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***Retrospective Cohort Study***

**Emergency department visits and hospital admissions in kidney transplant recipients during the COVID-19 pandemic: A hospital-based study**

Wongtanasarasin W *et al.* ED visit in KT during COVID-19

## **Abstract**

### **BACKGROUND**

Several studies have demonstrated that the coronavirus disease 2019 (COVID-19) affected daily living and the healthcare system. No previous evidence describes the consequences of COVID-19 on emergency department (ED) uses and hospital admissions among kidney transplantation (KT) recipients.

### **AIM**

To investigate the impact of the COVID-19 pandemic on the ED visits and hospital admissions within one year in patients who underwent KT in Thailand.

### **METHODS**

We conducted a retrospective study at a university hospital in Thailand. We reviewed hospital records of KT patients who visited ED during the outbreak of COVID-19 (from January 2020 to December 2021). We used the previous two years as the control period in the analysis. We obtained baseline demographics and ED visit characteristics of each KT patient. The outcomes of interest were ED visits and ED visits leading to hospital admission within the first year following a KT. The rate of ED visits and ED visits leading to hospital admissions between the two periods were compared using Stratified Cox's proportional hazard model.

### **RESULTS**

A total of 263 patients were included, 112 during the COVID-19 period and 151 during the control period. There were 34 and 41 ED visits after KT in the COVID-19 and control periods, respectively. The rate of first ED visit at one year was not significantly different in the COVID-19 period, compared with the control period [hazard ratio (HR) = 1.02, 95% confidence interval (CI): 0.54-1.92,  $P = 0.96$ ]. The hospital admission rate was similar between periods (HR = 0.92, 95% CI: 0.50-1.69,  $P = 0.78$ ).

## CONCLUSION

ED visits and hospital admissions within the first following year in KT recipients were not affected by the COVID-19 pandemic. Despite these findings, we believe that communication between post-KT patients and healthcare providers is essential to underline the importance of prompt ED visits for acute health conditions, particularly in post-KT patients.

**Key Words:** Emergency department visit; Hospital admission; Kidney transplantation; COVID-19

Wongtanasarasin W, Phinyo P. Emergency department visits and hospital admissions in kidney transplant recipients during the COVID-19 pandemic: A hospital-based study. *World J Transplant* 2022; In press

**Core Tip:** Coronavirus disease 2019 (COVID-19) affects kidney transplantation (KT) recipients in terms of hospital admission rates. This study highlights that despite emergency department (ED) visits not being changed during the COVID-19 spreading; hospital admission rates were increased. Although we could not establish the cause-effect relationship of these changes, we encourage healthcare providers to provide post-KT patients recommendations to visit ED promptly for acute health conditions.

## INTRODUCTION

In the United States, there were approximately 143 million total visits to an emergency department (ED) in 2018<sup>[1]</sup>. Over the last two decades, the rate of ED visits has increased, exceeding what could be accounted for by population growth<sup>[1]</sup>. Multiple factors, including extremes of age, women, public insurance, minority race/ethnicity, and country region, are associated with higher rates of ED visits in the general population<sup>[2]</sup>. Recently, there has been a significant increase in acute care delivery following hospitalization<sup>[3,4]</sup>. Acute care after hospital treatment is considered an

indication of poor quality of care in some contexts, including kidney transplantation (KT) patients<sup>[4,5]</sup>. Patients with end-stage renal disease (ESRD) account for 7.1% of total Medicare expenditures in the United States despite accounting for only 0.9% of Medicare treatments<sup>[6,7]</sup>. Patients with ESRD have visited the ED at a 6-fold higher rate than the general population; however, most previous studies excluded KT patients, who account for a growing proportion (around 22.8%) of prevalent ESRD patients<sup>[7]</sup>. The long-term advantages of KT are well documented and include improved survival and quality of life compared to dialysis<sup>[8,9]</sup>. On the other hand, the management of patients after KT is complex and resource-intensive, necessitating extensive care coordination, frequent laboratory monitoring, and ongoing patient engagement<sup>[9,10]</sup>. Furthermore, KT recipients frequently have multiple comorbidities, which complicates their care<sup>[11,12]</sup>.

In recent years, coronavirus disease 2019 (COVID-19) has become the most critical disease and influenced human health across the globe<sup>[13]</sup>. This pandemic affects not only physical health but also mental health and well-being<sup>[14]</sup>. Transplant recipients, including KT patients, who are receiving immunosuppressive therapy are at the highest risk of severe illness and, as a result, are at a higher risk of an adverse outcome from COVID-19<sup>[15]</sup>. One of the unique aspects of the transplant recipient's life is that, in the post-operative phase, the patient should live in an isolated space, pay special attention to their living environment, and prefer a limited social life because of the immunosuppressive treatment involves immunosuppression in the patient<sup>[14]</sup>. A previous study demonstrated that the COVID-19 pandemic is associated with a significant reduction in average daily ED visits; however, the admission rates were increased<sup>[16]</sup>. This research aimed to investigate the effect of COVID-19 and the consequences on the ED visits and admission rates among KT recipients within one year. In addition, this study aimed to assess the differences in KT patients' diagnoses who visited an ED between COVID-19 and regular periods.

## **MATERIALS AND METHODS**

### ***Protocol***

We conducted a single-center retrospective observational study at a university tertiary hospital between January 2018 and December 2021. The study protocol was approved by the Institutional Review Board (IRB) of the Faculty of Medicine, Chiang Mai University (EXEMPTION-8745/65). The IRB waived informed consent due to its retrospective design. Patient confidentiality was preserved by using anonymous health records. All methods employed in this study were performed following relevant guidelines and regulations.

### ***Setting and study population***

Maharaj Nakorn Chiang Mai Hospital (MNCMH) is a university hospital with 1500 beds, 151 intensive care units (ICU) and sub-ICU beds, 28 operating rooms, and doctors from all subspecialties on duty. According to the Canadian Triage and Acuity Scale, the triage categorization is based on a five-level scale, ranging from blue (level 1, resuscitation) to white (level 5, non-urgency). Our ED provides a 24-h service with emergency physicians and skilled nurses. We categorized seven types of dispositions in the current study: ICU admission, general ward admission, observational unit admission, referral to another hospital, discharge, discharge against doctor's recommendation, and death.

We included all adult patients (age  $\geq 18$  years) who underwent KT at MNCMH between January 2017 and December 2020. Patients who died in the hospital after KT before hospital discharge were excluded. We collected data only from KT patients who visited the ED of MNCMH within one year after the date of transplantation (between January 2018 and December 2021). Extreme outliers and high-volume ED visitors (KT patients using the ED more than ten times per year) were excluded from the study population and were not included in the study analysis.

### ***Data collection***

Data were collected through the Electronic Medical Records and chart review. To assess risk factors for ED visits and admissions following KT, age, gender, donor types, insurance, and Charlson comorbidity index were gathered. Specifically, for KT recipients who visited ED within a year after transplantation, we collected the data on: (1) Time to first and any ED visit since transplantation; (2) Triage level; (3) Total ED time; (4) Types of disposition; and (5) Invasive procedures during ED stay which were intubation and cardiopulmonary resuscitation. Diagnosis for each ED visit is also collected using the International Classification of Diseases code.

### ***Outcomes and data analysis***

The primary outcome of interest was ED visits in the first year following KT. All recipients were followed until death or out of the study period. In-hospital deaths were retrieved from hospital medical records. Patients who did not visit ED at the end of the study period were considered censors. For patients with recurrent ED visits, the time to ED visit was defined as the time from the index date of transplantation to the date of the recurrent ED visit. The risk interval was, therefore, set as marginal since we assumed that the patients were at risk of any ED visit from the date of their transplantation.

Secondary outcomes included ED visits leading to hospital admissions following KT's first year. The number of ED visits and hospital admissions for any reason was calculated and compared between January 2018-December 2019 and January 2020-December 2021. All responsible diagnoses from January 2018-December 2019 were compared to all diagnoses from January 2020-December 2021. We described continuous data using mean  $\pm$  SD for normally distributed. For skewed data, median and interquartile range were calculated. Categorical data were summarized using frequency and percentage. The independent *t*-test was used to compare continuous variables. For categorical variables, Fisher's exact probability test was performed. All tests were two-sided, with significance for all tests being determined as a *P*-value  $< 0.05$ . All analyses were performed using STATA 16 (StataCorp, College Station, TX, United States).

For the primary analysis, the rate of ED visits within a year after KT was compared using the stratified Cox proportional hazards model. We presented two analytic approaches for each survival outcome, the rate of first ED visits and any ED visits after transplantation. For the rate of the first ED visit, we restricted the analysis to only the first ED visit, whereas all ED visits during the first year period were considered in the analysis of the rate of any ED visits. We employed the modeling method for recurrent events described by Kelly and Lim<sup>[17]</sup>. The risk interval was defined as the total time (marginal). We used a restricted risk set and assumed event-specific baseline hazards. To quantify the effect of the COVID-19 pandemic period on the control period, hazard ratios (HRs) were estimated from the stratified Cox's regression model. They were reported with 95% confidence intervals (CI) and *P*-values. Kaplan-Meier curves were demonstrated, comparison of differences was made by log-rank test.

## **RESULTS**

### ***Patient characteristics***

A total of 263 KT recipients were enrolled in this study, 112 in the COVID-19 period (underwent KT between January 2019 and December 2020) and 151 in the control period (underwent KT between January 2017 and December 2018). No recipient died during the follow-up period. The mean ages were  $45.5 \pm 10.4$  and  $43.7 \pm 13.4$  years for COVID-19 and control groups, respectively. Most of the participants received deceased donors. There were no significant differences in baseline demographics between the two periods (Table 1). Baseline demographics of KT patients who visited an ED during the study periods were summarized in Table 1.

### ***ED visits***

A total of 17.1% of KT recipients visited ED within one year after transplantation (15.3% in the COVID-19 period and 18.5% in the control period), accounting for 75 ED visits. The mean times to first ED visit since transplantations were  $130.8 \pm 106.2$  and  $120.6 \pm 105.3$  d for COVID-19 and control periods, respectively. On the other hand, the rates of



invasive procedures were similar among both periods. Table 2 summarizes the clinical variables of KT patients who presented to the ED within one year after transplantation. The rate of first ED visit at one year was not different in the COVID-19 period, compared with the control period when adjusting for confounding variables (HR = 1.02, 95%CI: 0.54-1.92,  $P = 0.96$ , Figure 2). Similarly, the rate of any ED visit in the following year was also not different between the two periods (HR = 1.24, 95%CI: 0.73-2.10,  $P = 0.43$ , Table 3). The five most responsible diagnoses are demonstrated in Table 4. Fever and abdominal pain were ranked first during the control period, while abdominal pain was the top diagnosis during COVID-19.

### *Hospital admissions*

The admission rate in the COVID-19 period decreased significantly during the study period, compared with the control period (38.2% vs 65.9%,  $P = 0.02$ ). In addition, the rate of any ED visit leading to hospital admission in the following year was also not different (HR = 0.92, 95%CI: 0.50-1.69,  $P = 0.78$ , Table 3).

## **DISCUSSION**

In this retrospective study of KT patients, around one-sixth of KT recipients had at least 1 ED visit in the first year following transplantation. However, the rates of ED visits and hospital admissions were not affected by the impact of the COVID-19 pandemic. We also found that abdominal pain was responsible for most diagnoses across COVID-19 and control periods. The impact of COVID-19 on ED visits and hospital admissions is demonstrated across several previous studies<sup>[15-17]</sup>. To our knowledge, this is the first study investigating the effect of the COVID-19 pandemic on ED visits and admission rates among KT patients. KT recipients are usually advised to isolate themselves from the community because of the greater risk of being infected. Consequently, they might not visit ED promptly. Our earlier study found that an average daily ED visit was significantly reduced during the COVID-19 pandemic, probably due to the fear of reaching COVID-19 in the hospital<sup>[15]</sup>. However, these present findings showed the

difference. Despite the fear of contacting COVID-19, we found that ED visits by post-KT patients were not disturbed. The previous study has demonstrated that KT recipients had a higher chance of a more severe course of COVID-19 infection than hemodialysis patients<sup>[18]</sup>; however, another finding has shown that the severity and adverse outcomes were not different between KT recipients and those without for the COVID-19 infection<sup>[19]</sup>.

Recently, telemedicine has become one of the most powerful strategy using to follow-up kidney transplant recipients<sup>[18,19]</sup>. Results from Yadav and Singh's study found that application of telemedicine in the transplant population enhances medication compliance, reduces hospitalization rates, and makes living donor evaluation convenient<sup>[19]</sup>. Telemedicine could be recommended as an alternative method, especially in the pandemic era, to avoid and reduce the rate of transmission in the hospital in kidney transplant population.

Although ED visits are not different between the two groups in our study, hospital admissions were higher for the COVID-19 group. This may reflect the natural consequence of inappropriate and untimely ED visits, resulting in a higher severity of diseases. We proposed that the reasons for these findings could be multifactorial. First, KT patients have a higher baseline chance of visiting ED than other patients. Previous studies have shown that acute care utilization in the following year after KT is relatively high<sup>[4,7,9]</sup>. In one retrospective study conducted in the United States, nearly half of KT patients visited the ED within one year after KT<sup>[7]</sup>. Second, post-KT recipients are prescribed immunosuppressive agents. Usually, they are informed to seek medical evaluation even they have minor symptoms, such as low-grade fever or abdominal pain. Furthermore, fever and other unspecified symptoms could be one of the clinical features of COVID-19<sup>[20]</sup>. KT recipients might intend to visit ED as they considered themselves suspected of having this COVID-19 infection. Interestingly, our study found that hospital admissions were markedly increased in the COVID-19 group. Consistent with the previous evidence, hospital admission during this disastrous period is likely higher than usual, mainly because of untimely and delayed ED visits<sup>[15]</sup>.

Our findings regarding ED visits and admission rates during the COVID-19 pandemic may serve as a body of literature regarding the impact of COVID-19 in the various spectrum, including KT recipients. Not only the number of ED visits among post-KT patients were not less than the regular period, but also the admission rates were significantly high. Our data also suggests clinicians and healthcare professionals to encourage KT recipients to visit ED on time to reduce unfavorable outcomes.

### ***Limitations***

This study has some limitations to be considered. This method could not account for underlying trends in hospital admission and ED attendance despite comparing two time periods. Differences in hospital admission patterns may be associated with the epidemic or the limits by chance. This problem might be solved with additional time series analysis or regression modeling over a longer time. We only did the investigation at a single university hospital. As a result, the design may be valid and generalizable to the situation with the same degree of care. Furthermore, some baseline data were not recorded, including causes of ESRD and hospital length of stay during index transplantation. Moreover, another perspective that this study did not address was the quality of life of post-KT patients who visited ED in the first following year. Further research should evaluate this aspect of the patients.

### **CONCLUSION**

In conclusion, COVID-19 also affects KT recipients in terms of hospital admission rates. The present study points out that despite ED visits not being changed during the COVID-19 pandemic, hospital admission rates were increased. Although we could not determine the exact cause of this change, we believe that communication between post-KT patients and healthcare providers <sup>3</sup> is necessary to emphasize the importance of timely ED visits for acute health conditions, especially in immunocompromised hosts like post-KT patients.

## **ARTICLE HIGHLIGHTS**

### ***Research background***

Several investigations have shown that the coronavirus disease 2019 (COVID-19) has an impact on daily life and the healthcare system.

### ***Research motivation***

There has been no previous research on the effects of COVID-19 on emergency department (ED) visits and hospitalizations among kidney transplantation (KT) patients. We conducted this study to explore the effects of COVID-19 on ED visits among post-KT recipients.

### ***Research objectives***

The aim of this study was to investigate the impact of the COVID-19 pandemic on the ED visits and hospital admissions within one year in patients who underwent KT in Thailand.

### ***Research methods***

We conducted a retrospective study. We reviewed hospital records of KT patients who visited ED during the outbreak of COVID-19. We used the previous two years as the control period in the analysis. We obtained baseline demographics and ED visit characteristics of each KT patient. The outcomes of interest were ED visits and ED visits leading to hospital admission within the first year following a KT.

### ***Research results***

We included a total of 263 patients, 112 during the COVID-19 period and 151 during the control period. There were 34 and 41 ED visits after KT in the COVID-19 and control periods, respectively. The rate of first ED visit at one year was not significantly different in the COVID-19 period, compared with the control period. The hospital admission rate was also similar between periods.

### ***Research conclusions***

The COVID-19 pandemic had no effect on KT recipients' ED visits or hospital admissions in the first year after transplantations.

### ***Research perspectives***

Despite these findings, we suggest that communication between post-KT patients and healthcare professionals is crucial in emphasizing the significance of timely ED visits for acute health issues, especially in post-KT patients.

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