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Contents

Thrice Monthly Volume 12 Number 13 May 6, 2024

EDITORIAL

2138	8 Parenteral iron therapy in children with iron deficiency anemia	
	Roganovic J	
2143	Treatment-induced neuroendocrine prostate cancer and <i>de novo</i> neuroendocrine prostate cancer: Identi- fication, prognosis and survival, genetic and epigenetic factors	
	Wishahi M	
2147	Perioperative cardiac risks in myasthenia gravis	
	Nag DS, Chatterjee A, Mahanty PR, Sam M, Bharadwaj MK	

- 2151 Management of geriatric acetabular fractures: Contemporary treatment strategies Tosounidis T, Chalidis B
- 2157 Pioneering role of machine learning in unveiling intensive care unit-acquired weakness Dragonieri S

ORIGINAL ARTICLE

Case Control Study

Detection and analysis of serum bile acid profile in patients with colonic polyps 2160

Ji X, Chen H

2173 Clinical analysis of colistin sulfate in the treatment of pneumonia caused by carbapenem-resistant Gramnegative bacteria

Xu HC, Cui Y, Wang XY, Wu HB, Li W, Wang D, Lin N, Lin L, Zhang YH

Retrospective Cohort Study

2182 Establishment and evaluation of a prognostic model for patients with unresectable gastric cancer liver metastases

Chang ZY, Gao WX, Zhang Y, Zhao W, Wu D, Chen L

Retrospective Study

2194 Therapeutic effect of Wendan Decoction combined with mosapride on gastroesophageal reflux disease after esophageal cancer surgery

Zhang YJ, Wu SP

Clinical Trials Study

2201 Yiwei Xiaoyu granules for treatment of chronic atrophic gastritis with deficiency syndrome of the spleen and stomach

Chen WQ, Fan QF, He YJ, Li F, Wu X, Li YP, Yang XJ



Contents

Thrice Monthly Volume 12 Number 13 May 6, 2024

Observational Study

2210 Relationship between clinical belonging, professional identity, and nursing information ability among nursing interns: Model construction

Zhang G, Huang SJ, Li SF

META-ANALYSIS

2218 Efficacy and safety of Yangxinshi tablet for chronic heart failure: A systematic review and meta-analysis Lu SH, Yu YF, Dai SS, Hu YQ, Liu JH

CASE REPORT

- 2231 Intra-thyroid esophageal duplication cyst: A case report Lin HG, Liu M, Huang XY, Liu DS
- 2237 Magnetic resonance imaging findings of radiation-induced breast angiosarcoma: A case report Wu WP, Lee CW
- Rim ¹⁸F-fluorodeoxyglucose uptake of hepatic cavernous hemangioma on positron emission 2243 tomography/computed tomography: A case report

Hu YA, Guo YX, Huang QF

- 2248 Recovering from prolonged cardiac arrest induced by electric shock: A case report Zhang J, Qiao YR, Yang YD, Pan GZ, Lv CQ
- 2254 Young patient with a giant gastric bronchogenic cyst: A case report and review of literature Lu XR, Jiao XG, Sun QH, Li BW, Zhu QS, Zhu GX, Qu JJ
- 2263 Airway management of a patient with linear immunoglobulin A bullous dermatosis: A case report Nin OC, Hutnik R, Chheda NN, Hutchinson D
- 2269 Deferred revascularization in diabetic patient according to combined invasive functional and intravascular imaging data: A case report Al Nooryani A, Aboushokka W, Beleslin B, Nedeljkovic-Beleslin B
- 2275 Thymic carcinoid with multiple bone metastases: A case report Chen CQ, Huang MY, Pan M, Chen QQ, Wei FF, Huang H
- 2281 Atypical presentation of a posterior fossa tumour: A case report Narotam A, Archary M, Naidoo P, Naidoo Y, Naidu V
- 2286 Refractory autoimmune hemolytic anemia in a patient with systemic lupus erythematosus and ulcerative colitis: A case report Chen DX, Wu Y, Zhang SF, Yang XJ



Contents

Thrice Monthly Volume 12 Number 13 May 6, 2024

ABOUT COVER

Peer Reviewer of World Journal of Clinical Cases, Konosuke Nakaji, FACP, MD, Doctor, Endoscopy Center, Aishinkai Nakae Hospital, Wakayama-shi 640-8461, Japan. parupurikopui@yahoo.co.jp

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The primary aim of World Journal of Clinical Cases (WJCC, World J Clin Cases) is to provide scholars and readers from various fields of clinical medicine with a platform to publish high-quality clinical research articles and communicate their research findings online.

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CASE REPORT

Deferred revascularization in diabetic patient according to combined invasive functional and intravascular imaging data: A case report

Arif Al Nooryani, Wael Aboushokka, Branko Beleslin, Biljana Nedeljkovic-Beleslin

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Arif Al Nooryani, Wael Aboushokka, Department of Cardiology, Al Qassimi Hospital, Sharjah 1234, United Arab Emirates

Branko Beleslin, Department of Cardiology, Medical Faculty, University of Belgrade, Belgrade 11000, Serbia

Biljana Nedeljkovic-Beleslin, Institute of Endocrinology, Diabetes and Metabolic Disorders, University Clinical Center of Serbia, Belgrade 11000, Serbia

Biljana Nedeljkovic-Beleslin, Department of Internal Medicine/Endocrinology, Medical faculty, University of Belgrade, Belgrade 11000, Serbia

Corresponding author: Branko Beleslin, FACC, FESC, MD, PhD, Full Professor, Department of Cardiology, Medical Faculty, University of Belgrade, Dr Subotica 8, Belgrade 11000, Serbia. branko.beleslin@gmail.com

Abstract

BACKGROUND

Invasive functional evaluation by fractional flow reserve (FFR) is considered as a gold standard for the evaluation of intermediate coronary stenosis. However, in patients with diabetes due to accelerated progression of atherosclerosis the outcome may be worse even in the presence of negative functional testing.

CASE SUMMARY

We present a case of 55-year-old male diabetic patient who was admitted for chest pain. Diagnostic coronary angiography disclosed 2 intermediate stenoses of the obtuse marginal branch with no evidence of restenosis on previously implanted stent. Patient undergone invasive functional testing of intermediate lesion with preserved FFR (0.88), low coronary flow reserve (1.2) and very high index of microvascular resistance (84). Due to discrepancy in invasive functional parameters, intravascular imaging with optical coherence tomography showed fibrotic stenoses without signs of thin-sup fibroatheroma. Because of the preserved FFR and no signs of vulnerable plaque, the interventional procedure was deferred and the patient continued with optimal medications.

CONCLUSION

Combined functional and anatomic imaging of intermediate coronary stenosis in diabetic patients represent comprehensive contemporary decision pathway in the management of the patients.



Key Words: Fractional flow reserve; Coronary flow reserve; Index of microvascular resistance; Optical coherence tomography; Intermediate coronary stenosis; Case report

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Core Tip: We present a case of the diabetic patient with moderate-to-severe coronary stenosis with preserved fractional flow reserve, low coronary flow reserve, high index of microvascular resistance and intravascular optical coherence tomography image demonstrating fibrotic plaque without signs of thin-cup fibroatheroma. Combined functional and anatomic imaging of intermediate coronary stenosis represent comprehensive contemporary decision pathway in the management of the patients.

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INTRODUCTION

Fractional flow reserve (FFR) is considered as a functional gold standard for the evaluation of coronary stenosis, and the decision for management of intermediate coronary stenosis should be based on functional evaluation[1]. Still, there are some clinical conditions, such as patients with diabetes mellitus (DM) where the value of FFR is still not so well validated and defined. In fact, the previous data have shown that the outcome of deferred lesions according to negative FFR is worse in patients with DM in comparison to non-diabetic patients[2]. In addition, COMBINE optical coherence tomography (OCT) FFR trial has demonstrated additional value of intravascular OCT imaging in stratification of diabetic patients with the negative FFR[3].

CASE PRESENTATION

Chief complaints

We present a case of 55-year-old male diabetic patient who was admitted to our clinic for chest pain syndrome. His chest pain appeared occasionally during last 2 wk and had rather atypical presentation both at rest and after exertion. It was not accompanied by any other symptoms like dyspnea, fatigue or nausea. Due to recent appearance of chest pain of uncertain origin as well as previous implanted stent in left main coronary artery, the patient was referred directly to invasive coronary angiography without non/invasive functional testing.

History of present illness

Three years ago, he suffered from non-ST segment elevation myocardial infarction due to critical lesion of the ostial circumflex (Cx) artery which was treated with implantation of one drug-eluting stent (DES) with crossover from left main to Cx coronary artery (Figure 1). Following DES implantation from left main to Cx, 2 intermediate stenoses of the proximal part of the long second obtuse marginal branch (Figure 1A, orange arrows) were observed that were estimated to be not significant without a need for further intervention. Four years ago, he also had percutaneous coronary intervention with one DES implantation in the right coronary artery (RCA). Since the last coronary event, the patient has been free of any symptom, he was feeling physically and emotionally fit. His medications included aspirin, beta-blockers, and rosuvastatin.

History of past illness

Patient has been obese (102 kg), and has elevated HbA1c levels with no regular anti-diabetic therapy.

Personal and family history

Patient denies smoking, doesn't have hypertension, and denies alcohol abuse. He has elevated cholesterol levels treated with statins.

Physical examination

On physical examination there was no remarkable finding. His blood pressure was 130/80 mmHg. His heart sounds were normal, without significant murmurs. His lung fields were clear, and there were no signs of heart failure. His ECG was normal with sinus rhythm, heart rate of 62/min, without conduction and rhythm disturbances as well as ST-T changes.

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Figure 1 Angiographic images of left coronary artery before (A) and after (B) intervention. White arrow shows stenosis in the ostial part of circumflex artery stenosis, and two orange arrows depicts 2 tandem stenoses in the obtuse marginal branch. Coronary angiography after stenting from left main to circumflex coronary artery.

Laboratory examinations

The laboratory values showed normal blood count, elevated low density lipoprotein (LDL)-cholesterol of 2.3 mmol/L, HbA1c was 6.1%, and the renal function was preserved with estimated glomerular filtration rate of 75 mL/min/1.73m².

Imaging examinations

His cardiac ultrasound was normal with preserved ejection fraction, there were no wall motion abnormalities, the heart chambers were of normal dimensions as well as function and appearance of the valves. Coronary angiography showed no signs of restenosis in the left main to Cx stent, mild ostial stenosis/pinching of left anterior descending artery was unchanged to previous procedure, and 2 moderate-to-severe stenoses in the ostio-proximal part of the second obtuse marginal seem slightly tighter (Figure 2A) than 3 years ago. RCA was without signs of stenosis or restenosis. As there was no non-invasive functional testing for evaluation of myocardial ischemia, the decision was made to perform immediately invasive physiologic evaluation of the Cx obtuse marginal artery using Coroventis platform (Coroventis CoroFow Cardiovascular system, Abbot) and pressure wire (PressureWire X, Abbot) for the invasive functional assessment with FFR during hyperemia with i.v. adenosine, resting full-cycle ratio (RFR), coronary flow reserve (CFR) by thermodilution and index of microvascular resistance (IMR) for the assessment of microvascular dysfunction. FFR was preserved with a value of 0.88, RFR was also preserved with a value of 0.90 but CFR was very low (1.2) with a very high IMR of 84 (Figure 2B). As there was a discrepancy in functional parameters (FFR preserved, CFR low) pointing out to microvascular dysfunction, OCT imaging (DragonFly OpStar imaging catheter, Abbot) was also applied to evaluate the more detailed anatomical characteristics of the lesion. OCT imaging confirmed the significant fibrotic tandem stenoses but without signs of thin-cup fibroatheroma (Figure 3, co-registration of angiographic and OCT imaging, Ultreon 1.0 software for OCT intravascular imaging, Abbot) or plaque vulnerability.

FINAL DIAGNOSIS

The diagnosis was that the patient had non-flow limiting coronary stenoses without signs of vulnerable plaque by OCT, and with significantly deteriorated microvascular function according to low coronary CFR and high IMR, which might be clinically defined as microvascular disease producing angina or chest pain syndrome.

TREATMENT

As the patient had negative or normal FFR value with no intravascular imaging signs of vulnerable plaque, the revascularization was deferred and the patient was recommended to continue with aspirin and to increase dosage of lipid lowering therapy and add ezetimibe to optimize LDL-cholesterol levels. In addition, due to microvascular dysfunction, as demonstrated with very low CFR and very high IMR, diltiazem and ranolazine were added to his therapy. Gluformine was also added to control diabetes.

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Figure 2 Invasive functional parameters of the obtuse marginal stenoses. A: Coronary stenoses in obtuse marginal branch are shown in orange arrows; B: Corresponding fractional flow reserve, coronary flow reserve, and index of microvascular resistance values.



Figure 3 Co-registration images of angiography (left) and optical coherence tomography cross-section imaging (right).

OUTCOME AND FOLLOW-UP

His six-month follow-up was uneventful with very rare atypical chest pain episodes.

DISCUSSION

The optimal diagnostic strategy in patients with diabetes remains controversial and challenging. The previous studies have shown low event rates in asymptomatic patients but also disparities (different cardiovascular risk groups, different non-invasive and invasive testing) in the management and screening of those patients[4]. According to the latest European Society of Cardiology guidelines^[5], functional stress testing and computed tomography angiography may be considered and indicated in patients with diabetes as well. On the other hand, earlier studies with stress echocardiography have demonstrated that significantly more patients with diabetes have ischemia on stress testing, and that the prognosis in patients without ischemia is still worse in diabatic than non-diabatic patients [6]. In addition, noninvasively measured (2D echocardiography) CFR provides independent prognostic information in diabetic and non-diabetic patients and negative dipyridamole stress echocardiography - an abnormal CFR is associated with almost twice worse prognoses in diabetic in comparison to non-diabetic patients^[7]. Finally, Murthy *et al*^[8] have demonstrated that abnormal CFR by positron emission tomography is independent predictor of cardiac mortality in patients with diabetes and without known coronary artery disease.

Regarding invasive functional testing, Kennedy et al^[2] has shown that deferred revascularization in 250 patients according to FFR have been connected with worse outcome in patients with diabetes in comparison to non-diabetic patients, in terms of significant more revascularizations (16% vs 6%) in the mean follow-up time of 40 months. Although earlier studies [9,10] have not shown significant differences in the outcome between diabetic and non-diabetic patients according to FFR values this issue remains controversial. Therefore, COMBINE OCT FFR[3] investigated the impact of OCT-detected thin-cap fibroatheroma (TCFA) on clinical outcomes of patients with diabetes patients with FFR negative lesions. The study [3] revealed that among 550 enrolled diabetic patients with \geq 1 FFR-negative lesions, TCFA-positive patients represented 25% of this population and were associated with a five-fold higher rate of MACE despite the absence of ischemia as demonstrated by negative FFR > 0.80. This discrepancy between the impact of vulnerable plaque and



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ischemia on future adverse events may explain worse outcome in diabetic patients with negative FFR. On the other hand, diabetic patients with negative FFR and without TCFA remained to have very low event rate of 3.1% over mean followup of 18 months[3]. Still the question remains on the outcome of vulnerable plaques due to the low hard event rates and limited positive predictive value[11]. Regarding therapeutic options, serial intravascular imaging has demonstrated that lipid lowering therapy led to regression of coronary plaque volume and increase in fibroatheroma cup thickness proportional to decrease in cholesterol levels^[12].

Our patient presented with anatomically significant but functionally non-significant coronary lesion without signs of plaque vulnerability as shown by thick-cap fibroatheroma, but with significant disturbance of coronary microcirculation as shown by low CFR and very high IMR. The microcirculatory dysfunction may be present in diabetic patients and the cause of positive functional testing and chest pain[13]. In fact, the traditional risk factors may all contribute to coronary microvascular dysfunction and structural remodeling of the microcirculation. In the WISE study [14], chest pain, diabetes, smoking, CAD severity were all independent predictors of hard cardiovascular events. The evaluation of microcirculatory dysfunction has gained a lot of interest in the last years and specific algorithms have been developed to interrogate this clinical condition[15]. Optimal therapy for microcirculatory dysfunction has not been established yet but calcium channel blockers, nebivolol, ranolazine, trimetazidine, ACE inhibitors have been associated with beneficial effects in microvascular angina[15].

CONCLUSION

The invasive functional evaluation combined with intravascular imaging have complementary diagnostic and prognostic role and therefore a contemporary evaluation of coronary stenosis, particularly in diabetic patients whenever possible and available should include both functional and intravascular imaging data for the best patient management and decision making.

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ORCID number: Arif Al Nooryani 0000-0002-0329-7954; Wael Aboushokka 0000-0002-6746-6742; Branko Beleslin 0000-0001-9452-8241; Biljana Nedeljkovic-Beleslin 0000-0002-1687-9297.

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