### Timing of percutaneous endoscopic gastrostomy in COVID-19 infection: Endoscopic surgery unit experience

## COVID-19 enfeksiyonunda perkütan endoskopik gastrostomi zamanlaması: Endoskopik cerrahi ünite deneyimi

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#### Abstract

Aim: The COVID-19 pandemic has negatively affected the whole world and health systems. Although the D literature includes recommendations regarding the timing of Percutaneous endoscopic gastrostomy (PEG) in COVID-19 patients, there are no significant clinical studies yet. Therefore, we aim to contribute to the literature by sharing our data on this subject.

Materials and Methods: Patients who underwent PEG between March 2020 and March 2021 were retrospectively evaluated in our clinic. The patients were compared statically in terms of age, gender, medical indications, comorbid diseases, hospitalization in the intensive care unit (ICU), blood tests, and post-intervention complications. PEG was inserted routinely in PCR-negative patients. Patients who underwent PEG were compared as outpatients and inpatients in the ICU. Moreover, patients who underwent PEG while hospitalized in the ICU were divided into two groups according to the presence of COVID-19 infection; patients noninfected with COVID-19 (group 1) and COVID-19 infected patients (group 2).

Results: PEG was performed in 66 patients during the COVID-19 pandemic. These patients predominantly consisted of those with SVH, Alzheimer's, or traumatic brain injury. In the present study, thirty-two (%48.5) patients were female with a mean age of  $69.4\pm17.6$ , and forty-seven patients underwent PEG in the ICU. Furthermore, eleven of these patients were COVID-19, infected patient group (group-2). There was no statistical difference in blood albumin levels, CRP, hemogram results, and 30-day mortality results between group 1 and group 2 (P>0.05).

Conclusion: PEG is a minimally invasive intervention that is commonly used for enteral feeding. The timing of the procedure is crucial for inpatients with COVID-19. Although the most appropriate timing is the 30th day after the COVID-19 infection process, we think that PEG may insert on the 10th day in eligible patients.

Keywords: Percutaneous Endoscopic Gastrostomy, COVID-19, infection, Cerebrovascular disease, minimally invasive

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Ethics Committee Approval: This study was approved by the Sakarya University Medical Faculty Ethics Committee (ID: 71522473-050.01.04-21455-212, date: 05.03.2021).

Etik Kurul Onayı: Bu çalışma Sakarya Üniversitesi Tıp Fakültesi Etik Kurulu tarafından onaylanmıştır (ID: 71522473-050.01.04-21455-212, date: 05.03.2021).

Conflict of Interest: No conflict of interest was declared by the authors.

Çıkar Çatışması: Yazar çıkar çatışması bildirmemiştir.

Financial Disclosure: The authors declared that this case has received no financial support.

Finansal Destek: Yazarlar bu çalışma için finansal destek almadıklarını beyan etmişlerdir.

Geliş Tarihi / Received: 05.10.2021 Kabul Tarihi / Accepted: 02.12.2021 Yayın Tarihi / Published: 09.12.2021

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Amaç: COVID-19 pandemisi tüm dünyayı ve sağlık sistemlerini olumsuz şekilde etkisi altına almıştır. Literatürde COVID-19 hastalarında Perkütan endoskopik gastrostomi (PEG) zamanlaması ile ilgili öneriler yer alsa da henüz önemli bir klinik çalışma bulunmamaktadır. Bu konudaki verilerimizi paylaşarak literatüre katkı sağlamayı amaçlıyoruz.

Gereç ve Yöntem: Mart 2020-Mart 2021 tarihleri arasında PEG takılan hastalar geriye dönük olarak değerlendirildi. Hastalar yaş, cinsiyet, tibbi endikasyonlar, eşlik eden hastalıkları, kan testleri ve müdahale sonrası komplikasyonlar ile işlem sırasında hastanın yoğun bakım ünitesinde olup olmadığı kaydedildi. PCR negatif hastalarda rutin olarak PEG yapıldı. PEG uygulanan hastalar ayaktan ve yoğun bakım ünitesinde yatan hastalar olarak karşılaştırıldı. Daha sonra yoğun bakım ünitesinde yatarken PEG uygulanan hastalar, COVID-19 enfeksiyonu varlığına göre iki gruba ayrıldı. (grup-1 ve grup-2)

Bulgular: Pandemi sırasında uygun endikasyonları olan 66 hastaya PEG yerleştirildi. Bu hastalar ağırlıklı olarak SVH, Alzheimer veya travmatik beyin hasarı olanlardan oluşuyordu. Çalışmada otuziki (%48.5) hasta kadındı ve yaş ortalaması 69.4±17.6 idi. Ayrıca, kırkyedi hastaya yoğun bakımdayken PEG uygulandı; bu hastalardan 11'i COVID-19 enfekte hasta grubuydu (grup-2). Grup-1 ve grup-2 arasında kan albümin düzeyleri, CRP, hemogram sonuçları ve 30 günlük mortalite sonuçları açısından istatistiksel fark yoktu (P>0.05)

Sonuç: Minimal invaziv bir işlem olan peg enteral beslenme amacıyla yaygın olarak kullanılmaktadır. COVID-19 enfekte hastalara işlemin zamanlaması önemlidir. En uygun zamanlama enfeksiyon sürecinden sonraki 30. gün olmakla birlikte uygun hastalarda 10. Gün peg uygulanabilceğini düşünmekteyiz.

Anahtar Kelimeler: Perkütan Endoskopik Gastrostomi, COVID-19, enfeksiyon, Serebrovasküler hastalık, minimal invaziv

Atıf yazım şekli: How to cite:

Öz

m şekli: Fırat N, Aziret M, Mantoğlu B, Çapoğlu R, Demir H, Dikicier E, Akın E, Harmatepe T, Altıntoprak F. Timing of percutaneous endoscopic gastrostomy in Covid-19 infection: Endoscopic surgery unit experience. Arch Clin Exp Med 2021;6(3):158-161.

### Introduction

Coronavirus, which has spread rapidly worldwide, is a respiratory RNA virus that can lead to clinical issues with severe respiratory failure [1]. Symptomatic patients often transmit the infection through sizeable droplets produced during coughing and sneezing [2]. As a result of this global enigma, new guidelines to protect patients and healthcare professionals from confirmed and suspected cases are frequently issued. At the same time, health authorities strive to establish the optimal approach [3]. Recommendations for invasive surgical procedures in patients infected with COVID-19 have also been reviewed, and a delay to surgery of 3-4 weeks is recommended in the case of patients who cannot be fed orally and so require feeding tube insertion [4].

In 1980, Gauderer et al. described the percutaneous endoscopic gastrostomy (PEG) primarily for elderly patients with swallowing difficulties linked to neurological disorders [5,6]. This minimally invasive method is more advantageous than other procedures, providing a secure long-term feeding route with low morbidity and complication rates [7,8]. Therefore, its efficiency and practicality make PEG a preferred intervention for long-term enteral nutrition [9].

Currently, the literature includes minimal data on the topic of placing PEG in COVID-19 patients. Therefore, this study aims to evaluate PEG applications in our center over one year during the pandemic and present our data on PEG timing in patients infected with COVID-19.

### **Material and methods**

### **Patients and Ethics**

This study includes patients who underwent PEG in the General Surgery Endoscopy unit between March 2020 and March 2021, incorporating the first and second pandemic waves. The PEG procedure was performed on patients with gastrointestinal system continuity but could not take oral nutrition for about 30 days. Data concerning patient age, gender, medical indications, comorbid diseases, hospitalization in intensive care unit (ICU) at the time of the procedure, laboratory parameters, PCR tests, and complications were accessed from patient records and documented. First, the patients were compared as inpatients and outpatients in the ICU. Second, the patients were divided into two groups: patients in the intensive care unit not infected with COVID-19 (group 1) and patients infected with COVID-19 (Group 2). The groups were compared in terms of clinical features. Approval for this study was granted by the ethics committee of Sakarya University (05/03/2021; 71522473-050.01.04-21455-212).

## Patient selection process, diagnosis of COVID-19, and prevention of infection

PEG was administered to patients who were PCR negative. The procedure was not performed on patients with impaired bleeding profiles or symptoms of sepsis. COVID-19 diagnosis was made in symptomatic patients through PCR positivity and thorax computed tomography (CT) findings. All patients were given a COVID-19 PCR test before the procedure. In both PCR-positive patients and cases diagnosed with COVID-19 by tomography, PEG was delayed until after a negative PCR could be obtained, and symptoms of the disease had passed. While standard precautions were taken for non-virus-infected patients (wearing of masks by all team members and 30-minute room ventilation), special precautions were taken for infected patients (N95 mask, visor, disposable gowns, 1-hour room ventilation, and disinfection of the room after the procedure).

### **PEG procedure**

Before the procedure, patients were monitored, and nasal oxygen was initiated at 4-6 L/minute. The patients received 1-2 mg midazolam, and one mcg/kg fentanyl, with 0.5 mg/kg propofol added after the procedure had started for sedation. The cases got a lidocaine hydrochloride spray and 2cc prilocaine to the skin at the incision site for local oropharyngeal anesthesia. Antibiotic prophylaxis with cefazolin sodium was administered to the patients, except for cases already under antibiotic treatment. The patients did not receive enteral feeding for at least 8 hours before the procedure. An experienced team performed the process in the general surgery endoscopy unit.

The PEG procedure was performed by applying the "pull-through" technique by observing the indentation made by the finger and the transillumination created by the endoscope through the stomach. As a standard, a 20Fr PEG tube was placed in all patients. All interventions were made using a Fujinon VP-4450HD (Fujifilm company, Minato-ku, Tokyo, Japan) fiber endoscope. Enteral feeding was started 16 hours after the feeding tube was inserted.

In the present study, one patient whose PCR test was positive during the peg procedure and three patients who could not have the PEG procedure were excluded. In addition, while endoscopic gastric tube revisions were included in the study, gastric tube replacements performed without endoscopy were excluded.

### Statistical analysis

The frequency of sociodemographic, clinical data, and descriptive statistics were calculated as numbers, distribution, and percentages. The control groups, continuous variables with normal distribution were compared using the independent sample t-test; variables not showing a normal distribution with the Mann-Whitney U test, and independent group rates were compared using Fisher's exact test. A p-value <0.05 was considered significant. Analyses were performed using SPSS statistical software (IBM SPSS Statistics, Version 26.0. Armonk, NY: IBM Corp.).

### Results

During the pandemic, 66 patients had appropriate indications for PEG, and these procedures were planned. However, PEG was not performed because of co-morbidities, high-risk American Society of Anesthesiologists (ASA) scores, unsuitable anatomy or infection status. Patients undergoing PEG frequently had cerebrovascular disorder (CVD), Alzheimer's disease, or traumatic brain injury. The primary and comorbid conditions of these patients were shown in Table 1.

Most patients were in the ICU at the time of the procedure. A comparison was made between the intensive care patients and those who were prepared for PEG in the outpatients' clinic. (Table 2). The results of this comparison showed the 30-day mortality rate to be significantly higher in intensive care patients (p = 0.01). In addition, the C reactive protein (CRP) and white blood count (WBC) values of the patients hospitalized in the intensive care unit were significantly higher (p = 0.001 and p = 0.01, respectively), while albumin and hematocrit levels were significantly lower (p < 0.05). All of the virus-infected patients were inpatients in the intensive care unit (Table 2).

In the present study, ten patients had complications with their feeding tubes due to the deterioration of the tube in five patients and wound infection in the remaining five patients. In comparison, the antibiotic treatment was sufficient in the five patients who had wound infection; in the remaining cases, the tube had to be removed and replaced with a new feeding tube. The diagnosis of virus-infected patients (group 2), who were all in intensive care, was made by PCR in 4 patients and tomography in 7 patients. Patients with ICU have been divided into two groups: non-virus-infected (group 1) and virus-infected (group1) (Table 3). There was no significant difference between the two groups regarding age, gender, blood albumin levels, WBC and hematocrit values, blood gas, or lactate values (Table 3).

 Table1. Primary diseases and comorbid diseases of patients with PEG

 All Deticate (a)

 Patients with

	All Patients (n%)	COVID-19 (n%)	
Diagnosis			
Cerebrovascular disease	25 (37.9)	5 (45.4)	
Alzheimer disease	13 (19.7)	3 (27.3)	
Trauma	7 (10.6)		
General condition disorder	4 (6.1)	1.1 (9.1)	
Hypoxic or anoxic brain injury	5 (7.6)	1.1 (9.1)	
Cranial tumor	3 (4.6)	1.1 (9.1)	
Dementia	2 (3.0)		
Parkinson disease	2 3.0)		
Amyotrophic lateral sclerosis	1 (1.5)		
Cerebral palsy	1 (1.5)		
Polyneuropathy	1 (1.5)		
Lip or tongue cancer	1 (1.5)		
Esophageal cancer	1 (1.5)		
Total	66 (100)	11(100)	
Co-Morbidities			
Hypertension	20 (29.9)	4 (33.3)	
Senility	17 (25.4)	6 (50)	
Lung failure	9 (13.4)	5 (41.7)	
Coronary artery disease	8 (11.9)	1 (8.3)	
Diabetes Mellitus	8 (11.9)	2 (16.7)	
Chronic obstructive lung disease	7 (10.4)	1 (8.3)	
Malignity	6 (9.0)		
Congestive heart failure	5 (7.5)	2 (16.7)	
Thrombocytopenia	2 (3.0)		
Schizophrenia	1 (1.5)		
Substance abuse	1 (1.5)		
None	23 (34.3)		
Total	106	21	

### Discussion

The COVID-19 infection is associated with a high mortality rate in patients with multiple co-morbidities. Patients undergoing PEG procedures have had generally long-term and severe co-morbidities. Therefore, the timing of PEG insertion is essential in the presence of the co-existence of the COVID and PEG process.

The ICU-specific nutrition guidelines have suggested early nutrition (EN) within 24-48 hours following the [10]. intervention Furthermore, the influence of hypoalbuminemia, serum C-reactive protein levels, metastatic cancer, and other comorbid diseases on mortality after PEG is well-documented [11]. Both albumin levels and hemogram values of the patients undergoing PEG in the ICU were significantly lower than in the other patients; their CRP levels were significantly higher; and the 30-day mortality was also higher in these ICU patients, as expected. Thus, the importance of early enteral nutrition in this patient group is evident.

Wound infection is the most common complication associated with the PEG procedure. The surgical site infection can frequently respond to antibiotic treatment. [12]. In half of our patients, the feeding tube had to be removed due to wound infection, but after medical treatment, the PEG was reinserted in a second session.

A positive PCR test is used to make a COVID-19 diagnosis in clinically suspected patients. In case of a false negative, thorax CT may assist the diagnosis [13]. In this study, tomography was used to confirm the diagnosis of COVID-19 in 7 patients.

**Table 2.** Descriptive statistics of Outpatient and ICU patients

	Insertion of PEG	Insertion of PEG	
	in outpatient	in ICU	Р
	(n=19)	(n=47)	
Gender (M/F)	12(63.2%)/7(36.8)	21(44.7)/26(55.3)	0.155
Age (year)	68.5±22.8	69.7±15.4	0.616
CRP (mg/L)	45.8±45.7	98.7±61.0	< 0.01
Albumin (g/L)	32.5±6.9	26.4±4.7	< 0.01
WBC (K/uL)	7.2±2.6	9.2±2.9	0.01
Hematocrit (%)	35.1±4.2	29.4±5.4	< 0.01
Hemoglobin (g/dl)	$11.4 \pm 1.4$	9.6±1.7	< 0.01
Complication and mortality			
Diagnosis of COVID-19	0	11(23.4%)	0,014*
Leakage around the PEG	2 (10.5%)	7(14.9%)	0,901*
Alteration of the PEG	6 (31.6%)	4(8.5%)	0.017
First 30-day mortality (%)	0	12(25.5%)	0.01

CRP: C-Reactive protein: Intensive care unit, PEG: Percutaneous endoscopic gastrostomy, WBC: White blood cell, Mann Whitney U test, \*,Fisher Exact

Table 3. Descriptive statistics of	Group 1 ar	nd Group 2
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	Patients without COVID-19	Patients with COVID-19	Р
	(Group 1)	(Group 2)	
	(n=36)	(n=11)	
Patients (female/male)	14/22 (38.9/61.1)	6/5 (54.5/45.5)	0.369
Age (year)	68.3±14.9	72.6±16.9	0.421
CRP (mg/L)	99.8±61.8	102.8±58.8	0.885
Albumin (g/L)	26.5±4.6	25.0±4.1	0.511
WBC (K/uL)	9.2±3.1	9.5±2.5	0.757
Hematocrit (%)	29.6±5.3	27.8±4.9	0.199
Hemoglobin (g/dl)	9.7±1.7	9.0±1.5	0.150
Arterial blood gases ph	7.4±0.1	7.4±0.06	0.680
Complication and the			
hospital mortality			
Leakage around the PEG	7 (21.2%)	0	
The insertion time of PEG	-	30 (10/50) *	-
after COVID-19 diagnosis			
(days)			
Hospital mortality (First-	9 (25.0%)	3 (27.3%)	0.295
30 days) (%)	· ·		
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CRP: C-Reactive protein, PEG: Percutaneous endoscopic gastrostomy, WBC: White blood cell, Mann Whitney U test, \* median(min/max)

In our study, there was a strong association between virus-infected patients who underwent PEG and COVID-19. The literature suggests that COVID-19 may be more severe in individuals with a history of COVID-19 and describes the effects of accompanying cardiovascular diseases in these patients [14]. Morbidity and mortality rates are higher in COVID-19 infected patients with comorbidities such as diabetes, cardiovascular disease, chronic respiratory diseases, and in patients over the age of 60 [15]. Patients who undergo PEG have multiple risk factors. After the early insertion of PEG, an increase in 30-day mortality can be expected. In the cases in this study, the 30-day mortality rate was similar to other patients hospitalized in the ICU.

Patients can be infected by the environment retching, coughing, and drooling during upper GI endoscopy. Therefore, measures should be taken to protect both patients and healthcare professionals in the pandemic period [16, 17]. It should be emphasized that at the peak of the pandemic, only urgent endoscopic procedures should be performed and that the decision to place endoscopic feeding tubes should be made on a patientby-patient basis [18]. Moreover, PEG intervention should not be achieved unless complications related to the nasoenteral tube develop in active infection [19]. As perioperative COVID-19 conditions are known to carry an increased risk of death and pulmonary complications, international collaborative studies recommend delaying non-urgent procedures, where possible [20]. Meanwhile, current studies also suggest the optimal time for surgical intervention to be after an asymptomatic period of 7 weeks following virus infection [21]. Also, for PEG, a delay of 3-4 weeks with the pull technique is recommended [4].

This study has some limitations. First, there was a small sample size and retrospective planned research. On the other

hand, the present study can be critical for emphasizing the timing of the PEG procedure in patients with PEG.

PEG, which is a minimally invasive procedure, is widely utilized to supply enteral nutrition. Since surgical procedures increase the severity of the disease in COVID-19 patients, the timing of the intervention is essential in infected patients. Although the most appropriate timing is on the 30th day after the infection has cleared, it is possible to use PEG on the 10th day in proper patients who have a negative PCR and no disease symptoms..

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