Pattern of Stroke and Short Term Outcome of COVID-19 Patients admitted in a Dedicated Stroke Unit at Referral Neurology Hospital of Bangladesh

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Abstract
Background: COVID-19 was declared as a pandemic by WHO on March 11, 2020. Though the most common and important presentation is with respiratory disease, reports of neurological features are increasing. Objective: In this study it had been described the pattern of stroke in COVID-19 patients admitted in a dedicated stroke unit and their short term outcome. Methodology: This single arm, single centre prospective cohort study was conducted in the Stroke Unit of National Institute of Neurosciences and Hospital, Dhaka, Bangladesh from April to August, 2020 and was inclusive acute stroke patients irrespective of age and sex who were found RT-PCR positive for SARS CoV-2. The patients were evaluated clinically, biochemically and by imaging. After diagnosis patients were referred to a COVID dedicated hospital and were followed up over phone for 4 weeks from the day of admission in stroke unit to assess short term outcome. Results: Among the 41 patients, 21 were male and 20 female with an age range of 26 to 90 years where 58.5% patients were aged 60 and above. Majority (58.5%) of them came from different urban areas. Common co-morbidities were Hypertension (68.3%) and Diabetes mellitus (39%). Most common presenting features were hemiplegia (48.8%), and altered level of consciousness (39%). More than seventy percent (70.7%) patients had GCS of 10 or less and 56.1% patients had oxygen saturation 90% or less in room air. CT brain imaging revealed infarct in 39% and hemorrhage in 51.2%. Chest radiography revealed abnormality in 14.6% cases. Regarding the short term outcome, total 20 (48.8%) patients died where 7 died before referral to COVID-19 dedicated hospital. Death was higher in hemorrhagic stroke but there was no significant association between outcome and the type of stroke (p value>0.05). Conclusions: Hemorrhagic stroke is the most commonly occurring stroke among the COVID-19 patients and associated with poor outcome. Keywords: SARS CoV-2; COVID-19; stroke; hemorrhagic stroke

Introduction
Corona virus disease 2019 (COVID-19) caused by SARS-CoV-2 virus, first emerged from Wuhan city, Hubei province of China. The epidemic spread rapidly worldwide within 3 months and was declared as a pandemic by WHO on March 11, 2020. Severe acute respiratory illness with fever and respiratory symptoms, such as cough and shortness of breath, comprise the main clinical presentations. But unusual manifestations, such as patients without respiratory symptoms or only very mild symptoms are rising worldwide. Initially it was thought to involve primarily the respiratory system like other coronaviruses like severe acute respiratory syndrome (SARS) and the Middle East Respiratory Syndrome (MERS) later it was found to have involved the lungs in 17.1% cases in this study. The sensitivity of manifestation of COVID-19 patients. Chest diseases/novel-coronavirus-2019/events-as-they-happen (accessed 13 April 2020).
 Syndrome (MERS), later it was found to have involved other systems as well including the nervous system. Although the most common and important presentation is with respiratory disease, reports of neurological features are increasing. These features appear to be a combination of non-specific complications of systemic disease, the effects of direct viral infection, or inflammation of the nervous system and vasculature, which can be para-infectious or post-infectious. In one national registry of 125 patients with COVID-19 and neurological or psychiatric disease reported over a 3-week period, 62.0% patients had a cerebrovascular event: 46.0% ischemic strokes, 7.0% intracerebral hemorrhage. As the epidemic is ongoing, the pattern of neurological manifestations of COVID-19 especially CNS features still not well known worldwide. Whether the neurological features are directly attributed to the virus or it is just an associated infection is also a big question. In this study it had been described the pattern of stroke in COVID-19 patients admitted in a dedicated stroke unit and their short term outcome.

**Methodology**

This was a single arm, single centre prospective cohort study conducted in Stroke Unit of National Institute of Neurosciences and hospital, Dhaka, Bangladesh from April to August, 2020. The study included acute stroke patients irrespective of age and sex who were found RT-PCR positive for SARS CoV-2. The sampling method was purposive. All patients were clinically evaluated by Glasgow Coma Scale (GCS) and they were also evaluated by CT Brain imaging, Chest X-ray and routine hematology and blood biochemistry. After diagnosis patients were referred to a covid dedicated hospital and we followed up the patients over phone for 4 weeks from the day of admission in stroke unit to assess short term outcome. All the data were collected in a semi structured questionnaire by study physicians and analyzed by SPSS version 22. Chi-Square test was done to see the association of outcome with types of stroke.

**Results**

Total 41 patients were enrolled in the study. Among them 21 (51.2%) male and 20 (48.8%) were female with a male to female ratio of 1.05:1. Their age ranging from 26 to 90 year where most (26.8%) of the patients were in 60-69 age group and 58.5% patients were aged 60 and above. Majority (58.5%) of them came from different urban areas. Common co-morbidities were Hypertension (68.3%), Diabetes mellitus (39%) and chronic kidney disease (2.4%) (Table 1).

| Age Group       | Frequency | Percent |
|-----------------|-----------|---------|
| 20 to 29 Years  | 3         | 7.3     |
| 30 to 39 Years  | 3         | 7.3     |
| 40 to 49 Years  | 3         | 7.3     |
| 50 to 59 Years  | 8         | 19.5    |
| 60 to 69 Years  | 11        | 26.8    |
| 70 to 79 Years  | 8         | 19.5    |
| 80 to 89 Years  | 2         | 4.9     |
| More than 90 Years | 3   | 7.3     |

**Gender**

| Gender | Frequency | Percent |
|--------|-----------|---------|
| Male   | 21        | 58.5    |
| Female | 20        | 48.8    |

**Residence**

| Residence | Frequency | Percent |
|-----------|-----------|---------|
| Urban     | 24        | 58.5    |
| Rural     | 17        | 41.5    |

**Co-morbidities**

| Co-morbidities | Frequency | Percent |
|----------------|-----------|---------|
| HTN           | 28        | 68.3    |
| DM            | 16        | 39.0    |
| CKD           | 1         | 2.4     |

Common presenting features were hemiplegia (48.8%), altered level of consciousness (39%), headache (12.2%) and convulsion (7.3%). Clinical evaluation revealed GCS of 10 or less in 70.7% patients and oxygen saturation of 90% or less in 56.1% patients (Table 2).

| Features | Frequency | Percent |
|----------|-----------|---------|
| Loss of consciousness | 16 | 39.0 |
| Hemiplegia | 20 | 48.8 |
| Aphasia | 2 | 4.9 |
| Headache | 5 | 12.2 |
| Convulsion | 3 | 7.3 |
| Confusion | 10 | 24.4 |

**Examination findings GCS**

| <7 | 14 | 34.1 |
| 7 to 10 | 15 | 36.6 |
| >10 | 12 | 29.3 |

**SpO2**

| SpO2 | Frequency | Percent |
|-------|-----------|---------|
| 60 to 80% | 6 | 14.6 |
| 80 to 90% | 17 | 41.5 |
| >90% | 18 | 43.9 |

CT brain imaging revealed infarct in 39% and hemorrhage in 51.2% cases. Chest radiography revealed abnormality in 14.6% cases (Table 3).
Corona virus disease 2019 (COVID-19) caused by SARSCov-2 virus, first emerged from Wuhan city, Hubei province of China. The epidemic spread rapidly worldwide within 3 months and was declared as a pandemic. SARSCov-2 virus was a causative agent for stroke or it can be para-infectious or post-infectious. In one study, meningitis was reported in 1% cases. Another study showed that the mortality rate was 48.8% which is much higher than those reported in other studies (27.6%). This higher rate is probably due to the inclusion of both types of stroke (infarct and hemorrhage) patients. Death was relatively higher in hemorrhagic stroke in comparison to ischemic stroke and subarachnoid hemorrhage (SAH) but that was not statistically significant. It may be due to the sample size was comparatively small.

Regarding the short term outcome, total 20 (48.8%) patients died where 7 died before referral to COVID-19 dedicated hospital. Death was relatively higher in cases of hemorrhagic stroke but association of outcome with types of stroke was not statistically significant (p value>0.05) (Table 4).

Table 4: Association of Outcome with Types of Stroke

| Stroke type          | Recovered | Death (%) | P value |
|----------------------|-----------|-----------|---------|
| Ischemic stroke      | 12 (29.3%)| 8 (19.5%) |         |
| Hemorrhagic stroke   | 5 (12.2%) | 10 (24.4%)| 0.211   |
| SAH                  | 4 (9.7%)  | 2 (4.9%)  |         |
| Total                | 21 (51.2%)| 20 (48.8%)|         |

Discussion

SARS Cov-2 virus infection is a new disease worldwide. Though the virus primarily affects respiratory tract, there are several reports on various neurological presentation associated with COVID-19 worldwide. This small crosssectional study represents the pattern of stroke in a stroke unit of a tertiary care hospital. We found around almost equal number of patients in both sex. Their age ranging from 26 to 90 years where 58.5% patients were aged 60 and above. Study from China and Iran reported similar age of COVID-19 patients presenting with symptoms of stroke. Old age is an independent risk factor for stroke. Some case series also demonstrated stroke in relatively younger COVID-19 patients. In this study around one fifth (21.9%) of the patients are relatively younger (<50 years). Majority (58.5%) of our patients came from different urban areas which may be due to the fact that the initial spread of the virus was more in the urban region. Hypertension (68.3%) and diabetes (39%) were the most common co-morbidities among the patients. Similar findings were observed in other studies.

Common presentation among the stroke patients were hemiplegia (48.8%), altered level of consciousness (39%), headache (12.2%), convulsion (7.3%). Hussain et al from Bangladesh showed almost same clinical presentation among the neurological COVID-19 patients. CT brain imaging revealed infarct in 39% and hemorrhage in 51.2%. Asadi-Pooya and Simani reported acute ischemic stroke in 5% and acute hemorrhagic stroke in 0.5% of the patients in their systematic review of central nervous system manifestation of COVID 19 patients. Chest radiography by plain X-ray revealed opacities in the lungs in 17.1% cases in this study. The sensitivity of CT chest is obviously much higher. Ntaios et al found opacities in the lungs in 89.3% chest CT. The mortality rate in this study was 48.8% which is much higher than those reported in other study (27.6%). This higher rate is probably due to the inclusion of both types of stroke (infarct and hemorrhage) patients. Death was relatively higher in hemorrhagic stroke in comparison to ischemic stroke and subarachnoid hemorrhage (SAH) but that was not statistically significant. It may be due to the sample size was comparatively small.

There is some limitation of this study. It is a single centre small cross-sectional study. Whether SARSCov-2 virus was a causative agent for stroke or it was just a co-infection in this pandemic situation is not certain as brain autopsy was not done in any patient. It was not possible to establish whether Stroke and COVID 19 was a mere coincidence or there had been a causal relationship.

Conclusion

In conclusion hemorrhagic stroke was the most commonly occurring stroke among the COVID-19 patients and associated with poor outcome. Whether SARSCov-2 virus was a causative agent for stroke or it was just a co-infection in this pandemic situation is not certain. Therefore, the exact pathophysiology behind the stroke is still clearly unknown. Further large scale multicenter study is warranted.

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