Evaluation of the impact of upstream clopidogrel on time to coronary artery bypass grafting (CABG)

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INTRODUCTION

- Administration of dual antiplatelet therapy (DAPT) with aspirin and a P2Y12 inhibitor is an American College of Cardiology/ American Heart Association guideline recommendation for the management of non-ST elevation myocardial infarction acute coronary syndrome (NSTE-ACS) and ST elevation myocardial infarction (STEMI).^{1,2}
- The optimal timing of administration for P2Y12 inhibitors in the setting of ACS is unknown; however, some studies suggest "upstream" administration is associated with a reduction in cardiovascular events (e.g., stent thrombosis, periprocedural myocardial infarction) as compared to provisional administration after coronary angiography.¹
- The disadvantage of upstream administration is the potential to delay CABG while the P2Y12 inhibitor has a washout period (~5 days), resulting in added hospital costs and increasing the risk of iatrogenic complications.³

PURPOSE

To evaluate whether the time to in-hospital CABG was impacted by upstream clopidogrel therapy for patients presenting to the University of Maryland Medical Center with either NSTE-ACS or STEMI.

PATIENT POPULATION

Inclusion Criteria

- Ages <u>></u> 18 years old
- Admission diagnosis of NSTE-ACS or STEMI
- In-hospital CABG

Exclusion Criteria

- Admission diagnosis of unstable angina
- Prior P2Y12i maintenance therapy
- Elective CABG
- Delay to CABG resulting from complications by another disease state

METHODS

- Retrospective chart review conducted on records between June 2011 March 2017
- Pool of potential subjects identified through a search for ICD 9 and 10 codes for CABG, coronary angiography and cardiac catherization
- Primary endpoint:
 - Mean number of days from presentation to CABG between patients who did or did not receive upstream clopidogrel therapy
- Secondary endpoints:
 - Mean number of days from presentation to CABG stratified by STEMI vs NSTE-ACS patients
 - Mean number of days from last dose of clopidogrel to CABG stratified by STEMI vs NSTEMI patients
- Statistical analysis
 - Differences in outcomes between exposures were detected using two-way analysis of variance (ANOVA) and t-tests

RESULTS

Table 1 Patient Demographics						
Characteristics	Total (%), n=63	Upstream clopidogrel (n=34)	No upstream clopidogrel (n=29)	p value		
Age (years)						
	64.4	64	64.9	0.8		
Gender						
Male Female	71.4 28.6	70.6 29.4	72.4 27.6	0.9		
History of CAD						
Yes No	73 27	32.3 67.7	20.7 79-3	0.3		
History of MI		1	1			
Yes No	17.5 82.5	14.7 85.3	20.7 79-3	0.7		
Prior aspirin therapy						
Yes No	39.7 60.3	38.2 61.8	41.4 58.6	0.8		
Type of MI						
STEMI NSTE-ACS	28.6 71.4	23.5 76.5	34.5 65.5	0.3		
Percutaneous mechanical circulatory support use						
Yes No	50.8 49.2	38.2 61.8	65.5 34.5	0.03		

Table 2. Results						
Primary Endpoints	Mean Days (95% Cl)					
	Clopidogrel exposure	No exposure	p value			
From presentation to CABG	3.5 (2.8,4.3)	3.2 (2.3, 4.0)	0.5			
Secondary Endpoints	Mean Days (SD)					
	STEMI	NSTE-ACS	p value			
From presentation to CABG	2.2 (2.0)	3.8 (2.3)	0.009			
From last dose of clopidogrel to CABG	2.6 (1.6)	2.6 (1.7)	0.9			



CONCLUSIONS

REFERENCES

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- 63 patients included in the final analysis with 34 exposed to upstream clopidogrel and 29 without exposure.
- The mean number of days from presentation to CABG in patients exposed to clopidogrel vs not exposed was 3.5 days (95% Cl 2.8-4.3), and 3.2 days (95% CI 2.3-4.0; p=0.5, respectively
- Aside from fewer patients on percutaneous mechanical support receiving upstream clopidogrel, baseline characteristics were the similar across study arms

• Upstream administration of the P2Y12 inhibitor clopidogrel did not result in a delay to CABG for patients presenting with either NSTE-ACS or STEMI who were deemed surgical candidates

• Despite STEMI patients having an overall faster time to CABG from presentation, the mean number of days from last dose of clopidogrel did not differ between STEMI and NSTE-ACS patients, and therefore did not appear to be impacted by early exposure to clopidogrel

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