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Laura Neubauer

*Baptist Hospital of Miami*, LauraNeu@baptisthealth.net

Radhan Gopalani

*Baptist Hospital of Miami*, radhang@baptisthealth.net

Kristen de Almeida

*Baptist Hospital of Miami*, KristenDe@baptisthealth.net

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# Pharmacists' Role in the Management of Patients Receiving Dual or Triple Antithrombotic Therapy

Laura Provost, PharmD, BCPS  
PGY2 Cardiology Pharmacy Resident | Baptist Hospital of Miami  
[LauraNeu@baptisthealth.net](mailto:LauraNeu@baptisthealth.net)



# Disclosures

The authors for this project have nothing to disclose regarding any financial or nonfinancial relationships with the products described, reviewed, or evaluated in this presentation



# Abbreviations

- ACC • American College of Cardiology
- ACS • Acute Coronary Syndrome
- ADR • Adverse Drug Reaction
- APT • Antiplatelet
- ASA • Aspirin
- BHM • Baptist Hospital of Miami
- CAD • Coronary Artery Disease
- COVID • Coronavirus Disease
- DAT • Dual Antithrombotic Therapy
- DAPT • Dual Antiplatelet Therapy
- DOAC • Direct Oral Anticoagulant
- DVT • Deep Vein Thrombosis
- MVR • Mitral Valve Repair
- NVAf • Nonvalvular Atrial Fibrillation
- PAD • Peripheral Artery Disease
- PE • Pulmonary Embolism
- OAC • Oral Anticoagulant
- TAT • Triple Antithrombotic Therapy
- TAVR • Transcatheter Aortic Valve Replacement

# Objectives



1

Summarize the most recent literature describing optimal antithrombotic therapy management in patients with an indication for both APT and OAC therapies

2

Review reported DAT / TAT-related ADRs at BHM during the year 2019

3

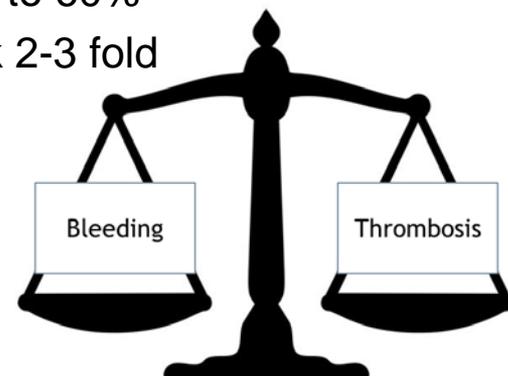
Discuss the clinical impact of pharmacist interventions in optimizing antithrombotic therapy in patients at BHM



# Background

# Background

- Patients requiring concomitant APT and AC therapy have  $\uparrow$  risk of both thrombotic and bleeding events
- Major bleeding is associated with 5x  $\uparrow$  risk of death after ACS
  - Addition of single APT to OAC  $\uparrow$  bleed risk up to 60%
  - Addition of dual APT to OAC further  $\uparrow$  this risk 2-3 fold
- Goal: Mitigate bleed risk while maintaining antithrombotic efficacy



# Summary of Literature

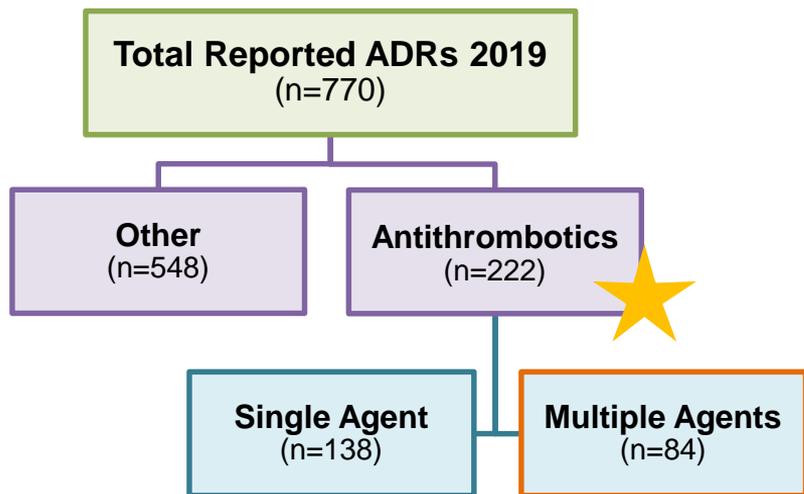
- Management of combination antithrombotic therapy is driven by:
  - Patient parameters, medication factors, guideline recommendations, indication, etc.
- Evolution of Triple Antithrombotic Therapy (TAT)



*Most recent recommendations state that TAT be used for shortest duration possible (most often limited to duration of hospitalization)*

# Antithrombotic-related ADRs at BHM

- Antithrombotics were #1 medication class associated with ADRs in 2019 (~29%)
- 28% of these ADRs included DAT or TAT regimens ( $\geq 1$  APT + IV/PO AC)



| Antithrombotic regimens including multiple agents | n=84 |
|---|------|
| DAT or TAT Regimens                               | 62   |
| • APT + IV AC                                     | 4    |
| • APT + OAC                                       | 40   |
| • 2 APT + OAC                                     | 6    |
| • 2 APT + IV AC                                   | 12   |
| DAPT  | 16   |
| AC + AC (Bridge therapy)                          | 6    |

>75% of ADRs were community-acquired



# Research Purpose



To evaluate pharmacists' role in the optimization of antithrombotic therapy in patients with concomitant indications for APT and OAC agents



# Methods



# Study Design



Single center



Quality  
Improvement



Prospective

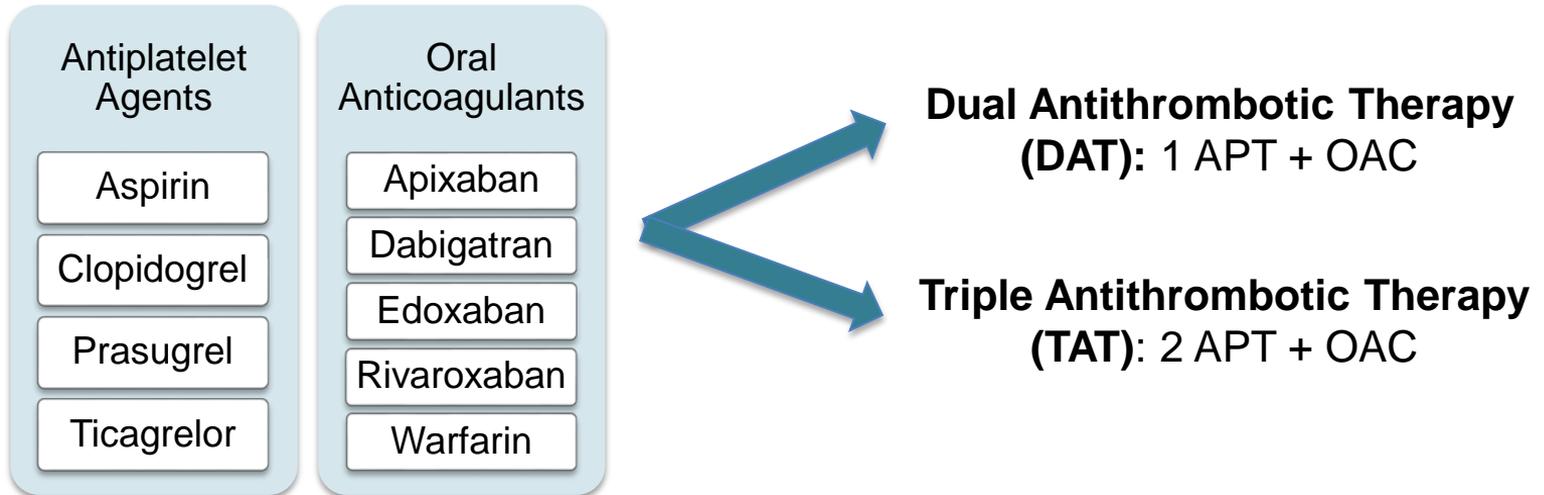


January – March  
31<sup>st</sup> 2021



# Eligibility Criteria

Hospitalized adult patients at Baptist Hospital receiving both an APT agent and OAC





# Data Collection

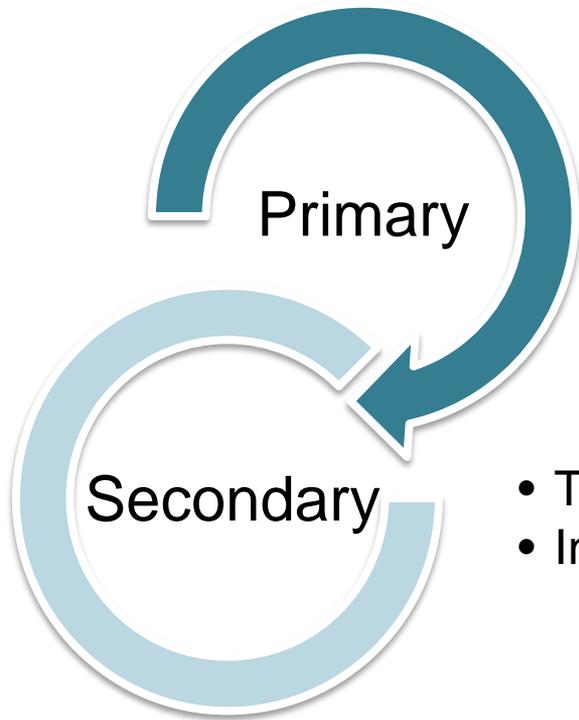
Daily report of hospitalized patients with active order for  $\geq 1$  antithrombotic agent

Filtered to include only patients receiving DAT or TAT

Recommendations made based on clinical guidelines, indications & patient characteristics



# Study Outcomes



- Appropriateness of AC and APT regimens based on specific indications

- Type of pharmacist intervention
- Intervention acceptance rate



# Results



# Baseline Demographics

| n= 239                                |             |
|---------------------------------------|-------------|
| Mean age, years (SD)                  | 74.3 ± 12.1 |
| Gender – male, n (%)                  | 136 (56.9)  |
| Regimen – n (%)                       |             |
| • Dual Therapy                        | 228 (95.4)  |
| • Triple Therapy                      | 11 (4.6)    |
| APT Indication – n (%)                |             |
| • CAD                                 | 130 (54.4)  |
| • COVID                               | 28 (11.7)   |
| • CVA                                 | 22 (9.2)    |
| • ACS                                 | 19 (7.9)    |
| • S/p Valve surgery                   | 16 (6.7)    |
| • PAD                                 | 11 (4.6)    |
| • Not documented                      | 7 (2.9)     |
| • 1 <sup>o</sup> prev. / ↑ ASCVD risk | 6 (2.5)     |
| OAC Indication – n (%)                |             |
| • Atrial Fibrillation                 | 172 (71.9)  |
| • DVT/PE treatment                    | 41 (17.2)   |
| • DVT/PE secondary ppx                | 15 (6.3)    |
| • Other*                              | 11 (4.6)    |

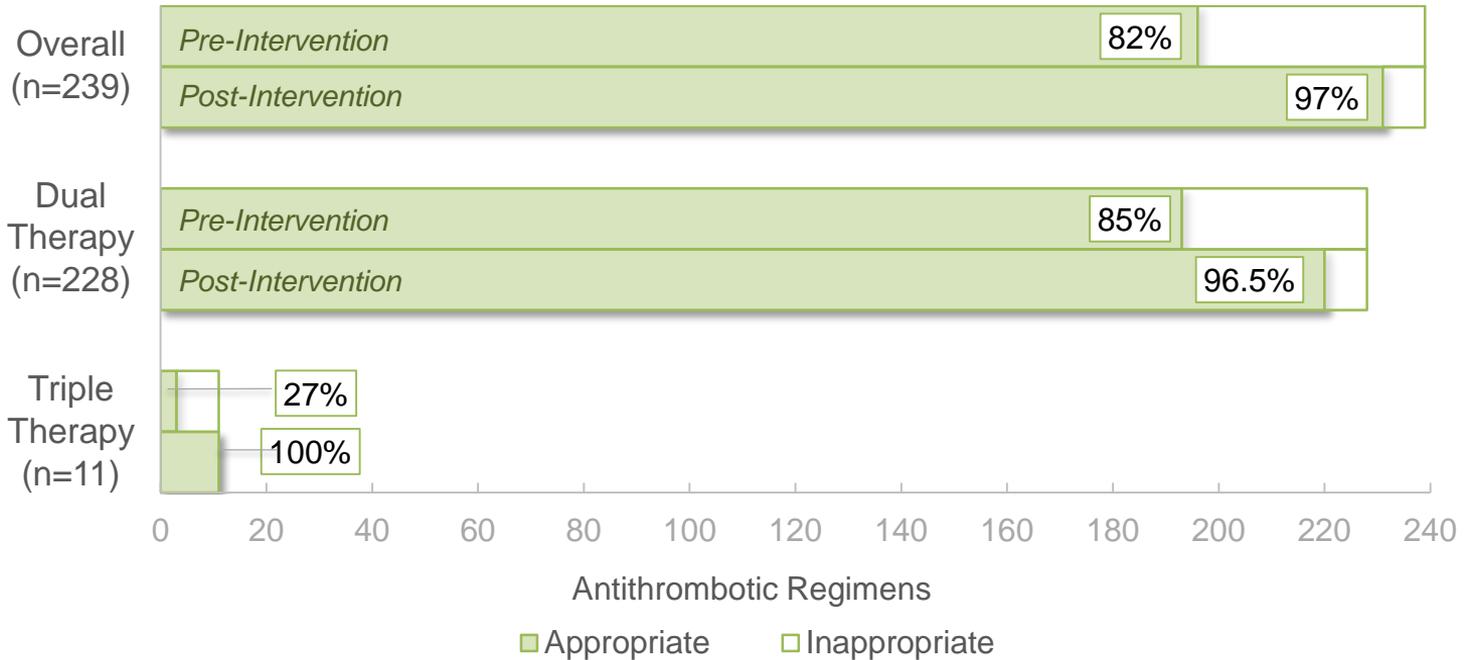
| Antithrombotic Agents n= 239                          |     |
|---|-----|
| Antiplatelet Agents                                   |     |
| • ASA   | 196 |
| • Clopidogrel   | 49  |
| • Ticagrelor  | 5   |
| <i>*11 patients were receiving 2 APT agents (TAT)</i> |     |
| Oral Anticoagulants                                   |     |
| • Apixaban  | 184 |
| • Rivaroxaban   | 36  |
| • Warfarin  | 12  |
| • Dabigatran  | 7   |

\*Intracardiac thrombus, valve surgery, PAD, portal vein thrombosis, s/p hip/knee replacement



# Primary Outcome

## Regimen Appropriateness





# Secondary Outcomes

## Pharmacist Interventions

|                               | Interventions Recommended | Interventions Accepted |
|-------------------------------|---------------------------|------------------------|
| Discontinue Agent             | 19                        | 18                     |
| • APT                         | 16                        | 15                     |
| • OAC                         | 3                         | 3                      |
| Increase OAC Dose             | 14                        | 11                     |
| • Apixaban                    | 11                        | 9                      |
| • Rivaroxaban                 | 2                         | 2                      |
| • Dabigatran                  | 1                         | 0                      |
| Decrease OAC Dose             | 10                        | 6                      |
| • Apixaban                    | 5                         | 2                      |
| • Rivaroxaban                 | 5                         | 4                      |
| Alternative Agent Recommended | 4                         | 4                      |
| <b>Total</b>                  | <b>47</b>                 | <b>39 (83.0%)</b>      |





# Secondary Outcomes

## Pharmacist Interventions

|                               | Interventions Recommended | Interventions Accepted |
|-------------------------------|---------------------------|------------------------|
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| • APT                         | 16                        | 15                     |
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# Secondary Outcomes

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| • Apixaban                    | 5                         | 2                      |
| • Rivaroxaban                 | 5                         | 4                      |
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# Secondary Outcomes

## Pharmacist Interventions

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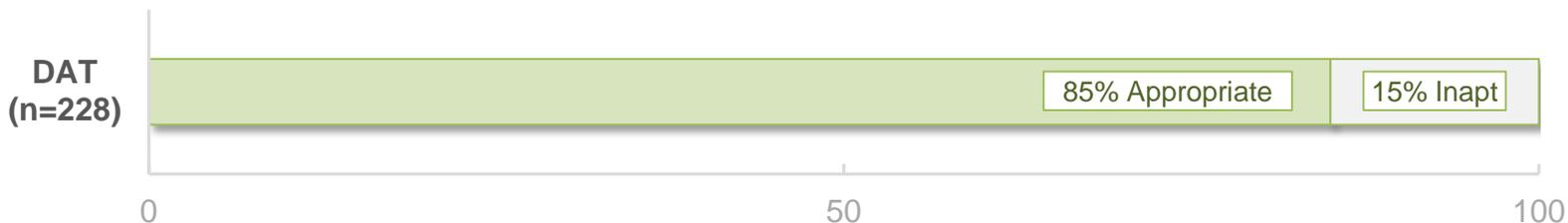
# Secondary Outcomes

## Pharmacist Interventions

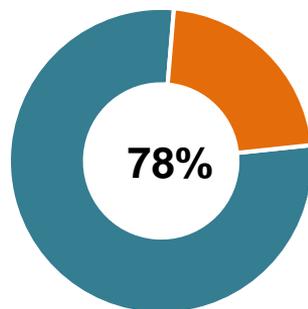
|                               | Interventions Recommended | Interventions Accepted |
|-------------------------------|---------------------------|------------------------|
| Discontinue Agent             | 19                        | 18                     |
| • APT                         | 16                        | 15                     |
| • OAC                         | 3                         | 3                      |
| Increase OAC Dose             | 14                        | 11                     |
| • Apixaban                    | 11                        | 9                      |
| • Rivaroxaban                 | 2                         | 2                      |
| • Dabigatran                  | 1                         | 0                      |
| Decrease OAC Dose             | 10                        | 6                      |
| • Apixaban                    | 5                         | 2                      |
| • Rivaroxaban                 | 5                         | 4                      |
| Alternative Agent Recommended | 4                         | 4                      |
| <b>Total</b>                  | <b>47</b>                 | <b>39 (83.0%)</b>      |



# Dual Antithrombotic Regimens



## Intervention Acceptance Rate



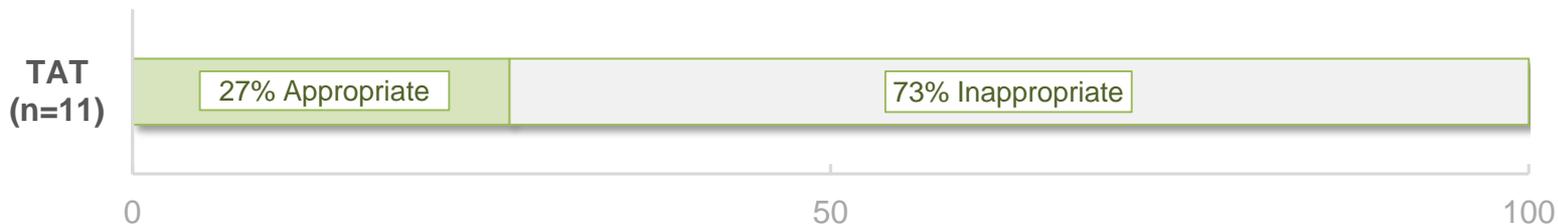
■ Accepted ■ Rejected



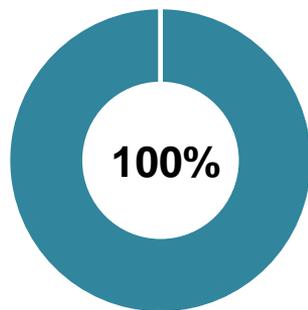
| Interventions Rec'd / Accepted |                |
|--------------------------------|----------------|
| Increase OAC                   | 12 / 9         |
| Decrease OAC                   | 9 / 5          |
| Discontinue Agent              |                |
| • APT                          | 9 / 8          |
| • OAC                          | 3 / 3          |
| Alternative Agent Rec'd        | 3 / 3          |
| <b>Total*</b>                  | <b>36 / 28</b> |

\*35 unique regimens with 36 interventions

# Triple Antithrombotic Therapy



## Intervention Acceptance Rate



■ Accepted



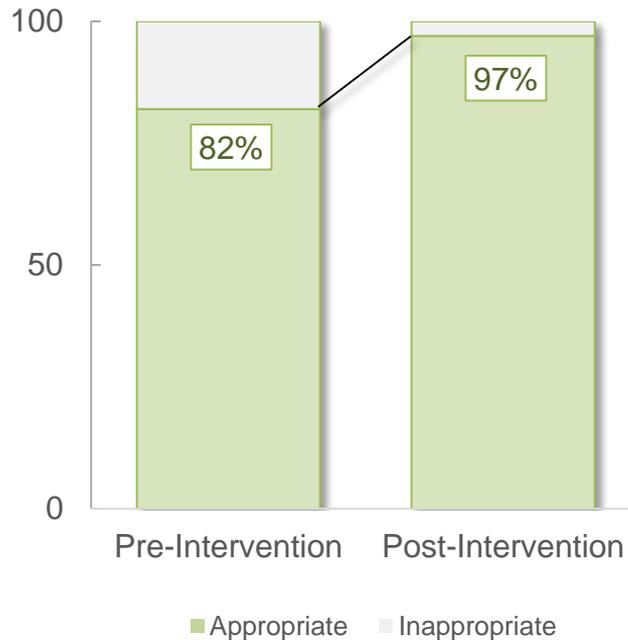
| Interventions Rec'd / Accepted |                |
|--------------------------------|----------------|
| Discontinue APT                | 7 / 7          |
| Increase OAC                   | 2 / 2          |
| Decrease OAC                   | 1 / 1          |
| Alternative Agent Rec'd        | 1 / 1          |
| <b>Total*</b>                  | <b>11 / 11</b> |

\*8 unique regimens with 11 interventions

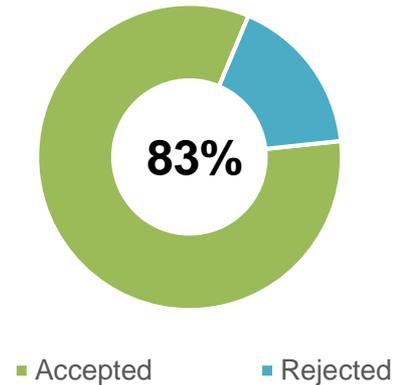


# Impact of Pharmacist Intervention

## Regimen Appropriateness



## Intervention Acceptance Rate





# Discussion

- Dual antithrombotic therapy (APT + OAC) accounted for 95% of all regimens reviewed
- Triple antithrombotic therapies had the greatest opportunity for optimization
  - 73% inappropriate pre-intervention, 100% of interventions accepted
- Type of pharmacist intervention
  - Most common: APT discontinuation
  - Most clinically significant: OAC discontinuation



# Limitations

- Short study duration (10 weeks)
- Warfarin pharmacy consult and pre-approved automatic dose adjustment protocol in place to facilitate OAC dose optimization
- Patients assessed for therapy optimization only once
- Unable to assess long-term impact of RPh intervention on adverse outcomes



# Conclusion

Pharmacist intervention resulted in optimization of dual / triple antithrombotic therapies by 15%, with greatest impact seen with triple antithrombotic therapy

- Next steps:
  - Implement an automated alert that requires providers to acknowledge the use of triple therapy as well as intended duration
  - Pharmacist re-education regarding indication-specific dosing of OACs and indicated duration of therapy



# Acknowledgements

- Radhan Gopalani, PharmD, BCPS, BCCP
- Kristen de Almeida, PharmD, BCCP
- Ian Del Conde-Pozzi, MD



# Self Assessment Question

When indicated, what is the most appropriate duration of triple antithrombotic therapy for most patients?

- A. Until hospital discharge
- B. Indefinitely
- C. 1 year
- D. Triple therapy is never appropriate



# References

- Angiolillo DJ, Goodman SG, Eikelboom JW, et al. Antithrombotic Therapy in Patients with Atrial Fibrillation With Oral Anticoagulation Undergoing Percutaneous Coronary Intervention. *Circulation*. 2018;138:527-536.
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PGY2 Cardiology Pharmacy Resident | Baptist Hospital of Miami  
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