exercise	Control/active control	Supervision	Intensity	Duration/session/Repetitions/Sets/intervals	Outcome time point	Outcome measures
9	Karstoft et al., 2013 <sup>[23]</sup>	Interval walking training; JD Mate, used as a pedometer, group had the target energy expenditure rate set for 70% with speed walking. Continuous walking training; target energy expenditure rate set at 55% with slow walking	Habitual life style and JD mate pedometer data uploaded monthly	Yes	IWT: Peak energy ex 70% low to high CWT; 55% peak energy ex moderate intensity	4 months, 60 min per session, 5 days a week,	0 and 4 months	Hba1c, FG, vo2 max, Body com, lipids, Bp, CGM glucose
10	Tudor-Locke et al., 2004 <sup>[29]</sup>	First step programs; pedometer use walking hour while energy used activities goal setting self-monitoring	Group received post card for their participation	NR	PA by approximately 3000-12000 step/day	16 weeks, 30 min session, 3-4 days a week	0 and 16 weeks	Hba1c, FG, cholesterol, 2-h glucose, w.c, h.c, Heart rate, Bp
11	Snel et al., 2012 <sup>[19]</sup>	Adding ex; training at home for 30 min and a 1 h hospital training aerobic exercise + low calorie diet	Very low calorie diet approximately 450 Kcal/day	Yes	70% max capacity cycloergometer	16 weeks, 30 min session, 4 days a week	0 and 16 weeks	Hba1c, FG, Bp, cholesterol, average insulin, BMI, Waist cm
12	Kadoglou et al., 2012 <sup>[14]</sup>	Aerobic exercise; walking or running on treadmill, cycling, calisthenics of upper and lower limbs	Walking 3-5 times per week with low intensity	Yes	60-75% max heart rate	12 weeks, 45-50 min per session, 4 times a week	0 and 12 weeks	Hba1c, FG, Bp, cholesterol, BMI, Waist cm
13	Hansen et al., 2009 <sup>[13]</sup>	Endurance type exercise; walking, cycling, cross-country and sky type exercise. High-intensity exercise 40 min, 118 ± 3 beat per min	Low intensity 55 min, 105 ± 3 heart beat	Yes	HI-75% vo2 Peak moderate to high LI-50% vo2 Peak low to moderate 50-70%	6 months, 40-55 min session, 3 session per week	0, 2 and 6 months	Hba1c, FG, Blood lipid, body com, vo2 max
14	Kadoglou et al., 2010 <sup>[30]</sup>	Brick walking; goal 150 min/week of self controlled moderate intensity	No exercise instruction maintain usual habitual activities	Yes	Aerobic 60% HRR Resista-60-80%	16 weeks, 30-60 Min per session, 4 days a week	0 and 16 weeks	Hba1c, FG, Bp, W/H ratio, insulin, cholesterol, vo2 peak
15	Kang et al., 2016 <sup>[31]</sup>	aerobic = resistance ex; treadmill 30 min, and resistance use weight machine (upper and lower body)	Standard care	NR	50-70%	12 weeks, 60 min per session, 3 times a week, (R) 7-12(S) 2	0 and 12 weeks	Hba1c, FG, Wc, insulin cholesterol, Bp, c-reactive protein, HOMA_IR, C peptide
16	De Lade et al., 2016 <sup>[32]</sup>	Aerobic ex; treadmill ex, bikes, elliptical and upper body cycle ergometer	Strength training: rowing squat, bench press, rowing with dumb, bells, shoulder ex, dumbbells and circuit exercise	Yes	80% Bore scale rate (15-18)	20 weeks, 50-60 min per session, 3 times weekly, (R) 12-15 (S) 2-3	0, 10 and weeks	Hba1c, FG, PP glycemc, cholesterol, seric insulin, HOMA-IR
17	Mavros et al., 2013 <sup>[26]</sup>	Progressive resistance training; High-intensity, pneumatic resis Equipment power training was concentric and eccentric for 4 s	Low intensity with usual care	Yes	aerobic 85% vo2 max Resistance - 45-65% Hr	12 months, 3 days a week, R-8, S-3	0 and 12 months	Hba1c, HOMA2-IR, body com, skeletal muscle mass, Vat, mid thigh muscle attenuation
18	Annibalini et al., 2017 <sup>[33]</sup>	Aerobic ex; treadmill (walking) & strength ex- horizontal, leg puss & pull down, lat machine, chest press	Usual diabetics care	Yes	aerobic 85% vo2 max Resistance - 45-65% Hr	16 weeks, A 5-10 min, Resis 30-60 min, 2-3 times a week, R 15-20, S 2-4	0 and 16 weeks	Hba1c, FG, BMI, hc, 1-RM, Bp, cholesterol

(Contd...)

**Table 2:** (Continued)

S. No	Author/Year	exercise	Control/active control	Supervision	Intensity	Duration/session/ Repetitions/Sets/ intervals	Outcome time point	Outcome measures
19	Taghizadeh et al., 2018 <sup>[20]</sup>	Endurance type exercise; moderate intensity treadmill walking or running	Not involve any kinds of training	NR	60–75%	8 weeks, 35–50 min per session, 3 session a week	0 and 8 weeks	Hba1c, FG, HOMA -IR, vo2 max, body fat, BMI
20	Liu et al., 2015 <sup>[94]</sup>	Aerobic ex; home based ex start with slow and low intensity, Resis ex elastic band (borg scale 11–14) High-intensity interval training; 11 week bicycle intervention consisting of 20 min/session and 95% of W peak Endurance training; 11 week bicycle intervention consisting of 40 min/session and 50% of W peak	Convention and drug therapy, psychological counseling Energy intake low Compare to both groups	Yes	A 40–60% R 50–60%	12 weeks, 3 session per week, R-2-5 times a week, Re 8–10, S-2	0 and 12 weeks	Hba1c, FG, cholesterol, HOMA-IR, Fin, Pbg
21	Winding et al., 2018 <sup>[12]</sup>	Endurance RT low repetition 75% of 1 RM RT3; Endurance RT low intensity high repetition 50% of 1 RM Aerobic ex; walking, jogging, cycling, cross-country, occupation and recreational activity	Compare Energy intake low Compare to both groups	NR	HIIT- cycling of 1 min at 95% W peak ENDU-cycling at 50% W peak	11 weeks, 3 session per week, In-1 min	0 and 11 weeks	Hba1c, FG, vo2 max, Body com, lipids, CGM
22	Yang et al., 2017 <sup>[95]</sup>	Endurance RT low repetition 75% of 1 RM RT3; Endurance RT low intensity high repetition 50% of 1 RM Aerobic ex; walking, jogging, cycling, cross-country, occupation and recreational activity	RT1; low intensity, high repetition lowest ex volume 50% of 1 RM Standard care	Yes	R2-High-intensity (7-RM) R3-Low Intensity (15-RM) 30-60%	6 months, 5 days a week, R2-(Re) 7, R3-(Re) 15, S 2–3	0 and 6 Months	Hba1c, FG, cholesterol, vo2 max, BMI, hc, body fat, lean mass
23	Vanninen et al. 1992 <sup>[4]</sup>	Aerobic ex; walking, jogging, cycling, cross-country, occupation and recreational activity	Standard care	Yes	30-60%	12 months, 3–4 times a week, 30–60 min per session,	3, 0, 12 months	Hba1c, FG, FP insulin, cholesterol, max oxygen uptake, serum triglycerides

RT: Resistance training, F.Ex: Functional exercise, T.Ex: Traditional exercise, Hba1c: Glycosylated hemoglobin, FG: Fasting blood glucose, RPE: Rating of perceived exertion, NR: Not report, Sv: Stroke volume, Co: Cardiac output, Bp: Blood pressure, PR: Peripheral resistance, Ex: Exercise, PA: Physical activity, HOMA-IR: Homeostasis model assessment, OHAS: Oral hypoglycemic agents, BMI: Body mass index, HT: Height, WT: Weight, EF: Max, ejection fraction, Hip.C: Hip circumference, Wait. C: Waist circumference, HDL: High-density lipoprotein, LDL: Low-density lipoprotein, CGM: Continuous glucose monitor, HRR: Heart rate reserve, 1RM: Maximum repetition, FIN: Fasting insulin



**Table 3:** Risk of bias assessment in included study as per Cochrane bias tool

S. No	List of studies	Random sequence generation bias	Allocation concealment bias	Blinding of participants and performance bias	Blinding of outcome assessment bias	Incomplete Outcome Data bias	Selecting reporting bias	Other sources of bias
1	Rech <i>et al.</i> , 2019 <sup>[27]</sup>	Low	Low	Low	Low	Unclear	Low	Low
2	Botton <i>et al.</i> , 2018 <sup>[22]</sup>	Low	Unclear	Unclear	Low	Unclear	Low	Low
3	Cassidy <i>et al.</i> , 2019 <sup>[24]</sup>	Low	Unclear	Unclear	Low	Low	Low	Low
4	Stefano <i>et al.</i> , 2010 <sup>[28]</sup>	Low	Low	Low	Low	Low	Low	Low
5	Rahbar <i>et al.</i> , 2017 <sup>[17]</sup>	Low	Unclear	High	Low	Low	Low	Low
6	Motahari-Tabari <i>et al.</i> , 2014 <sup>[15]</sup>	Low	Unclear	Unclear	Unclear	Low	Low	Low
7	Shakil-ur-Rehman <i>et al.</i> , 2018 <sup>[18]</sup>	Low	Unclear	High	Unclear	Low	Low	Low
8	Praet <i>et al.</i> , 2008 <sup>[16]</sup>	Low	Low	Low	Low	Low	Low	Low
9	Karstoft <i>et al.</i> , 2013 <sup>[23]</sup>	Low	Unclear	Low	Low	Low	Low	Low
10	Tudor-Locke <i>et al.</i> , 2004 <sup>[29]</sup>	Low	Unclear	Low	Unclear	Low	Low	Low
11	Snel <i>et al.</i> , 2012 <sup>[19]</sup>	Low	Unclear	Low	Low	unclear	Low	Low
12	Kadoglou <i>et al.</i> , 2012 <sup>[14]</sup>	Low	Unclear	High	Unclear	Low	Low	Low
13	Hansen <i>et al.</i> , 2009 <sup>[13]</sup>	Low	Unclear	High	Unclear	Low	Low	Low
14	Kadoglou <i>et al.</i> , 2010 <sup>[30]</sup>	Low	Unclear	High	Unclear	Low	Low	Low
15	Kang <i>et al.</i> , 2016 <sup>[31]</sup>	Low	Unclear	Unclear	Unclear	Low	Low	Low
16	De Lade <i>et al.</i> , 2016 <sup>[32]</sup>	Low	Unclear	High	Unclear	Low	Low	Low
17	Mavros <i>et al.</i> , 2013 <sup>[26]</sup>	Low	Unclear	Low	Low	Low	Low	Low
18	Annibalini <i>et al.</i> , 2017 <sup>[33]</sup>	Low	Unclear	High	Unclear	Low	Low	Low
19	Taghizadeh <i>et al.</i> , 2018 <sup>[20]</sup>	Low	Unclear	Unclear	Unclear	Low	Low	Low
20	Liu <i>et al.</i> , 2015 <sup>[34]</sup>	Low	Unclear	Unclear	Unclear	Low	Low	Low
21	Winding <i>et al.</i> , 2018 <sup>[12]</sup>	Low	Unclear	Unclear	Unclear	Low	Low	Low
22	Yang <i>et al.</i> , 2017 <sup>[35]</sup>	Low	Unclear	Low	Low	Unclear	Low	Low
23	Vanninen <i>et al.</i> , 1992 <sup>[4]</sup>	Low	Unclear	Low	Low	Unclear	Low	Low

used high-intensity of 80% could have significant difference with compare groups. Meta-analysis trails showed that resistance band smaller effect on Hba1c and found in meta-analysis in participants with type 2 diabetes mellitus. The resistance band studies were training period 12–16 weeks in duration, with a little reduction glucose, but free weight or weight machine used duration of 3 and 6 months have showed significant reduction. The resistance band training was not assessing resistance precisely to ensure optimal progression to higher loads. However, free weight or weight machine exact quantities of weight were assessed. However, resistance band training was not important effect, there is reason to key difference at studies used free weight and weight machine, and has contributed to this lesser reduction in Hba1c.<sup>[36]</sup> Endurance based interval training<sup>[24,12]</sup> exercise, this reviews based 12 weeks training high-intensity interval exercise well difference compared groups on Hba1c and FG, but minimum improvement in cardiac function. Therefore, intensity 50–60%, three sessions weekly with 40–60 min and considered exercise supervision. The walking training three studies<sup>[30,23,29]</sup> nine and ten were showed that first step program is an effect physiological adaptation to type 2 diabetes mellitus patients with cardiac problem. Karstoft study was three groups, one had interval walking training reduced hyperglycemic level but there were no significance different and post-intervention parameter was measured at least 48 h after and up 7 days. Measures would have to lead under estimation of benefits and three groups exercise pre-post values of Hba1c and FG were almost reduced, but difference had not compared two intervention groups except conventional group. Clinical messages that walking training frequency should be fairly high on average 3–5 times per week, in session lasting from 20 to 60 min, to be effective in improving walking performance.<sup>[37]</sup> Walking exercise patients had persistent motive to goal follow supervised sessions and strictly diet control would be significant. The capacity for glycogen synthesis increases with physical training.<sup>[5]</sup> Type 2 diabetes patients’ two group comparisons which have larger

effect size associated with exercise prescription, fitness testing, supervised exercise, group session, and recommendations for longer duration exercise session.<sup>[38]</sup> The aim of Italian diabetes and exercise study was assessed whether a strategy combining prescribed and supervision mixed (aerobic and resistance) training program alone with structured exercise counseling impact lead to Hba1c level and cardiovascular risk to reduce.<sup>[28]</sup> American diabetes association as per 150 min/week of moderate intensity activity corresponding to MET- h/wk and new information, this study highly intensive exercise combine form (aerobic and resistance training) strategy effective Hba1c with lifestyle change which was supervised that superior continuous reinforcement counseling. The study found that combined resistance and aerobic regimens improved glycemia control to greater extent than either aerobic or resistance alone.<sup>[39]</sup> The study 6 months longer training of combines both aerobic and resistance training impact was feasible weekly supervised intensity 50–75% (1-RM) and volume 2–3 sets and 7–15 repetitions advantage an AT routine before initiating RT.<sup>[35]</sup> The review shown that the implementation of physical activity programs (aerobic, resistance, flexibility, and combined exercise), based on nutrition with low glycemia index, calorie restriction, and education session about type 2 diabetes mellitus patients improve blood glucose level.<sup>[40]</sup> The clinical favorable impact of diabetes self-management education on reduction in glycogenic control is critical important because glycemic control is among the strongest predictors of disease.<sup>[41]</sup> Aerobics and resistance both combine<sup>[31-35,28]</sup> form exercises above 8 weeks of exercise with 80% intensity, 40–60 per session and 3 days a week that might be significant difference in hemoglobin a1c among compared group. However, 8 weeks low to moderate intensity or weekly twice, 30–40 min exercises were not significant difference in compared group although within pre-post FG to reduce minimum. Authors’ statement that systematic review indicates the evidence the effectiveness of resistance exercise compared to aerobic exercise was no difference in values of Hba1c and cholesterol; however, it

was increase in  $vo_{2\max}$  favoring of type 2 diabetes mellitus.<sup>[42]</sup> In the systematic review, core study analysis might be combine exercise possible with prescribed medicine, diet, and supervision. The meta-analysis found that lifestyle intervention showed significant benefit in risk factors to which associated with development of cardiovascular disease of type 2 diabetes mellitus. Meta-analysis included 16 trails reported data were Hba1c significantly favored the intervention groups compared with control groups thus intervention such as exercise, diet, and education.<sup>[8]</sup>

### Strength and Limitation

The systematic review, impact of intervention such as aerobic exercise, resistance exercise, and combine form (strength and endurance based) of exercises, Hba1c, and FG control among type 2 diabetes mellitus patients. Therefore, combine basis exercises were significant difference on glucose reduced than the resistance exercise no reduction appropriated glucose level and co-intervention of diet, medication, and education counseling. Supervision and exercise proper guideline gave motivation on diabetes patients to additional impose training program. Exercise before resistance exercise, began with endurance based exercise. All the exercises program various duration from 8 weeks to 12 months to which vary age groups from 30 to 88 years. This systematic review, study conducted. and based on Web of Science and PubMed database obtained paper only and varied country place where as different geographical setting. Some of the randomized controlled trails adverse affect among type 2 diabetes mellitus patients, who were absent following training program. The study has 25% to loss the follow-up of training program.<sup>[28]</sup>

### CONCLUSION

The worldwide burden of diabetes mellitus is highly and growing that which contributors to morbidity and mortality. The study conclusion that high-intensity interval exercise were conducted 12 weeks programs following 3–4 session for more than 30 min weekly and 3 days per week training was significance reduction in Hba1c and FG. High-intensity exercise sufficient duration for post-exercise energy expenditure, aerobic exercise of moderate to vigorous intensity (60–80% max heart rate). Resistance training result was less significant energy expenditure than aerobic exercise. Combine training of aerobic and resistance exercise, the intensity of at least 60% of  $HR_{peak}/HR_{max}$  or 55% of  $vo_{2\max}$  for aerobic exercise and 50% of 1RM for resistance training generally showed, session must followed 40–60 min statistically significant result. This study suggestion starting aerobic exercises before resisted exercise would be significant benefits. Study can be combine (endurance and strength) form best result for diabetes type 2 patients with proper regular guideline exercise, supervision, diet following by dietician and education counseling might be occurred Hba1c and FG reduced significantly. It could minimize the risk of cardiovascular disease.

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Area clarified with specialists' person emerging answer on varied exercise regimen impact on glucose control for type 2 diabetes mellitus patients.

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