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Research Article





Dual-Antiplatelet Therapy vs Single Antiplatelet Therapy after Coronary Artery Bypass Grafting : Meta-Analysis of Randomized Controlled Trails

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Abstract

Introduction: Every year, around 400,000 patients in the United States and Europe undergo CABG surgery. The risks and benefits of antiplatelet regimens in these patients have been studied in various trials. In patients with a stable clinical state, aspirin monotherapy is the recommended standard of care. The present systematic literature review and meta-analysis is aimed to compare the efficacy between Dual Antiplatelet Therapy (DAPT) and Single Antiplatelet Therapy (SAPT) after CABG.

Method:We searched PubMed/Medline, EMBASE and Cochrane Central Register of Controlled Trials (CENTRAL) for relevant published Randomized Controlled Trials (RCTs). In total, 2,152 studies from PubMed, Science Direct and Scholar were obtained and screened for the relevance of our topic for the last 10 years (2014 to 2024), in which 18 studies passed after the inclusion and exclusion criteria using PRISMA.

Result: We included 18 randomized controlled trial studying 218,125 patients. DAPT was associated with significantly lower rates of Cardiovascular mortality (OR 0.71, 95% CI 0.54, 0.93, p=0.01), Major Adverse Cardiovascular And Cerebrovascular Events (MACCE) (OR 0.68, 95% CI 0.61, 0.76, p=0.0001), and Myocardial Infarction (OR 0.78, 95% CI 0.64, 0.94, p=0.009). However, there was no significant association with graft occlusion (OR 0.86, 95% CI 0.68, 1.10, p=0.23), stroke (OR 0.82, 95% CI 0.63, 1.45, p=0.82), All-cause Death (OR 1.06, 95% CI 0.61, 1.84, p=0.84) and major bleeding (OR 1.21, 95% CI 0.81, 1.81, p=0.0001).

Conclusion: In conclusion, there were no significant differences in the incidence of graft occlusion, all-cause death, stroke, or serious bleeding between Dual Antiplatelet Therapy (DAPT) and Single Antiplatelet Therapy (SAPT). The p-values exceeded the 5% significance level. However, there were significant differences in the outcomes of cardiovascular mortality, MACCE, and myocardial infarction, with p-values showing statistical significance at a level of less than 5%.

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Keywords: Aspirin; Antiplatelet; Coronary Artery Bypass Grafting; Cardiovascular myocardial infarction; Coronary artery bypass graft surgery; Dual antiplatelet therapy; Graft occlusionclopidogrel; Graft occlusion; Major bleeding; Mortality; Prasugrel; Ticagrelor; Ticagrelor; Stroke

Introduction

Coronary Artery Bypass Grafting (CABG) has been proven to be an efficacious intervention for individuals suffering from severe coronary artery disease [1]. For patients with CABG, aspirin is advised as a basic secondary preventive drug to preserve the advantages of revascularization and avoid significant adverse cardiovascular events [2]. Nevertheless, individuals who undergo coronary artery bypass grafting (CABG) still face a significant likelihood of experiencing serious ischemic cardiac and cerebrovascular events following the surgery. This risk can surpass 10% within the initial 6 to 12 months post-operation [3]. The diminished postoperative sensitivity to aspirin, along with increased platelet activation and blood clot formation, leads to a state of systemic hypercoagulability and early graft failure. These elements have been recognized as crucial contributors in this specific circumstance [4,5]. Each year, about 400,000 individuals in the United States and Europe receive coronary artery bypass grafting (CABG). The risks and benefits of antiplatelet regimens in these patients have been the focus of numerous trials [6]. It is recommended to use aspirin alone as the usual treatment for patients who are in a stable clinical condition. In the past ten years, multiple trials have examined the possible benefits of dual antiplatelet treatment (DAPT) compared to using only aspirin in terms of effectiveness and safety outcomes [7,8]. A meta-analysis conducted by [9] indicated that patients who present with ACS (Acute Coronary Syndrome) may experience favorable outcomes from Dual Antiplatelet Therapy (DAPT) following Coronary Artery Bypass Grafting (CABG). This data aligns with previous research on patients with Acute Coronary Syndrome (ACS) who received medical treatment with Dual Antiplatelet Therapy (DAPT), specifically clopidogrel in combination with aspirin [10]. Additional investigations explicitly examined the effectiveness of dual Antiplatelet Therapy (DAPT) compared to aspirin in maintaining the openness of bypass grafts following Coronary Artery Bypass Grafting (CABG). These trials found that DAPT resulted in higher rates of bypass graft patency, as reported by [11,12].

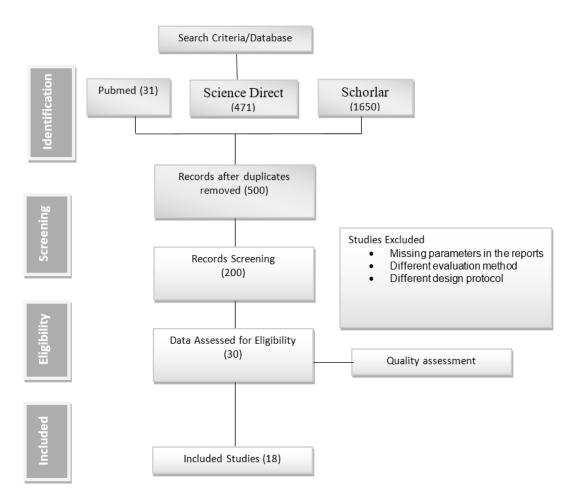
Dual Antiplatelet Therapy (DAPT) using aspirin and a P2Y12 receptor antagonist (such as clopidogrel or ticagrelor) has been shown to enhance the antiplatelet effect. This therapy has

been reported to slow down the progression of native coronary stenosis and improve graft patency in patients who have undergone Coronary Artery Bypass Grafting (CABG). Additionally, DAPT can help prevent recurrent strokes in patients with ischemic cerebrovascular disease. Nevertheless, the inquiry into whether the advantages linked to DAPT, particularly the potential enhancement in graft patency, result in superior clinical outcomes has not been thoroughly examined and has yielded inconsistent findings [13-15]. Administering aspirin within 48 hours after Coronary Artery Bypass Grafting (CABG) significantly decreased the risk of death, Myocardial Infarction (MI), and stroke during the patient's hospital stay (Mangano et al., 2002). These findings led to other trials involving the use of Dual Antiplatelet Treatment (DAPT) with clopidogrel and aspirin. However, the results regarding clinical outcome and graft patency were contradictory. Ticagrelor bypasses the need for the two-step CYP-dependent metabolism that hindered clopidogrel's effectiveness. The PLATO research demonstrated that ticagrelor is more effective than clopidogrel in lowering cardiovascular events [16]. People who had Coronary Artery Bypass Grafting (CABG) and then started taking the study drug again after the surgery had a lower risk of cardiovascular death and overall mortality when they were treated with ticagrelor [17]. Historically, aspirin has consistently been the preferred medication for preventing graft occlusion and severe cardiac events during coronary artery bypass grafting [2]. The Clopidogrel vs. Aspirin in Patients at Risk of Ischemic Events trial [18] showed that clopidogrel alone was more effective than aspirin alone in lowering the occurrence of ischemic stroke after CABG surgery. It is thought that adding a P2Y12 inhibitor will help keep the graft open and lower the risk of bad cardiac events by stopping the platelet-driven progression of graft disease. However, the effectiveness of Dual Antiplatelet Therapy (DAPT) in preserving graft openness and reducing negative cardiac events is not clearly demonstrated [2,19,20]. Hence, this present study compare the effectiveness of Dual Antiplatelet Therapy (DAPT) and Single Antiplatelet Therapy (SAPT) with Aspirin after CABG.

Research Methodology

This review will adhere to the Preferred Reporting Items for Systematic Reviews and Meta-Analyses standards. There were several stages to the method. Prior to developing the inclusion and exclusion criteria, the methodology was developed. Then, using the criteria of the abstract, title, and full text, we searched databases for research. Data extraction was followed by a synthesis of the earlier literature. A flowchart showing the stages is shown in Figure 1.

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Research Protocol

The protocol was created in accordance with the Cochrane Handbook for Systematic Review and it established the primary research questions that directed the selection of papers to search for, the data sources to use in the search, the inclusion and exclusion criteria, and the outcomes. The study questions served as the foundation for this evaluation, and they were used as a guide to finding high-quality papers and resources across several databases.

Inclusion and Exclusion Criteria

In view of the present study, only studies that addressed the research topic Monotherapy (Aspirin) Versus Dual Antiplatelet

Therapy (Prasugrel vs Ticagrelor) Following Coronary Artery Bypass Grafting were taken into consideration. The inclusion papers were randomized control design studies, studies published in the last 10 years (2014-2024) and only the studies written in English were taken into consideration in order to provide the most valid literature review possible. Systematic literature review papers, studies, and conference papers are excluded criteria.

Research Question

The research question was developed using PECOS (Participant, Exposure, Comparator, Outcome and Study Design) framework (Table 1). This framework is used to inform the search string and the entire search strategy.

PICO								
Р	Participant	Patients with coronary artery disease, CABG						
Ι	Intervention	dual antiplatelet therapy with either aspirin + clopidogrel, +ticagrelor or prasugrel)						
С	Comparative	single antiplatelet therapy (SAPT) with Aspirin						
		All-cause mortality, Cardiovascular mortality,						
O Outcome		Major adverse cardiac and cerebrovascular events (MACCE),						
		myocardial infarction, stroke, or major bleeding, Graft occlusion						

Table 1: PESCOS framework for research question

Using the above framework, this review aim to answer one research questions we trying to answer from this research was : "Is Single Antiplatelet Therapy (SAPT) better in comparison to dual antiplatelet therapy after CABG in preventing graft occlusion aswell as other secondary prevention".

Data Source and Search strategy

Reporting Items for Systematic Reviews and Meta-Analysis (PRISMA), we use PubMed, Science Direct and EMBASE and Cochrane Central Register of Controlled Trials (CENTRAL) for relevant published randomized controlled trials (RCTs) to get the studies that were used for this systematic review from year 2014 to 2024 following the inclusion and exclusion criteria earlier mentioned. The following precise keywords were used in the literature search: ("single Antiplatelet Therapy" OR "Aspirin Monotherapy") AND ("Dual Antiplatelet Therapy" OR Prasugrel OR Ticagrelor) AND ("Coronary Artery Bypass Grafting" OR "heart bypass surgery")). Table 2

Database	Number of study
Science Direct	471
Google Shorlar	1650
PubMed	31
Total	2152

Table 2: Number of hit generated from database

Quality Assessment or Risk of Bias

Considering the nature of this research, quality assessment of the eligible studies was performed using Cochrane risk of bias assessment via review manager (RevMan 5.4). Each of the study was appraised based on the CASP checklist for qualitative studies (Table 3). The appraisal was basically "Low, Moderate and High" each study was asses. The checklist criteria of assessment were 7 questions. The risk of bias is termed low if the study has 70% and above assessment score, moderate if it have within 50% to 70% score and high if it has below 50%.

Question ID	Appraisal Questions
Q1	Random sequence generation
Q2	Allocation concealment
Q3	Blinding of participants and personnel
Q4	Blinding of outcome assessment
Q5	Incomplete outcome data
Q6	Selective reporting
Q7	Other bias

 Table 3: Quality Assessment Questions

Study Selection

The exclusion criteria for the selected studies were reported as case reports, review articles, case series and articles in that is not written in English language were excluded. Full text studies on the research topic were included. Randomized controlled trials were the study types that were taken into consideration. Each article's title and abstract were examined for inclusion and exclusion criteria.

Data Extraction and Quality Assessment

A pre-defined sheet was used to collect the following information from the included studies: research design, author's first name, year of publication, country, study design, all-cause mortality, cardiovascular mortality, Major Adverse Cardiac And Cerebrovascular Events (MACCE), myocardial infarction, stroke, or major bleeding and graft occlusion. the prisma flowchart was then used to display the final list of studies that will be included. after the data were extracted, the risk bias range was divided into three categories: low, high, and uncertain.

Data Extraction and Analysis

The data were transferred to Excel, where RevMan was used to analyze the features of the chosen studies and check for the presence of heterogeneity using the inconsistency index. The numerical data extracted from the included studies were analyzed using number, event, and total. To analyse the primary and secondary

outcomes, we adopted a random effect meta-analysis with a odd ratio OR that measured the effect size. The results was visualized using a forest plot, the plot shows the overall OR for each studies. The heterogeneity across each studies was measured using (I2). A 95% Confidence level and 5% level of significance was chosen for decision making. The null hypothesis of no significant difference between the two groups would be rejected if the estimated p - value is less than the significance level. The publication bias was assessed using funnel plot. The criteria was that, if all studies were within the funnel, and the funnel is symmetric, there is no evidence of publication bias across the included studies.

Results

Study selection and Screening process

After the keywords were entered into the aforementioned database, a total of 2,152 publications were retrieved from it. The 2,152 papers were first entered into the reference manager Endnote for review. We read through the titles and abstracts of the studies during the screening stage, eliminating any duplicates and studies that did not pertain to the aim of our study. Studies that were

concentrated on a different intervention were also disregarded. 500 publications remained after the identification and screening phases; these 30 papers underwent full-text screening before being assessed for methodology and data analysis, both of which required to be done in accordance with the current study's scope. There were 18 studies left after the eligibility procedure.

Quality Assessment Result

In total, 18 studies were included, out of which 18 studies is at low risk bias under the random sequence generation. 17 studies were of low-risk for the allocation concealment and 1 study is unclear for the allocation of concealment. 15 studies are of lowrisk for blinding of participants, 2 studies is at high risk and a study is unclear. 15 studies blinding of outcome assessment were at lowrisk and 3 studies are unclear. 13 studies are of incomplete outcome data are of low-risk, 4 study are unclear and a study is at high risk. 14 studies were of low risk of selective reporting and 4 studies are unclear of selecting reporting. Lastly, other bias associated with 17 studies are of low risk while a study associated with other bias is unclear as shown in Figure 2. The percentage of the grading is shown in Figure 3.

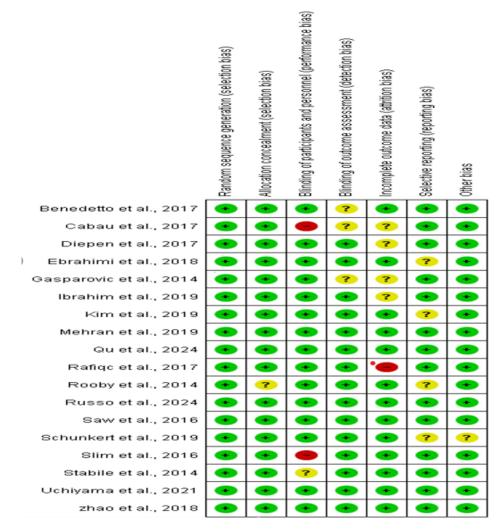


Figure 2: The quality assessment summary of the included studie

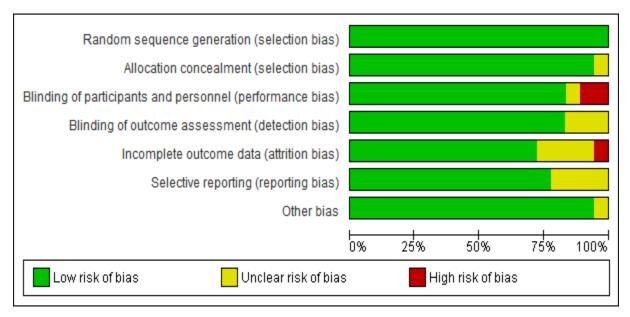
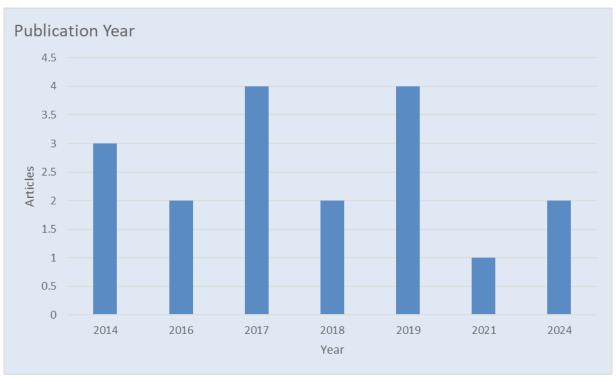
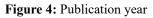


Figure 3: Percentage of the quality assessment level of the included studies

Distribution of studies with respect to year of publication

Figure 4 display the publication year of the included studies, 4 articles of the included studies was published in the year 2017 and 2019 respectively, 3 articles published in year 2014, 2 articles published each in year 2016, 2018 and 2024 respectively while a study was published in year 2021.





Meta Analysis

Data evaluation and analysis

RevMan 5.4 was used to conduct the meta-analysis. We adopted a more cautious approach and used a random-effects model because it was unknown whether there was a "true" effect size underlying all studies. The standard mean difference as well as its associated 95% Confidence Intervals (CLs) was used to estimate the effect size. The degree of heterogeneity was evaluated using the I2 test.

Data Extraction and Analysis

The data was extracted to excel and Review Manager was used to analyze the selected studies characteristics and the intervention measure also test for the heterogeneity's presence using the inconsistency index. Also was included in the forest plot is the chi statistic and the inconsistency index I2 statistics to access if the studies were heterogeneous. An indication of I2 that was less than fifty percent indicated low heterogeneity. A value between fifty and seventy-five was a sign of mild heterogeneity and above

seventy-five showed a high level of heterogeneity. Therefore, the researcher had high confidence on whether to generalize the studies to a larger sample depending on the results. Table 4, present the pooled analyses comparing DAPT with SAPT with 121004 and 97121 patients, respectively, in 18 studies. DAPT, compared with SAPT, was associated with significantly lower rates of Cardiovascular mortality (OR 0.71, 95% CI 0.54, 0.93, p=0.01), MACCE (OR 0.68, 95% CI 0.61, 0.76, p=0.0001) and Myocardial infarction (OR 0.78, 95% CI 0.64, 0.94, p=0.009). While there is no association with lower rates of graft occlusion (OR 0.86, 95% CI 0.68, 1.10, p=0.23), stroke (OR 0.82, 95% CI 0.63, 1.45, p=0.82), but high rates of All cause of death (OR 1.06, 95% CI 0.61, 1.84, p=0.84) and major bleeding (OR 1.21, 95% CI 0.81, 1.81, p=0.0001). In summary there were no significant difference in the outcomes of graft occlusion, all cause of death, stroke and major bleeding with p values greater than 5% significance level while there is significant differences in the outcome of cardiovascular mortality, MACCE and myocardial infarction with p values lesser than 5% significance level.

Outcome or Subgroup	Studies	DAPT	DAPT rate (%)	SAPT	SAPT rate (%)	Pooled OR (M-H, 95% CI)	P value	Heterogeneous test p vale	I2 statsistics (%)
Graft occlusion	7	13461	5.267	11124	7.992	0.86 [0.68, 1.10]	0.23	0.09	46
All cause of Dealth	13	18973	4.87	17225	3.64	1.06 [0.61, 1.84]	0.84	0	90
Cardiovascular mortality	8	16278	3.79	12603	2.785	0.71 [0.54, 0.93]	0.01**	0.59	0
MACCE	4	14209	7.995	8856	8.006	0.68 [0.61, 0.76]	0.001**	0.46	0
Myocardial infarction	14	22529	4.385	17736	3.822	0.78 [0.64, 0.94]	0.009**	0.18	26
Stroke	8	18451	6.216	14741	6.329	0.95 [0.63, 1.45]	0.82	1E-05	78
Major bleeding	10	17103	4.303	14836	3.013	1.21 [0.81, 1.81]	0.35	0.003	65

Table 4: Pooled Analysis results for DAPT against SAP.

P values <0.05 are asterik. DAPT, dual antiplatelet therapy; MACCE, major adverse cardiovascular and cerebrovascular events; SAPT, single antiplatelet therapy

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Figure 5: Forest plot showing the Graft occlusion of DAPT and SAPT

The statistics estimated is not statistically significant with overall odd ratio of 0.86, [95% CI: 0.68, 1.10] when p- value (0.23) is greater than the level of significant (0.05) which implies that there is no significant difference between DAPT and SAPT for across all studies. But a significant lower event rates was found. The heterogeneity, $I^{\wedge} = 46\%$ across the studies is moderate with a insignificant value (p = 0.09) indicating that the heterogeneity is not statistically significant, which makes the graft occlusion of DAPT and SAPT to be consistent across the different studies with insignificant variability.

	DAF	т	SAP	т		Odds Ratio		Odds Ratio		
Study or Subgroup	Events	Total	Events	Total	Weight	M-H, Random, 95% Cl		M-H, Random, 95% C	1	
Qu et al., 2024	1	10854	1	7215	0.7%	0.66 [0.04, 10.63]				
Rafiqc et al., 2017	17	66	15	67	7.5%	1.20 [0.54, 2.67]				
Rooby et al., 2014	135	1015	264	1802	30.5%	0.89 [0.71, 1.12]		+		
Saw et al., 2016	9	87	22	120	7.0%	0.51 [0.22, 1.18]				
Schunkert et al., 2019	512	931	486	928	33.5%	1.11 [0.93, 1.33]		•		
Slim et al., 2016	6	21	9	19	3.1%	0.44 [0.12, 1.64]				1
zhao et al., 2018	29	487	92	973	17.7%	0.61 (0.39, 0.93)				
Total (95% CI)		13461		11124	100.0%	0.86 [0.68, 1.10]		•		
Total events	709		889							
Heterogeneity: Tau ² = 0.	.04; Chi² =	: 11.08,	df = 6 (P =	= 0.09); I	² = 46%		0.005		20	
Test for overall effect: Z	= 1.21 (P	= 0.23)					0.005	Favours (DAPT) Favours (10

Figure 5: Forest plot showing the Graft occlusion of DAPT and SAPT.

Figure 6: Forest plot showing the all cause of death of DAPT and SAPT.

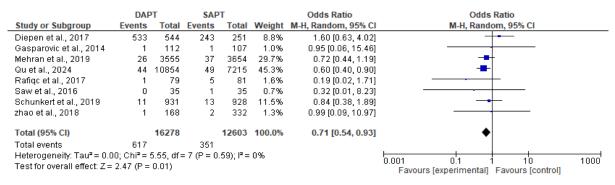
The statistics estimated is statistically not significant with overall odd ratio of 1.06, [95% CI: 0.61, 1.84] when p- value (0.84) is greater than the level of significant (0.05) which implies that the overall effect has no significant impact. But a significant high event rates was found. The heterogeneity, $I^{\wedge} = 90\%$ across the studies is high with a significant value (p < 0.00001) which makes the all cause of death of the DAPT and SAPT to be inconsistent across the different studies.

	DAP	т	SAP	т		Odds Ratio	Odds Ratio
Study or Subgroup	Events	Total	Events	Total	Weight	M-H, Random, 95% Cl	M-H, Random, 95% Cl
Benedetto et al., 2017	7	609	9	609	8.0%	0.78 [0.29, 2.09]	
Cabau et al., 2017	7	111	4	111	6.9%	1.80 [0.51, 6.33]	
Diepen et al., 2017	538	544	244	251	7.5%	2.57 [0.86, 7.73]	+- -
Ebrahimi et al., 2018	13	511	28	1014	9.3%	0.92 [0.47, 1.79]	
Gasparovic et al., 2014	2	112	4	107	5.3%	0.47 [0.08, 2.61]	
Ibrahim et al., 2019	173	2634	100	925	10.5%	0.58 [0.45, 0.75]	+
Kim et al., 2019	61	2121	96	2340	10.4%	0.69 [0.50, 0.96]	-
Mehran et al., 2019	34	355	45	3564	10.0%	8.28 [5.23, 13.12]	
Qu et al., 2024	61	10854	66	7215	10.3%	0.61 [0.43, 0.87]	
Rafiqc et al., 2017	1	79	5	81	4.0%	0.19 [0.02, 1.71]	
Russo et al., 2024	2	52	0	20	2.5%	2.03 [0.09, 44.14]	
Schunkert et al., 2019	22	931	23	928	9.6%	0.95 [0.53, 1.72]	
Stabile et al., 2014	3	60	3	60	5.5%	1.00 [0.19, 5.16]	
Total (95% CI)		18973		17225	100.0%	1.06 [0.61, 1.84]	★
Total events	924		627				
Heterogeneity: Tau² = 0.7 Test for overall effect: Z =	•		df = 12 (P	< 0.000	01); I ² = 9	0%	0.005 0.1 1 10 200 Favours [DAPT] Favours [SAPT]

Figure 6: Forest plot showing the all cause of death of DAPT and SAPT.

Figure 7: Forest plot showing the Cardiovascular mortality of DAPT and SAPT

The statistics estimated is statistically significant with overall odd ratio of 0.71, [95% CI: 0.54, 0.93] when p- value (0.01) is less than the level of significant (0.05) which implies that the overall effect has a significant impact and a significant low event rates was found. The heterogeneity, $I^{\wedge} = 0\%$ across the studies is very low with a insignificant value (p 0.59) which makes the Cardiovascular mortality of the DAPT and SAPT to be consistent across the different studies.



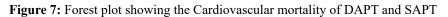


Figure 8: Forest plot showing the MACCE of DAPT and SAPT

The statistics estimated is statistically significant with overall odd ratio of 0.68, [95% CI: 0.61, 0.76] when p- value (0.00001) is less than the level of significant (0.05) which implies that the overall effect has a significant impact and a significant low event rates was found. The heterogeneity, $I^{\wedge} = 0\%$ across the studies is very low with a insignificant value (p 0.46) which makes the MACCE of the DAPT and SAPT to be consistent across the different studies.

	DAP	т	SAP	т		Odds Ratio	Odds Ratio
Study or Subgroup	Events	Total	Events	Total	Weight	M-H, Random, 95% Cl	M-H, Random, 95% Cl
Benedetto et al., 2017	34	609	34	609	4.9%	1.00 [0.61, 1.63]	
Gasparovic et al., 2014	7	112	11	107	1.2%	0.58 [0.22, 1.56]	
Ibrahim et al., 2019	783	2634	359	925	48.2%	0.67 [0.57, 0.78]	
Qu et al., 2024	312	10854	305	7215	45.7%	0.67 [0.57, 0.79]	
Total (95% CI)		14209		8856	100.0%	0.68 [0.61, 0.76]	◆
Total events	1136		709				
Heterogeneity: Tau ² = 0.0	00; Chi ² = :	2.57, df=	= 3 (P = 0	l.46); I²	= 0%		
Test for overall effect: Z =	6.94 (P <	0.00001)				0.2 0.5 1 2 5 Favours [DAPT] Favours [SAPT]

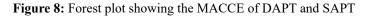


Figure 9: Forest plot showing the MI of DAPT and SAPT.

The statistics estimated is statistically significant with overall odd ratio of 0.78, [95% CI: 0.64, 0.94] when p- value (0.009) is less than the level of significant (0.05) which implies that the overall effect has a significant impact and a significant low event rates was found. The heterogeneity, $I^{\wedge} = 26\%$ across the studies is low with a insignificant value (p 0.18) which makes the MI of the DAPT and SAPT to be consistent across the different studies.

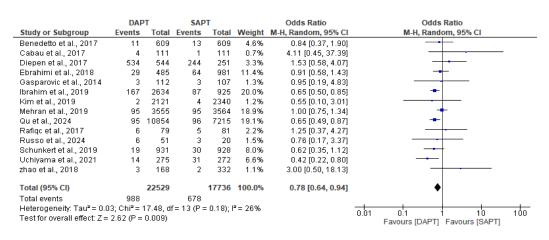


Figure 9: Forest plot showing the MI of DAPT and SAPT.

Figure 10: Forest plot showing the stroke of DAPT and SAPT.

The statistics estimated is statistically not significant with overall odd ratio of 0.95, [95% CI: 0.63, 1.45] when p- value (0.82) is less than the level of significant (0.05) which implies that the overall effect has a significant impact and a significant low event rates was found. The heterogeneity, $I^{\wedge} = 78\%$ across the studies is high with a significant value (p <000001) which makes the stroke of the DAPT and SAPT to be inconsistent across the different studies

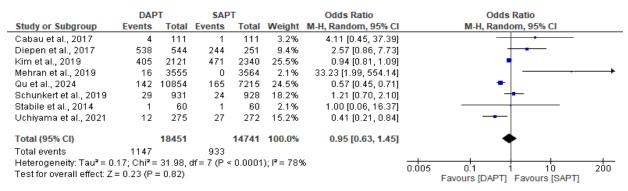


Figure 10: Forest plot showing the stroke of DAPT and SAPT.

Figure 11: Forest plot showing the bleeding of DAPT and SAPT.

The statistics estimated is statistically not significant with overall odd ratio of 1.21, [95% CI: 0.81, 1.81] when p- value (0.35) is greater than the level of significant (0.05) which implies that the overall effect has a significant impact and a significant low event rates was found. The heterogeneity, $I^{\wedge} = 65\%$ across the studies is moderate with a significant value (p 0.003) which makes the bleeding of the DAPT and SAPT to be inconsistent across the different studies bleeding.

	DAP	т	SAP	т		Odds Ratio	Odds Ratio
Study or Subgroup	Events	Total	Events	Total	Weight	M-H, Random, 95% Cl	M-H, Random, 95% Cl
Benedetto et al., 2017	14	609	25	2308	12.7%	2.15 [1.11, 4.16]	
Cabau et al., 2017	12	111	4	111	7.4%	3.24 [1.01, 10.39]	
Diepen et al., 2017	538	544	244	251	7.9%	2.57 [0.86, 7.73]	+ -
Gasparovic et al., 2014	28	112	20	107	12.9%	1.45 [0.76, 2.77]	+
Mehran et al., 2019	39	3555	72	3564	16.3%	0.54 [0.36, 0.80]	-
Quietial., 2024	53	10854	30	7215	15.6%	1.18 [0.75, 1.84]	+
Russo et al., 2024	1	52	2	20	2.4%	0.18 [0.02, 2.07]	
Schunkert et al., 2019	45	931	44	928	15.9%	1.02 [0.67, 1.56]	+
Stabile et al., 2014	4	60	3	60	5.0%	1.36 [0.29, 6.34]	
Uchiyama et al., 2021	2	275	3	272	4.0%	0.66 [0.11, 3.96]	
Total (95% CI)		17103		14836	100.0%	1.21 [0.81, 1.81]	•
Total events	736		447				-
Heterogeneity: Tau ² = 0.2	22: Chi ² = :	25.42. di	f=9(P=	0.003); P	² = 65%		
Test for overall effect: Z =			0				0.001 0.1 1 10 1000 Favours [DAPT] Favours [SAPT]

Figure 11: Forest plot showing the bleeding of DAPT and SAPT.

Funnel Plot

For all research, funnel plots are commonly used to identify publication bias; visual inspection of a funnel plot was done on all comparisons for the papers included in the meta-analysis of graft occlusion forest plot in Figure 12,13 below. The two funnel plot for the DAPT associated with autistic people revealed a symmetrical funnel with outliers, although this implies true heterogeneity rather than publication bias. Base on this, we can conclude none of the studies could be considered to be overly influential.

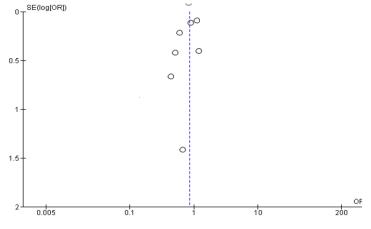


Figure 12: Graft occlusion funnel plot

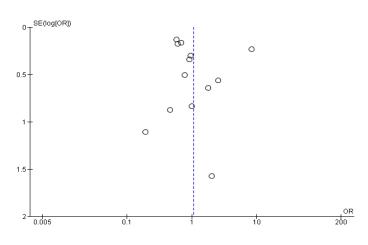


Figure 13: All cause of death funnel plot.

Discussion

The meta-analyses comparing Dual Antiplatelet Therapy (DAPT) with Single Antiplatelet Therapy (SAPT) in 18 studies, including a total of 218,125 patients, yielded valuable information regarding the effectiveness of these treatment plans in patients with cardiovascular conditions. Out of the 18 included studies, 7 studies was use to plot the forest plot of graft occlusion, 13 studies for all cause of death, 8 studies foe cardiovascular mortality, 4 studies for MACCE, 14 studies for myocardial infarction, 8 studies for stroke and 10 studies for major bleeding. The analysis yielded intriguing findings with substantial implications for clinical practice. A notable discovery from the current study was the considerably reduced occurrence of graft occlusion when using dual Antiplatelet Therapy (DAPT) in comparison to Single Antiplatelet Therapy (SAPT). The odds ratio of 0.71, with a 95% confidence range ranging from 0.54 to 0.93 and a p-value of 0.01, clearly demonstrates the significant benefit of DAPT in preventing Cardiovascular mortality. Cardiovascular patients receiving therapy is a critical event that could result in severe complications and jeopardize the procedure's effectiveness. Another significant discovery was the decreased occurrence of major adverse cardiovascular and cerebrovascular events (MACCE) and myocardial infarctions associated with dual Antiplatelet Therapy (DAPT). The odds ratios of 0.68 and 0.78 for major adverse cardiac and Cerebrovascular Events (MACCE) and myocardial infarction, with p-values of 0.0001 and 0.009, respectively, make it clear that Dual Antiplatelet Therapy (DAPT) is much better at lowering these bad outcomes than single antiplatelet therapy (SAPT), this was in line with the result of [15] found DAPT to be associated with cardiovascular mortality and a trend towards lower all-cause mortality. These results make it clear how important Dual Antiplatelet Therapy (DAPT) is for improving patient outcomes and lowering the risk of major cardiovascular events. Nevertheless, this research also revealed that there was no substantial disparity in the occurrence

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of graft occlusion, this result however contradict the result of Agarwal (2018) who found that DAPT appears to be associated with a reduction in graft occlusion compared with aspirin monotherapy in patients undergoing CABG [9], also there was no substantial disparity in the occurrence of all-cause death, stroke, and major bleeding between Dual Antiplatelet Therapy (DAPT) and single antiplatelet therapy (SAPT). Although DAPT has shown evident advantages in specific outcomes, such as MACCE and myocardial infarction, it has not established superiority in other domains. This suggests that the choice between DAPT and SAPT should be carefully considered based on the specific needs and risks of individual patients. The investigation included forest plots, which visually depicted the data and offered additional support for the study's findings. The forest plots for cardiovascular mortality, MACCE, and myocardial infarction demonstrated statistically significant outcomes in favor of DAPT, with reduced occurrence rates and a notable influence.

However, there were no statistically significant differences between dual antiplatelet therapy (DAPT) and Single Antiplatelet Therapy (SAPT) in the forest plots for graft occlusion, all-cause death, stroke, and major bleeding. This suggests that these outcomes were similar in both treatment plans. This result is inline with Schunkert, 2019 whose found MI, stroke, and cardiovascular death and all-cause death showed no significant differences between the two treatment groups [21]. The combined analyses, which compared Dual Antiplatelet Therapy (DAPT) with single antiplatelet therapy (SAPT) in patients with cardiovascular conditions, highlight the importance of tailoring treatment approaches to each patient's desired outcomes. Although DAPT had evident advantages in decreasing Major Adverse Cardiovascular And Cerebrovascular Events (MACCE) and myocardial infarction, it did not indicate superiority in all outcomes. Healthcare professionals should thoroughly evaluate the potential advantages and disadvantages of every available treatment choice in order to enhance the quality of patient care and the overall results for individuals with cardiovascular conditions. LimitationIt is very important to mention some of the limitations encountered during the database search and interpretation of the meta-analysis. First, the study included only articles published in English; as such, relevant studies might be found in studies published in other languages, which may improve the performance of the analysis. As a result, future researchers should include other languages [22-35]. Second, despite large cohorts, limiting occurrences per outcome resulted in insufficient analytical strength, particularly when analyzing only 18 randomized trials. Thus, these data are hypothesis-generating and demand bigger randomized trials. We may need further research and clinical trials to validate these findings and provide more comprehensive evidence to guide treatment decisions in this patient population.

Conclusion

In conclusion, there were no significant differences in the occurrence of graft occlusion, all-cause death, stroke, and major bleeding between the use of Dual Antiplatelet Therapy (DAPT) and Single Antiplatelet Therapy (SAPT). The p-values were greater than the 5% significance level. However, there were significant differences in the results of cardiovascular mortality, MACCE, and myocardial infarction, with p-values indicating statistical significance at a level below 5%. Overall, DAPT proved to be more effective than SAPT in reducing cardiovascular mortality, MACCE, and myocardial infarction. However, there was no notable distinction between the two regimens in terms of graft occlusion, all-cause death, stroke, and major bleeding. These findings provide clinicians with valuable insights into choosing the most suitable antiplatelet therapy based on the anticipated outcomes for individuals receiving treatment.

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