Need for repeat revascularisation in hybrid coronary revascularisation vs. percutaneous coronary intervention

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Abstract

Hybrid coronary revascularisation (HCR), being a treatment path combining both coronary artery bypass grafting and percutaneous coronary intervention (PCI) approaches, offers the advantages of both methods in patients with multi-vessel coronary artery disease. Since available literature provides few studies comparing the need for repeat revascularisation (RR) after HCR in comparison to PCI, our review aimed at summarising the latest data on this topic from the last 5 years (2018–2023). The search was conducted within the PubMed and Embase databases, followed by application of inclusion and exclusion criteria and providing a summary of data and characteristics of eligible studies. On the basis of 7 records included in the final analysis, RR and/or follow-up target vessel revascularisation (TVR) were significantly less frequently required in the case of HCR than in PCI in 3 out of 7 records, whereas the remaining four provided no significant differences in analysed rates between the 2 therapeutic pathways. When it comes to lowering the necessity for follow-up TVR and/or RR in a fraction of instances, HCR demonstrates a significant differences were observed between the 2 methods in the remaining 4 records.

Key words: percutaneous coronary intervention, repeat revascularisation, hybrid coronary revascularisation.

Introduction

Hybrid coronary revascularisation (HCR) is a treatment method with the inclusion of coronary artery bypass grafting (CABG) and percutaneous coronary intervention (PCI) targeting patients suffering from multivessel coronary artery disease (CAD) [1].

HCR typically integrates the minimally invasive CABG procedure, where the left internal mammary artery (LIMA) is adhered to the left anterior descending coronary artery (LAD), with PCI aimed at non-LAD vessels [2]. The aim of this approach is to combine the advantages of both surgical and percutaneous revascularisation while minimising some of their respective drawbacks [1]. There are several clinical situations, where HCR is notably recommended: history of prior CABG, inadequate or poor-qual-

ity venous conduits, non-LAD lesions amendable to PCI, or LAD lesion not amendable to PCI [3]. Essentially, this novel revascularisation method manages the survival benefits associated with the LIMA-to-LAD graft while offering a comprehensive and minimally invasive cure for coronary artery revascularisation, which includes PCI for arteries other than the LAD [2].

In the literature and clinical practice, we distinguish 3 different sequences of performing the procedure: PCI before surgery, PCI after surgery (both known as 2-stage HCR), and in the case of performing both procedures in a single approach – single-stage or simultaneous HCR [2]. The possible paths for performing the surgical revascularisation during HCR include the following: conventional on-pump and off-pump coronary artery bypass grafting, minimally invasive direct coronary artery bypass (MIDCAB),

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Krzysztof Sanetra, Department of Cardiac Surgery, American Heart of Poland, Bielsko-Biala; Division of Cardiovascular Surgery, Collegium Medicum, Andrzej Frycz Modrzewski Krakow University, Krakow, Poland, e-mail: krzyssan@poczta.onet.pl **Received:** 3.05.2024, **accepted:** 10.06.2024, **online publication:** 17.06.2024. endoscopic, atraumatic coronary artery bypass grafting (EACAB), robotic-assisted CABG (RACAB), and totally endoscopic coronary artery bypass (TECAB) [1]. All the mentioned techniques aim to perform LIMA-LAD anastomosis with superior long-term patency. While the available literature focuses mainly on summarising a wide range of multicentre results, patients' characteristics, or data comparing the results depending on the selected path of revascularisation, our review aims to provide the most detailed discussion on the need for repeat revascularisation (RR) in patients after undergoing HCR, in comparison to PCI, based on recent (2018–2023) publications.

Material and methods

The studies summarised in this review were exclusively sought in English language within the PubMed and Embase databases, as well as through manual extraction of referenced literature within the previously identified manuscripts. Throughout the search, we used the following terms: (((hybrid coronary revascularization[Title/ Abstract]) OR (HCR[Title/Abstract]))) AND ((PCI[Title/ Abstract]) OR (PERCUTANEOUS CORONARY INTERVEN-TION[Title/Abstract]) OR (repeat revascularization[Title/ Abstract]) OR (REPEAT PCI[Title/Abstract]) OR (REPEAT CABG[Title/Abstract]) OR (MACCE[Title/Abstract]) OR (FOLLOW-UP[Title/Abstract]) OR (LONG-TERM[Title/ Abstract]) OR (SHORT-TERM[Title/Abstract]) OR (MID-TERM[Title/Abstract]) OR (OUTCOME[Title/Abstract]) OR (OBSERVATION[Title/Abstract])), and a specific time frame from January 2018 to August 2023 was applied. We decided to summarise the results of eligible studies published in the last 5 years to ensure findings that reflect the most current evidence on the analysed topic. Furthermore, both technology (new generation drug-eluting stents, improved surgical vision devices) and approach have changed in the past 10 years, and recent results cannot be compared to outdated databases from when mainly bare-metal stents were used.

After the duplicates were removed, the inclusion of the articles was performed upon screening and applying the following eligibility criteria: the study participants were patients undergoing hybrid coronary revascularisation; the control interventions consisted of PCI alone; primary or secondary endpoints in the selected studies had to include data reporting RR, regardless of the follow-up time.

Exclusion criteria included insufficient detail concerning RR and/or target vessel revascularisation (TVR) and lack of accessibility. Systematic reviews and metanalyses from previous years were also excluded from the study.

A flow diagram illustrating the records qualification process for the review is presented in Figure 1.

Data collected from the studies eligible for the review included general information about the publications (title, first author, country, DOI) as well as further details, i.e. the number of patients, inclusion criteria, interventions' techniques, RR and/or TVR rates depending on the primary procedure, and follow-up data. Two authors independently conducted the literature search, decided on inclusion of the eligible articles, and addressed any differences and quality considerations by collaborative discussion and following inclusion or exclusion.

Results

Study selection

The initial search strategy, based on carefully selected terminology, as outlined in the Methods section, resulted in a total of 401 records, available through advanced search in the PubMed and Embase databases. After removing duplicates and following a selection process guided by specific eligibility criteria, 7 records (6 original studies [4–10], 1 follow-up research letter [6]) conducted between the years 2018 and 2023 were selected for the final analysis.

Characteristics

Our study involves a total of 28,672 patients, including 704 (2.46%) patients in the HCR group and 27,968 (97.54%) subjects in the PCI group. Because our primary objective was to consolidate the data regarding RR and/ or TVR, we have provided a summary of RR and/or TVR, depending on the outcome measure reported in each of the records. Characteristics of eligible studies presented in Tables I and II, besides RR and/or TVR rates, include



Figure 1. Study flowchart

Table I. Summary of the results of studies assessing repeat revascularisation in hybrid coronary interventions vs. percutaneous coronary intervention

Ref;	First author, publication year, country	Patient enrolment	Study type	RR and/or TVR for HCR	RR and/or TVR for PCI	Other
[4]	Hannan E, 2021, USA	2010 – 2016	Retro- spective observa- tion	Time [years] (No RR, No. at risk [n]; N = 335): 1 (330) 2 (272) 3 (215) 4 (158) 5 (117) 6 (80) Freedom from RR in tl 4-year fo HCR 91.13% v p-value aHR = 95% CI: 0	Time [years] (No RR, No. at risk [n]; N = 27557): 1 (23388) 2 (18442) 3 (14314) 4 (10882) 5 (7773) 6 (5243) he LAD artery (median bollow-up) ys. PCI 83.59% e 0.001, = 0.51, 0.34-0.77	RR – any unstaged revascularisation (PCI or CABG surgery) in the LAD artery; 335 HCR: 320 off-pump surgery, 5 off- pump surgery followed by on-pump surgery, 10 on-pump surgery; Interaction between RR and 6 highest volume HCR hospitals (aHR = 0.42, 95% CI: 0.26–0.69, <i>p</i> -value = 0.01). Examination of pre-selected subgroups of patients indicates that no patient subgroups had significant interactions between revascularisation strategy and RR in the LAD artery.
[5, 6]	Ganyukov V, 2020, Poland and Russia [5] Ganyukov V, 2021, Russia [6]	2012	Ran- domised controlled trial	30 days: 1.9% (1) (<i>N</i> = 49); 12 months: clinically driven TVR 1.9% (1); Angiography – driven TVR 11.5% (6); total TVR 13.5% (7) [6] F/u time [months]: 52.5 (min. 36) Clinically driven TVR: 16.6 (8)	30 days: 0% (0) (N = 51); 12 months: clinically driven TVR 5.7% (3); Angiography – driven TVR 11.3% (6); total TVR 17.0% (9) [6] F/u time [months]: 52.5 (min. 36) Clinically driven TVR: 20.0 (10)	 HCR: incomplete TLR (per patient) 7.7% (4) Incomplete TLR (per total number target lesions in study group) 2.7% (4/149); PCI: incomplete TLR (per patient) 5.7% (3) Incomplete TLR (per total number target lesions in study group) 2.1% (3/146).
				p-val p-value	ue NS e NS [6]	
[7]	Basman C, 2020, USA	2009–2016	Retro- spective observa- tion	Time [years] (Freedom from RR, No. at risk [n]): 2 (69) 4 (30) 6 (17) 8 (0) (8-year f/u) RR: 16 TVR: 6 (3 problems with the LIMA to LAD graft, 3 in non-LAD vessels), 10 (62.5%) de novo lesions	Time [years] (Freedom from RR, No. at risk [<i>n</i>]): 2 (63) 4 (37) 6 (18) 8 (0) (8-year f/u) RR: 18 TVR: 10	TVR – a repeat intervention for a prior stented lesion, either within the stent itself or within 5 mm of the stent, and/ or a repeat procedure for a lesion that was previously surgically bypassed. In the TVD patient population with in- termediate SYNTAX scores, although midterm survival is comparable across treatment arms, morbidity may be higher after PCI, particularly with re- spect to the increased incidence of RR and new MI.
				RR <i>p-</i> va TVR <i>p-</i> v	alue NS ⁄alue NS	
[8]	Modrau IS, 2020, Denmark	2010–2012	Retro- spective observa- tion	(3-year f/u) 21.4% (index hospitalisation (8/16), prescheduled 1-year angiography (8/16); 9/16 (56%) during the first year were driven by angio- graphic findings w/o associated symptoms of ischaemia) Time [years] (Freedom from RR, No. at risk [<i>n</i>]): 1 (85) 2 (76) 3 (75)	(3-year f/u) 12.6% Time [years] (Freedom from RR, No. at risk [<i>n</i>]): 1 (93) 2 (89) 3 (87)	Multivessel PCI was performed "one- stop" in 75 (73%) patients and staged in 28 (27%) patients. HMR was converted to CABG in 3 pa- tients and censored as RR and analysed as intention to treat (failed PCI for total chronic occlusion in 2 patients, LIMA graft thrombosis and procedure-related myocardial infarction in 1 patient).
<i>p</i> -value NS						

Patient enrolment	Study type	RR and/or TVR for HCR	RR and/or TVR for PCI	Other
2009–2016	Retro- spective observa- tion	TVR Perioperative: 0/52, Mid-term 59 months (interquartile range, 42 to 79 months): 2/44 (4.5%) <i>p</i> -valu	TVR Perioperative: 0/44, Mid-term 59 months (interquartile range: 42 to 79 months): 10/45 (22.2%) e 0.015	-
2013–2016	Retro- spective observa- tion	(100% complete angio- graphic follow-up at 12 months); TVR (4): 2 in-stent restenoses of left main-Cx stent, 1 poststent stenosis, 1 in-stent restenosis on RCA lesion; no procedures on LAD for LIMA-LAD graft failure or stenotic anasto- mosis Time [months] – Surv (HCR/F 3 – 6 6 – 6 9 – 5 12 – 4 15 – 4 Survival freed (HCR: 93.3 (4.6)%	7 patients: plain old balloon angioplas- ty (POBA; kissing balloon) on left main for bifurcation initial restenosis; 3 in-stent restenoses treated by new PCI, 4 poststent stenosis, 2 incomplete distal stent expansion, 8 in-stent restenosis on a RCA lesion ival freedom from TVR PCI) [n]: 7/102, 51/97, 4/90, 43/81, 11/64, 10/57; lom from TVR: 6 PCI: 75.5 (5.6) % e 0.002	HCR vs. PCI studies demonstrated that the complexity of the coronary lesion directly affects the outcomes of PCI, es- pecially the TVR (mainly concentrated in the LAD), whereas PCI with DES for non-LAD offered low and similar TVR rates in both HCR and PCI groups. PCI stenting on left main was an inde- pendent predictor of MACCEs (hazard ratio 4.1, 95% CI 2.4–11.3; <i>p</i> -value 0.001) and TVR (hazard ratio 3.9, 95% CI 1.36– 9.64; <i>p</i> -value 0.002). Female sex was an independent predictor of TVR (haz- ard ratio 2.1, 95% CI: 1.12–4.65; <i>p</i> -value 0.049).
	Patient enrolment	Patient enrolmentStudy type2009–2016Retro- spective observa- tion2013–2016Retro- spective observa- tion	Patient enrolmentStudy typeRR and/or TVR for HCR2009–2016Retro- spective observa- tionTVR Perioperative: 0/52, Mid-term 59 months (interquartile range, 42 to 79 months): 2/44 (4.5%)2013–2016Retro- spective observa- tion(100% complete angio- graphic follow-up at 12 months); TVR (4): 2 in-stent restenoses of left main-Cx stent, 1 poststent stenosis, 1 in-stent restenosis on RCA lesion; no procedures on LAD for LIMA-LAD graft failure or stenotic anasto- mosisTime [months] – Survi (HCR/I 3 – 6 6 – 6 9 – 5 12 – 15 – 4 18 – 4	Patient enrolmentStudy typeRR and/or TVR for HCRRR and/or TVR for PCI Retro- spective observa- tionRR and/or TVR for HCRRR and/or TVR for PCI2009–2016Retro- spective observa- tionTVRTVR Perioperative: 0/52, Mid-term 59 months (interquartile range, 42 to 79 months): 2/44 (4.5%)TVR Perioperative: 0/44, Mid-term 59 months (interquartile range: 42 to 79 months): 10/45 (22.2%)2013–2016Retro- spective observa- tion(100% complete angio- graphic follow-up at 12 months); TVR (4): 2 in-stent restenoses of left main-Cx stent, 1 poststent stenosis, 1 in-stent restenosis on RCA lesion; no procedures on LAD for UIMA-LAD graft failure or stenotic anasto- mosis7 patients: plain old balloon angioplas- ty (POBA; kissing balloon) on left main for bifurcation initial restenosis; 3 in-stent restenosis, 2 incomplete distal stent stenosis, 0 a RCA lesionTime [months] – Survival freedom from TVR (HCR/PCI) [n]: 3 – 67/102, 6 – 61/97, 9 – 54/90, 12 – 43/81, 15 - 41/64, 18 – 40/57; Survival freedom from TVR: (HCR: 93.3 (4.6)% PCI: 75.5 (5.6) % P-value 0.002

aHR – adjusted hazard ratio, CABG – coronary artery bypass grafting, CI – confidence interval, DES – drug-eluting stents, HCR – hybrid coronary revascularisation, HR – hazard ratio, LAD – left anterior descending artery, LIMA – left internal mammal artery, MACCE – major adverse cerebral and cardiac events, MI – myocardial infarction, PCI – percutaneous coronary intervention, POBA – plain old balloon angioplasty, RCA – right coronary artery, RR – repeat revascularization, TLR – target lesion revascularisation, TVD – triple vessel disease, TVR – target vessel revascularisation.

study type, time of patients' enrolment, outcome measures other than RR and/or TVR (MACCE, death, myocardial infarction, stroke, all-cause mortality) with median follow-up time and HCR approach and sequence (simultaneous or 2-stage HCR). The geographic distribution of studies encompassed North America, Europe, and Asia.

Repeat revascularisation and/or follow-up target vessel revascularisation

In 3 out of 7 records, the RR and/or follow-up TVR rates were more favourable for HCR compared to PCI. It was observed that RR was significantly less frequently required in the case of HCR than in PCI. Hannan *et al.* observed after a follow-up period of 4 years freedom from RR in the LAD artery in 91.13% in the HCR group, whereas in the PCI group the rate was 83.59% with a *p*-value of 0.001 (aHR = 0.51, 95% CI: 0.34-0.77) [4]. Moreover, they reported an

interaction between RR and the 6 highest-volume HCR hospitals (aHR = 0.42, 95% CI: 0.26-0.69, p = 0.01). An analysis of pre-selected patient subgroups revealed that there were no significant interactions between the patient subgroups and the revascularisation strategy or risk reduction in the LAD artery [4].

In a study from 2019, the perioperative TVR rate was 0 for both groups, whereas after 59 months in a midterm follow-up TVR was performed in 2/44 (4.5%) for the HCR group and 10/45 (22.2%) in the PCI group with *p*-value at 0.015 [9].

In 2018, Repossini *et al.* reported a survival freedom from TVR at 93.3% (4.6%) for HCR and 75.5% (5.6%) for PCI (p = 0.002). According to their analysis, HCR vs. PCI trials showed that the intricacy of the coronary lesion strongly influences PCI results, particularly the TVR (which is mostly concentrated in the LAD). In contrast, PCI with DES for non-LAD provided low and comparable TVR rates in both HCR and PCI groups. Also, they observed female sex as an independent predictor of TVR (HR = 2.1, 95% CI: 1.12-4.65; p = 0.049) [10].

In the remaining 4 studies, no significant differences in RR and/or TVR rates between the 2 treatment strategies were observed [5–8]. A summary of the results is presented in Table I. Additional data regarding characteristics of the patients in the included studies are presented in Table II.

Quality considerations

In a study by Hannan *et al.*, the issue of selection bias caused by lack of randomisation was minimised by employment of Cox proportional hazards models so as to control for differences in patient risk factors among patients undergoing the analysed procedures. Given that the study included only patients who survived long enough to have the second treatment, it was noted that for 2-stage procedures, survival bias could be present. Another constraint is that HCR encompasses a wide

Table II. Additional da	ta regarding charad	cteristics of patients	in included studies
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Ref	First author, publication year, country	Eligibility criteria	Number of patients HCR/PCI	HCR ap- proach	Si- mul- tane- ous HCR	HCR: PCI after surgery	HCR: PCI before surgery	Other outcomes HCR/ PCI
[4]	Hannan E, 2021, USA	 MV-D (≥ 70% stenosis in ≥ 2 major epicardial CA), in- cluding diseased LAD artery (≥ 70% stenosis) Minimally invasive CABG surgery (no sternotomy) or PCI in the LAD artery Elective PCI in one or more other diseased arter- ies within 60 days before or after the LAD procedure without any other concomi- tant major cardiac surgery 	335 (1.20%) /27,557 (98.80%)	Isolated CABG surgery performed on the LAD artery with minimally in- vasive surgery (off-pump/on- pump) + PCI procedures performed within 60 days before or after the CABG surgery in non-LAD vessels	44%	18%	38%	Median f/u time [years]: 3.81 (HCR), 4.20 (PCI) [Prevalence HCR/PCI; Four-year mortality: hazard ratio (95% CI) of HCR/PCI, <i>p</i> -value] MI within 20 days 28.96%/37.53%; 0.76 (0.44–1.32), 0.44 Cerebrovascular disease 10.15%/8.63%; 1.14 (0.62–2.10), 0.37 BMI < 25 kg/m ² 27.76%/21.67%; 0.63 (0.36–1.12), 0.003
[5, 6]	Ganyukov V, 2020, Poland and Russia [5] Ganyukov V, 2021, Russia [6]	 Angiography-confirmed MV-CAD involving LAD and a significant (≥ 70% diameter stenosis, DS, on quantitative coronary angiography, QCA) lesion in at least one major non-LAD epicardial vessel of ≥ 2.5 mm in diameter, amenable to PCI and CABG and HCR 2) Lesions of 50–70% DS were subjected to func- tional evaluation and were considered the study target lesions (i.e, were labelled for revascularization) if lesion-related myocardial ischemia was present on functional testing (fractional flow reserve, FFR, or SPECT stress imaging 	52/ 53 [6] 3-year F/u (randomized) 48 (52)/50 (53)	MIDCAB LIMA-LAD + PCI for non- LAD vessel/s		HCR patients, except 5 (9.8%) who required conver- sion to CABG had per-pro- tocol PCI within 3 days (in most cases at 24–48 h) after per- forming MIDCAB LIMA-LAD anasto- mosis that was always the first stage of HCR	_	MACCE (death/stroke/ MI/clinically driven re- peat revascularisation) F/u time [days]: 30 Death 1.9% (1)/0% (0) Stroke 1.9% (1)/0% (0) MI 5.8% (3)/3.8% (2) F/u time [months]: 12 Death 5.8% (3)/3.8% (2) Stroke 3.8% (2)/0% (0) MI 5.8% (3)/7.5% (4) [6] F/u time [months]: 52.5 (min. 36) All-cause mortality 6.3 (3)/6.0 (3) MI 6.3 (3)/12.0 (6) Stroke 4.2 (2)/8.0 (4)

Table II. Cont.								
Ref	First author, publication year, country	Eligibility criteria	Number of patients HCR/PCI	HCR ap- proach	Si- mul- tane- ous HCR	HCR: PCI after surgery	HCR: PCI before surgery	Other outcomes HCR/ PCI
[7]	Basman C, 2020, USA	1) Stable TVD (w/o con- comitant non-coronary procedure, previous coro- nary and/or valve surgery, emergency/salvage surgery, hemodynamic instability)	100 (after propensity match)/100 (after propen- sity match)	Off-pump robotic-assist- ed LIMA to LAD bypass (MIDCAB component of HCR) + PCI standard techniques (~50% radial approach); either second- or third-gen- eration DES	0	72 (MIDCAB first approach, followed by interval PCI, typically within 4 to 6 weeks of surgery)	28 (coronary syndrome in which the culprit lesion was deemed to be within one of the non-LAD vessels, or angio- graphic severity and clinical import of at least one of the non-LAD stenosis greater than that of the disease within the LAD itself. For these patients, subsequent LIMA to LAD graft- ing was undertaken on unin- terrupted DAPT)	F/u time [days]: 30 30-d mortality 0/0 Stroke 0/0 Periprocedural MI 0/0 New-onset renal failure 0/0 Length of stay, days, mean \pm SD 5.7 \pm 7.5/2.0 \pm 2.2 ($p < 0.0001$) Residual SYNTAX score, mean \pm SD 4.5 \pm 4.4/7.1 \pm 6.5 ($p < 0.001$) Mean (SD) f/u [years]: 7.14 (0.12) Mortality 5/9 ($p = 0.41$) Myocardial infarction 4/5 ($p = 1.0$) MACE (death, repeat revascularisation, and myocardial infarction) 21/25 ($p = 0.61$)
[8]	Modrau IS, 2020, Den- mark	1) Age: 18 years 2) MVD involving the LAD	103/103	Offpump anastomosis of the LIMA to the LAD through a left inferior J-hemis- ternotomy (JOPCAB)	-	11 (11%)	92 (89%)	F/u time [years]: 3 MACCE (all-cause death, myocardial infarction, stroke, and repeat re- vascularisation at 3-year follow-up) Death 6.8%/5.8% Myocardial infarction 3.9%/3.9% Stroke 3.9%/2.9%
[9]	Qiu J, 2019, China	 The patient underwent HCR, isolated OPCAB or isolated PCI The patient had two-ves- sel CAD including proximal LAD stenosis LIMA-to-LAD anastomosis was performed in patients underwent HCR or OPCAB The stents used in HCR or PCI were drug eluting stents (DES) Exclusion criteria: the operation was emergent; the patient had undergone coronary revascularisation before 	47 (after pro- pensity score matching)/ 47 (after pro- pensity score matching)	LIMA-to-LAD anastomosis; the stents used in HCR or PCI were drug-eluting stents (DES)	-	-	_	MACCE (death, MI, stroke, TVR) F/u time [days]: 30 Death 0/0 MI 0/1 (p = 0.365) Stroke 0/0 F/u time [months]: 59 [42–79] Death 1/2 (p = 0.811) MI 1/3 (p = 0.411) Stroke 2/3 (p = 0.874)

Table II. Cont.

Ref	First author, publication year, country	Eligibility criteria	Number of patients HCR/PCI	HCR ap- proach	Si- mul- tane- ous HCR	HCR: PCI after surgery	HCR: PCI before surgery	Other outcomes HCR/ PCI
[10]	Repossini A, 2018, Italy	1) Critical left main stenosis or equivalent left main lesion, with or without mul- tivessel coronary lesions 2) Primary/rescue PCI for acute coronary syndrome on non-LAD lesions with residual lesions on left main (excluded: distal heavy cal- cified lesions and isolated ostial or proximal-mid-body left main disease, concom- itant surgical procedures in addition to myocardial revascularisation)	67 (pre- operative matched)/ 108 (pre- operative matched)	LIMA-LAD and PCI on other target vessels (MIDCAB was performed as the first step of the hybrid revascularisa- tion strategy, followed by PCI stenting of circumflex artery and non-LAD lesions)	0	62 (un- protected LMCD, a surgical revascu- lariza- tion via MIDCAB was per- formed as the first step of the hybrid revascu- larization strategy, followed by PCI stenting of cir- cumflex artery and non-LAD lesions) (a time- frame of about 1–4 weeks)	5 (left main equiv- alent lesions with ostial stenosis of both LAD and circum- flex artery (Cx), PCI stenting from Cx to left main was per- formed before MIDCAB)	MACCEs (cardiac death, stroke, AMI, repeated TVR) F/u time [days]: 30 In-hospital mortality 0/3 (2.7) ($p = 0.603$) Stroke 0/1 (0.9) ($p = 0.839$) Myocardial infarction 0/1 (0.9) ($p = 0.839$) Postoperative atrial fibrillation 8 (11.9)/1 (0.9) ($p = 0.008$) Pericardial effusion 3 (4.4)/ 5 (4.6) ($p = 0.984$) Mean (SD) f/u [months]: HCR 15.4 (2.6)/PCI 15.2 (2.8) Mortality at 18 months' O/0 Major cerebral adverse events 0/2 AMIs 0/7 Survival freedom from MACCEs at 12 months' 97.2 ±2.5%/86.3 ±3.2 Survival freedom from MACCEs at 18 months' 93.3 ±4.6%/ 72.3 ±6.3 ($p = 0.001$)

AMI – acute myocardial infarction, BMI – body mass index, CABG – coronary artery bypass grafting, CA – coronary arteries, DAPT – dual antiplatelet therapy, DES – drug eluting stents, FFR – fractional flow reserve, HCR – hybrid coronary revascularization, LAD – left anterior descending artery, LIMA – left internal mammal artery, LMCD – left main coronary artery disease, MACCE – major adverse cerebral and cardiac events, MACE – major adverse cardiac events, MI – myocardial infarction, MIDCAB – minimally invasive direct coronary angiography, MVD – multivessel disease, PCI – percutaneous coronary intervention, TVD – triple vessel disease.

range of procedures, and we could not evaluate the effects of the pharmacological therapies administered between and after the procedures or assess how their usage impacted the outcomes [4].

Due to variations in SYNTAX scores, Basman *et al.* stated in their study that their data is not appropriate for drawing conclusions regarding the superiority of one revascularisation technique over another [7]. The Repossini *et al.* study did not fully document the need for clinical versus angiographic reasons for RR, which means that it is not possible to completely rule out the possibility of an excessively high rate of prudential RR in the event of initial restenosis [10].

In the HREVS RCT study the issue of patient's choice towards a less invasive procedure was considered with result of 1 in 4 refusal rates to random treatment allocation due to preference of PCI. However, the overall recruitment rate was over 75% [5]. It was also observed that while considering the applicability of the results, the moderate angiographic complexity of multi-vessel disease (MVD) must be taken into account. This complexity reflects the necessity for the technical feasibility of HCR and multi-vessel PCI (MVPCI). Due to this criterion, cases involving left main coronary artery stenosis not amenable to HCR, severely calcified lesions, complex bifurcations, or chronic total occlusion were excluded (all of which may favour surgical interventions) [5].

The principal constraint of the Modrau study was its sufficient statistical power to draw definitive conclusions regarding MACCE endpoints. To derive meaningful insights, the results should be considered alongside other studies that contribute to the broader evidence on the topic. It is worth noticing that the retrospective SYNTAX calculation was conducted retrospectively and, therefore, did not serve as a matching criterion for control patients undergoing PCI or CABG. The notably high repeat revascularisation rate observed in the hybrid revascularisation (HMR) group was primarily a result of angiography following the protocol rather than symptom-driven, introducing a bias against HMR, because this protocol was not applied to the CABG and PCI groups [8].

Discussion

In cases where both the LAD artery and at least one other significant coronary artery are involved, HCR is an infrequent treatment path as an alternative to PCI for CAD patients [4]. In 2021 Hannan *et al.* reported that, because no significant difference in mortality in a median follow-up of 4 years was noted, HCR exhibits a lower rate of RR as opposed to PCI (91.13% vs. 83.59%, p = 0.001, aHR = 0.51 [95% CI: 0.34-0.77]), which creates a field for analysing whether and why it is worth to consider HCR as an alternative treatment pathway. Especially, since there is a limited number of publications regarding this criterium [4].

Several research articles have examined diverse outcomes of HCR compared to traditional PCI. In patients with MVD, both HCR and PCI showed similar 6-year risk-adjusted survival, whereas HCR patients were less likely to require a repeat LAD revascularisation at that time [4]. Eight-year survival outcomes in patients with TVD treated with HCR compared with multivessel PCI showed similar mortality rates, with HCR having a lower residual SYNTAX score [7]. In a different study, after a 3-year follow-up period, HCR has shown similar results to PCI in terms of all-cause mortality, myocardial infarction, stroke, TVR, and major adverse cardiac and cerebrovascular events (MACCE). To fully ascertain the potential of HCR as a coronary revascularisation technique in patients with MV-CAD, the study highlights the necessity for more extensive studies and longer follow-up [6].

In a study by Repossini *et al.*, following an 18-month period, HCR demonstrated a markedly reduced incidence of MACCEs, mostly as a result of greater independence from TVR. The improved results in terms of MACCEs may be explained by the advantages of the left internal mammary artery to left anterior descending artery (LIMA-LAD) bypass versus PCI in terms of patency rates, according to the research. It also draws attention to the sequential staged method used in HCR, highlighting its possible benefits in lowering the risk of bleeding and thrombotic events in comparison to single-step revascularisation techniques [10].

HCR is gaining popularity as an alternative to CABG and PCI in the treatment of the left main artery, providing a solution that integrates the advantages of both techniques while minimising surgical trauma and postoperative complications [4, 10]. HCR is a safe procedure, providing promising midterm results, also in patients with high risk, and long-term in patients with multivessel coronary artery disease, especially in patients with proximal LAD stenosis [9, 11, 12]. The benefits of HCR are numerous; however, the decision to choose this treatment path over PCI must be made on an individual basis, taking into account factors such as lesion complexity and risk for the patient. It must be remembered that the hybrid revascularisation protocol is not unified, and many divergences can be observed in different institutions. Those may include not only alterations in procedure sequence (PCI-first; surgery-first; one-stage), but also timing of divided procedures (days/months) and qualification criteria. Obviously, institutional experience in such procedures also contributes to the results. As such, significant variations in study results can be observed.

Clinical guidelines emphasise the limited evidence from randomised control trials to support hybrid revascularisation [13]. As such, it is essential to conduct such multicentre studies with a unified protocol, including antiplatelet treatment, and patients' eligibility criteria. This would produce the data that would finally result in modification of European Associations guidelines. It is also essential to properly address the patient groups, because hybrid coronary revascularisation may be particularly beneficial in both young, active patients with relatively low risk of complications and in patients for whom standard surgical treatment is of great risk of complications. As such, a clear and effective protocol for such study is required.

Study limitations

The analysis has following limitations: Firstly, hybrid revascularisation protocols are different in various institutions, which leads to differences in the results. Secondly, the devices used for both percutaneous and surgical stage of hybrid revascularisation are different in different institutions, as is clinical experience. Qualification criteria vary in those facilities, suggesting that different groups of patients are included in the selected studies. Finally, mainly retrospective studies were included, which increases the risk of bias.

Conclusions

HCR as a treatment method exhibits a significant advantage over PCI in decreasing the necessity for RR and/ or follow-up TVR in a subset of cases. Nevertheless, it is important to remember that in the remaining 4 out of 7 studies, no statistically significant differences were reported between the 2 therapeutic methods, emphasising the complexity and variability of outcomes associated with these therapeutic techniques.

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Ethical approval

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Conflict of interest

The authors declare no conflict of interest.

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