Effect of the COVID-19 Pandemic and Other Predictors of True Therapeutic Inertia on Patients with Hypertension in a Primary Care Clinic in Thailand

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Introduction: Hypertension (HT) has a significant impact on health care worldwide. Therapeutic inertia (TI) is defined as the failure to intensify therapy in the absence of an optimal goal and is widely used as a quality of care parameter. The coronavirus disease 2019 (COVID-19) pandemic has affected many health-care systems, including HT care. Therefore, the present study assessed the impact of the COVID-19 pandemic on TI and its predictors in patients with HT.

Methods: The electronic medical records of patients with HT who attended a primary care clinic at a tertiary hospital during pre-COVID-19 (February 2019 to February 2020) and COVID-19 (March to August 2020) periods were reviewed.

Results: Our study included 6089 visits during the 12-month pre-COVID-19 period and 2852 visits during the 6-month COVID-19 period. Most of the baseline characteristics of the HT patients were not significantly different between the two time periods. During the COVID-19 period, the percentage of uncontrolled HT visits decreased from 43% to 31%. Similarly, the prevalence of TI decreased from 81% to 77%. False TI was predominantly due to physicians’ concerns regarding the in-clinic blood pressure measurement being inaccurate during both the periods.

Conclusion: After readjustment for the physicians’ reasons, the true TI was 64% and 60% in the pre-COVID-19 and COVID-19 period. For adjusted physician and patient-related factors, multilevel modeling was used. Senior medical staff visits, elderly patients, prior diabetes mellitus diagnosis, patients who used more than one type of anti-HT medication, and patients with systolic blood pressure >150 mmHg were all predictors of TI. The COVID-19 period, on the other hand had no effect on TI with an adjusted odds ratio of 0.82 (95% confidence interval, 0.67–1.01).

Keywords: hypertension, therapeutic inertia, COVID-19, quality of care, multilevel modeling, primary care

Introduction
Hypertension (HT) is a major noncommunicable disease (NCD) that causes significant mortality and morbidity worldwide. Less than half of all patients have controlled HT, despite being aware of the importance of optimal blood pressure (BP). As a result, the World Health Organization (WHO) has incorporated HT treatment quality into the Global Monitoring Framework for NCDs. A previous study highlighted three major factors that influence the quality of HT care: patients,
Therapeutic inertia (TI) is defined as failure of the health-care provider to intensify therapy in the absence of an optimal goal, such as inadequately adjusting anti-HT medications to counteract increasing BP.6,7 TI is widely used as a parameter to monitor the continued quality of HT care.8–10 Reducing TI in primary care might reduce its prevalence in populations with uncontrolled BP.11–13 The severity of HT, concern about adverse events, comorbidities, complexity of HT treatment guidelines, mean higher age of the patients, insurance, and treatment cost have been identified as predictors of TI.13,14

Patients’ access to care has been affected by the coronavirus disease 2019 (COVID-19) pandemic due to the need for social distancing and lockdown measures.15,16 A rapid assessment survey from the WHO showed that >50% of HT services were completely or partially disrupted due to the pandemic.17 Furthermore, 40% of NCD-related staff were deployed to provide COVID-19 relief. No previous studies have investigated whether the COVID-19 pandemic has influenced TI in HT care and whether TI predictors were the same during pre-COVID and COVID-19 periods. Understanding these issues is essential for health-care planners to adjust the monitoring of quality of care in response to COVID-19.

The present study evaluated the effect of the COVID-19 pandemic on TI among patients with HT in a primary care clinic (PCC) at a tertiary hospital in southern Thailand. The relationship between patient and physician-related factors on TI was also analyzed.

Methods

Study Design and Setting, Participants, and the COVID-19 Situation

We performed a retrospective cross-sectional study that included HT patients’ visits to a PCC at the medical school at a tertiary hospital in Thailand. Most physicians in the PCC were family physicians, general practitioners, and internists working at the Department of Family and Preventive Medicine. All medical records in the hospital were fully computerized and retrievable through the Division of Digital Innovation and Data Analytics (DIDA) at the authors’ institute.

COVID-19 cases were identified in Thailand at the beginning of 2019 and the disease spread widely from approximately March to April 2019. Therefore, the Thai government implemented measures to restrict interprovincial transportation and lock down high-risk areas.15 Postal prescriptions were provided to NCD patients undergoing continuous care who were unable to attend the hospital for any reason from April 2019 at the PCC.

Data Collection

We defined two study periods as the pre-COVID-19 period (February 2019 to February 2020) and COVID-19 period (March to August 2020). Data were anonymously extracted from the hospital database by DIDA. We identified HT visits from records with ICD-10 of Essential (primary) HT (I10) and/or use of anti-HT prescription. All medical records were reviewed.

The inclusion criteria were HT patients receiving continuous care at the PCC for more than two visits within the last six months. The exclusion criteria were records without BP records, patients whose relatives received medicine on their behalf, postal prescription, and nurse home visit prescription.

Data were under relational structure with each visit used as the main unit of analysis. Data regarding the patients’ characteristics, such as gender, age, religion, insurance schemes, and comorbidities, were retrieved from the first registration visit. Data regarding the physicians’ characteristics were extracted from the hospital database. On different occasions, each patient was seen by more than one physician and a physician usually attended to more than one patient.

Operational Definitions of Key Variables

1. Uncontrolled HT was defined as systolic blood pressure (SBP) ≥140 mmHg and/or diastolic blood pressure (DBP) ≥90 mmHg according to current guidelines.18,19 Uncontrolled HT was identified by the selection of records in which these criteria were met.

2. TI was defined as uncontrolled HT in a patient who did not receive anti-HT treatment. This criterion did not apply to false TI in which the physician was fully aware of the uncontrolled HT yet did not add anti-HT medication for the following reasons (adapted from a previous study):7
   (i) self-measured blood pressure monitoring (SMBP) at home data suggested that BP was well controlled at home;
   (ii) concerns of in-clinic BP being inaccurate (data for repeat BP measurements in-clinic were shown in medical records);
   (iii) patients refused to take additional medication;
(iv) patients with a history of orthostatic hypotension; and
(v) anti-HT medication ran out (the patient had not been properly medicated before the visit).

The present study defined these patients as true TI if these reasons could not be found in the medical records. The calculation of TI score was modified from the previous study. The patients who came to hospital more than two times in the 6 months of period were used in this study. Furthermore, true TI was used to calculate the TI score, which was defined as the adapted TI score.\textsuperscript{8,20}

**Statistical Analysis**

Analyses were performed using R software version 4.0.2 with epiDisplay and lme4 packages.\textsuperscript{21,22} Baseline characteristics of the patients were compared across the two COVID-19 periods using \textit{t} test for continuous variables and chi-squared test for categorical variables to examine potential confounding roles of characteristics of the patients and the physician. The COVID-19 period was used as the stratification factor to examine the effects of other variables on TI, and predicting variables were compared between true TI and non-TI visits. The odds ratios (OR) from two strata were pooled together to calculate the period-adjusted OR of each variable using the Mantel–Haenszel method. Stratified linear regression was used for continuous data to estimate the pooled mean of that variable stratified by the COVID-19 period. Finally, multilevel logistic regression simultaneously considering the two factors was performed using the Linear Mixed-Effects Models in the lme4 package to adjust the effects of physician-related and patient-related variables. The final adjusted OR of COVID-19 period on TI, adjusted for both physician and patient-related variables and their 95% confidence interval (95% CI) were reported. Statistical significance was considered when \( P \)-values were <0.05.

**Results**

There were 6089 visits during the 12-month pre-COVID-19 period and 2852 during the 6-month COVID-19 period, and the average monthly visits were only reduced by 6% (Figure 1). However, during the 6-month COVID-19 period, 17.5% of the records indicated patients’ absence at the clinics and 13% of visits included relatives who came to collect the medication prescribed physicians on behalf of the patients. Furthermore, 4% of patients had their medication delivered by post. These patients were not eligible for assessment of TI.

Table 1 shows a comparison of the baseline characteristics of HT patients who visited an attending physician at the PCC during the pre-COVID-19 and COVID-19 periods. There were no significant differences between the variables during the two periods except the percentage of thiazide prescribed and the percentage of prescriptions by

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**Figure 1** Patient enrollment between the pre-COVID-19 and COVID-19 periods.

**Note:** *Relative* = patients whose relatives received.
Table 2 shows the BP parameters and TI during the pre-COVID-19 and pre-COVID-19 periods. The mean SBP and DBP were significantly lower during the COVID-19 period. The percentage of uncontrolled HT visits decreased from 43% to 31% during the COVID-19 period. Similarly, the prevalence of TI decreased from 32.5% to 24.5% during the COVID-19 period. The percentage of uncontrolled HT visits decreased from 43% to 31% during the COVID-19 period.
81% to 77%. False TI was predominantly due to physicians’ concerns about the in-clinic BP being inaccurate during both periods and were not significantly different ($P$-value = 0.434). After readjustment for the physicians’ reasons, the true TI was 64% during the pre-COVID-19 period and 60% during the COVID-19 period. The true TI and TI scores were significantly lower during the COVID-19 period. For convenience, we used the true TI value as TI in subsequent analyses.

Table 3 illustrates the relationships between TI and patient and physician-related factors stratified by COVID-19 period. Since the interaction between the COVID-19 period and these variables was not significant (details of testing interaction effects are omitted), the pooled OR based on the Mantel–Haenszel method for categorical variables and the period-adjusted effect of continuous variables are displayed. TI visits were an average of 5.8 mmHg (95% CI 5.0–6.5) lower in SBP and 2.3 mmHg (95% CI 3.0–5.0) lower in DBP compared with non-TI visits. Patients using ACE-inhibitors, second-line anti-HT medications, and those with a prior diagnosis of diabetes mellitus (DM) were associated with TI. TI visits were also attended by physicians with an average of 2.5 years more work experience and who were 2.5 years older than those attending non-TI visits. Senior medical staff (>10 years’ experience) showed an increased risk of true TI. On the other hand, internists were associated with a protective effect, with an adjusted OR of 0.5 (95% CI 0.32–0.85).

Table 4 shows the final model predicting TI using multilevel (lme4) modeling after simultaneously and mutually adjusting for physician and patient-related factors. Five significant predictors for TI included senior medical staff visits, elderly patients, patients with prior DM diagnosis, patients who used more than one type of anti-HT medication, and patients with SBP >150 mmHg. The COVID-19 period had an adjusted OR of 0.82 (95% CI 0.67–1.01). Thus, there was insufficient evidence to state whether the COVID-19 period was associated with an increase in TI.

Discussion
Statement of Principal Findings
In our clinical setting, the COVID-19 pandemic resulted in nearly one-sixth of prescriptions being made without the physician really seeing the patients. The demographic backgrounds of the patients and the physicians during the COVID-19 period were not significantly different from those in the preceding period. On the other hand, patients visiting the PCC during the COVID-19 period had lower SBP, DBP, uncontrolled HT and greater use of thiazides. Most cases of false TI were due to physicians’ concerns about the inaccuracy of the in-clinic BP during both periods. During both periods, TI was consistently more common in elderly patients and those with a prior DM diagnosis. Patients with SBP >150 mmHg were three times less likely to experience TI compared with those...
### Table 3 Therapeutic Inertia by Patient and Physician-Related Characteristics and COVID-19 Period

#### I. Patients' characteristics

| Characteristics                  | Pre-COVID-19 | COVID-19 | P-value | MH–OR (95% CI) * |
|----------------------------------|--------------|----------|---------|------------------|
|                                  | No TI (n = 916) | TI (n = 1653) |         |                  |
| Mean age (SD), years             | 67.1 (10.4)  | 68 (10.5) | 0.050   |                  |
| Male, no. (%)                    | 290 (31.7)   | 558 (33.8) | 0.299   |                  |

| Medical schemes, no. (%)         |               |          |         |                  |
|----------------------------------|---------------|----------|---------|------------------|
| Universal coverage               | 369 (40.3)    | 674 (40.8) | 0.931  |                  |
| Civil servant medical benefit    | 416 (45.4)    | 753 (45.6) | 0.999  |                  |
| Social security                  | 33 (3.6)      | 49 (3)   | 0.999  |                  |
| Cash                             | 65 (7.1)      | 120 (7.3) | 0.999  |                  |
| Other                            | 33 (3.6)      | 57 (3.4) | 0.999  |                  |
| Median body mass index (IQR)     | 26.3 (23.9,30.1) | 26.2 (23.6,29.5) | 0.048 |                  |
| Median SBP, (IQR) mmHg           | 152 (146,160) | 146 (142,153) | <0.001 |                  |
| Median DBP, (IQR) mmHg           | 78.8 (10.8)   | 78.8 (10.5) | <0.001 |                  |

| Comorbidity, no. (%)             |               |          |         |                  |
|----------------------------------|---------------|----------|---------|------------------|
| Diabetes mellitus                | 362 (39.5)    | 755 (45.7) | 0.003  |                  |
| Dyslipidemia                     | 856 (93.4)    | 1526 (92.3) | 0.327 |                  |
| Cerebrovascular disease          | 26 (2.8)      | 62 (3.8) | 0.269 |                  |
| Chronic kidney disease stage ≥3  | 43 (26.2)     | 75 (25.7) | 0.542 |                  |

| Anti-HT medications, no. (%)    |               |          |         |                  |
|----------------------------------|---------------|----------|---------|------------------|
| ACE-I                            | 329 (35.9)    | 531 (32.1) | 0.056  |                  |
| ARBs                             | 344 (37.6)    | 573 (34.7) | 0.155  |                  |
| CCBs                             | 595 (65)      | 1011 (61.2) | 0.963 |                  |
| Thiazide                         | 257 (28.1)    | 454 (27.5) | 0.873  |                  |
| Alpha-blocker                    | 43 (4.7)      | 68 (4.1) | 0.554 |                  |
| Second-line drugs                | 22 (2.4)      | 17 (1)   | 0.011  |                  |
| Combined pill                    | 40 (4.4)      | 87 (5.3) | 0.363 |                  |
| ≥2 types of anti-HT medication   | 561 (61.2)    | 850 (51.4) | <0.001 |                  |

| 2. Physicians' characteristics   |               |          |         |                  |
|----------------------------------|---------------|----------|---------|------------------|
| Median age (IQR), years          | 36 (30,41)    | 37 (34,41) | <0.001 |                  |
| Median work experience (IQR), years | 12 (6,18)    | 13 (12,18) | <0.001 |                  |

* MH–OR (95% CI): Multivariate hazard ratio (95% confidence interval).
with SBP of 141–150 mmHg. The adjusted OR of TI also increased two-fold among patients attended by senior staff. Patients who used more than one type of anti-HT therapy were less likely to experience TI compared with those using only one type. Finally, there was no relationship between COVID-19 period and TI after adjusting for factors using the multilevel model.

**Interpretation Within the Context of the Wider Literature**

The effect of COVID-19 may not contribute to demographic factors. However, our study found that BP parameters, percentage of uncontrolled HT visits, and TI of patients during the COVID-19 period were significantly better than during the pre-COVID-19 period. Patients who are socioeconomically vulnerable are more likely to become non-adherent in the absence of effective public health interventions. In the present study, the government measures during the COVID-19 period included only those patients who were able to come to the hospital. Therefore, vulnerable patients who were unable to access health care during the pandemic may have been excluded from the study. The main cause of false TI was physicians’ concerns about the BP measured in the clinic being inaccurate, which is consistent with the reports of previous studies. HT treatment should be designed as a framework for physicians to improve the outcome of HT care and reduce TI. However, appropriated TI should be more accurate if patients are encouraged to use SMBP at home, as the present study revealed that this is the second greatest cause of false TI. Furthermore, current clinical guidelines indicate that SMBP is effective in lowering BP.

The effect of the COVID-19 period on TI was not found to be significant after adjustment for patient and physician-related factors. On the other hand, the findings of our study were consistent with those of previous studies, which stated that TI could be influenced by the patient-associated factors, such as comorbidities, severity of HT, and age. Novel factors, such as senior medical staff and one type of anti-HT medication usage, were found to strongly affect TI. There have been no direct studies on the association between senior staff and TI; however, it is possible that senior medical staff are concerned about the impact of long-term doctor–patient relationships and acknowledge patient preferences, resulting in a slower rate of medication increase.
Table 4 Factors Associated with Therapeutic Inertia in Hypertension Treatment Using Final Multilevel Modeling

| Factor                                      | Adjusted OR (95% CI) |
|---------------------------------------------|----------------------|
| Intercept                                   | 1.05 (0.40–2.79)     |

1. Patient-related

| Factor                                      | Adjusted OR (95% CI) |
|---------------------------------------------|----------------------|
| COVID-19 vs pre-COVID-19 period             | 0.82 (0.67–1.01)     |
| Female vs male                              | 0.98 (0.81–1.18)     |
| Increase in 1 year of age                   | 1.01 (1.00–1.02)     |
| SBP >150 mmHg vs SBP 141–150 mmHg           | 0.34 (0.28–0.41)     |
| DM vs non-DM                                | 1.43 (1.19–1.72)     |
| Using 2 vs 1 type of HT medication          | 0.65 (0.54–0.78)     |
| Using second-line vs other anti-HT medication | 0.56 (0.31–1.03)   |
| Using ACE-I vs other anti-HT medication     | 1.01 (0.84–1.22)     |

2. Physician-related

| Factor                                      | Adjusted OR (95% CI) |
|---------------------------------------------|----------------------|
| Senior staff vs other physicians            | 1.87 (1.06–3.11)     |
| Female vs male                              | 0.96 (0.65–1.42)     |
| Internist vs other physician                | 0.83 (0.45–1.52)     |

Abbreviations: ACE-I, angiotensin-converting enzyme inhibitors; CI, confidence interval; DM, diabetes mellitus; HT, hypertension, OR, odds ratio; SBP, systemic blood pressure.

Strengths and Limitations

One limitation of this study is that it was a retrospective cross-sectional study in which data were reviewed from medical records and the reasons that physicians prescribed anti-HT were assumed from their notes. However, in clinical practice, physicians did not always record the reasons. Therefore, this retrospective study was limited by incomplete information in the medical records. A strength of the present study is that it is the first to identify a relationship between the COVID-19 pandemic and TI in HT care. However, COVID-19 outcomes are not directly relevant to TI. Nevertheless, we revealed intriguing effects of COVID-19 that contribute to HT care, such as an increased percentage of uncontrolled BP and other predictors for TI.

Implications for Policy, Practice, and Research

Our findings revealed that physicians were concerned about increasing anti-HT in patients with SBP >150 mmHg, even though the current HT guidelines recommend a lower optimal BP. This encourages physicians to follow current HT guidelines more rigorously. The guidelines demonstrate the clear benefits of combining anti-HT types for BP control, particularly in uncontrolled HT. However, in the present study, TI was higher with use of one type of anti-HT medication compared with use of two or more types. This demonstrates the reluctance of physicians to increase anti-HT use from one to two types. On the other hand, the prompt addition of a second type of anti-HT medication makes the next step simpler. Our findings showed a positive effect of senior staff on increasing TI. A previous study suggested that one of main problems was providers’ own clinical judgment and experience. However, there has been no prior research directly describing TI among senior staffs. We hypothesized that senior staff may have had a long relationship with their patients, due to which rapid medication adjustment would not have been performed but they might suggest the patient to lifestyle modifications.

To resolve this issue, we propose that the implementation and revision of a protocol for improved HT control in primary care in the HT clinic as part of routine evidence-based practice. Further prospective studies are required using TI or TI score to represent health-care system inertia in other NCDs, including HT. In addition, we propose that future studies should include an intervention to improve TI in NCD care. Finally, in our study setting, we demonstrated that resilience and adapted health care was strongly exhibited during the pandemic. This developing systematic approaches are in accordance with the digital healthcare solution for NCD services during the pandemic of WHO.

However, while out-of-patient prescriptions may have a negative impact on the quality of HT care, it may benefit coverage during a pandemic. Finding a balance between treatment quality and access to treatment should be considered concurrently for COVID-19 as well as the next pandemic. The health-care system should be well-prepared and efficient.

Conclusion

The present study revealed that the COVID-19 pandemic had no significant effect on TI in HT care. Senior staff were positive for TI among the physician-related factors. Furthermore, patient-related factors, such as elderly patients, prior DM diagnosis, lower SBP, and use of only one medication were associated with TI. However, the medical health care in this study was adapted to deal with the pandemic.
Key Messages

- The coronavirus disease-2019 (COVID-19) pandemic has affected many health-care systems, including hypertension care (HT). Therapeutic inertia (TI) is defined as failure to intensify therapy in the absence of an optimal goal. This study assessed the impact of the pandemic on TI and its predictors in patients with HT in a primary care clinic (PCC) in Thailand.
- The study found that the demographics of patients and physicians during the COVID-19 period were not significantly different from those in the preceding period. During the COVID-19 period, patients visiting the PCC had lower blood pressure and uncontrolled HT.
- TI was consistently more common in elderly patients and those with a prior diabetes mellitus diagnosis during both the periods. The adjusted odds ratio of TI increased by a factor of two among patients treated by senior staff. Patients who received more than one type of anti-HT therapy were less likely to develop TI than those who received only one type.
- However, after adjusting for factors using the multilevel model, no relationship was found between the COVID-19 period and TI.

Data Sharing Statement

The data presented in this study are available from the corresponding authors upon reasonable request.

Ethics and Other Permissions

The study was approved by the Human Research ethics committee (HREC), Faculty of Medicine, Prince of Songkla University issued on October 20, 2020. (Approval Number: REC 63-396-9-1). HREC is an official institute responsible for ethical review and supervision for human research. The study was conducted in accordance with the Declaration of Helsinki. All medical records were fully computerized by the Division of Digital Innovation and Data Analytics and confidentially reported.

Informed Consent Statement

The requirement for informed consent was waived because of the retrospective study design. This retrospective study was conducted using electronic medical records. The clinical data that could be used to identify patients was kept private, and the record form was assigned a code number. The patients’ code numbers were used during the analysis.

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Author Contributions

All authors made a significant contribution to the work reported, whether that is in the conception, study design, execution, acquisition of data, analysis and interpretation, or in all these areas; took part in drafting, revising or critically reviewing the article; gave final approval of the version to be published; have agreed on the journal to which the article has been submitted; and agree to be accountable for all aspects of the work.

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Disclosure

The authors report no conflicts of interest in this work.

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