The effect of the thrombolytic therapy on the early rehabilitation of patients with acute ischemic stroke – study report

MUTU Cătălin Cosmin 1,2, CEREI Larisa-Georgiana 1

Corresponding author: MUTU Cătălin Cosmin, E-mail: cosmin.mutu@ulbsibiu.ro

Abstract
Introduction. r-TPA intravenous thrombolytic therapy is a big step in acute management of ischemic stroke (IS) but is burdened by a small therapeutic window (max. 4.5 hours) that restricts patient access. NIH stroke scale (NIHSS) is the largest worldwide tool used by doctors in order to objectively quantify the severity of neurological impairment in acute IS, in the initial stage as well as in the subsequent stages. Material and method. The aim of this paper is to present the results of an observational, prospective, analytical study on a number of 110 patients with acute IS, hospitalized within two months in the Neurology Clinic of Sibiu (19 patients with thrombolytic therapy applied, 91 patients with classic therapy). NIHSS has been applied to all patients on the first day of the admission and at the discharge date. Results and discussions. The mean value of NIHSS at admission and at discharge (A/D) were 12,84/4.06 points for the patient with thrombolytic therapy and 7,73/4.45 points for the patients without thrombolytic therapy. In hospital NIHSS score reduction, meaning early recovery of patients, was consistent with type of applied therapy (8.78 vs 3.28 points). The pattern of neurological impairment is specific for each subtype of IS. Conclusions. The degree of early recovery was superior for patients with versus without thrombolytic therapy (68.4% versus 42.4%). Average hospital care period was reduced with 8,33% for patients with thrombolytic therapy (11,4 versus 12,2 days care in hospital).

Keywords: ischemic stroke, early rehabilitation, NIH stroke scale, neurological pattern.

Introduction
Stroke represents one of the leading causes of morbidity, mortality and persistent disability worldwide. Ischemic stroke is a heterogeneous disease with distinct subtypes, each of them presenting specific etiological, pathogenesis and clinical aspects (1). The widespread introduction of IV thrombolytic therapy with r-TPA in Romania, starting with January 2019, has probably significantly improved the chances of patients with acute ischemic stroke for a favorable evolution. Unfortunately, patients access to thrombolytic therapy is limited by the reduced window of time to only 4.5 hours after the onset of a stroke. As clinician-reported scales of global disability, modified Rankin Scale (mRS) and National Institute of Health Stroke Scale (NIHSS) are the tools used by doctors in order to objectively quantify the severity of neurological impairment in acute ischemic stroke, in the initial stage as well as in the subsequent stages (1,2,3).

Patient-reported scales to be used in the primary care medicine are: The Patient-Reported Outcome Measure Information System (PROMIS) and Quality of Life in Neurological Disorders (NeuroQoL) scales (4,5).

Purpose
The aim of this study was to evaluate the neurological impairment, as well as the assessment of the short-term evolution of inpatients in the Neurology Clinic of Sibiu, by analyzing the dynamics of all 11 items of the NIHSS in the two key moments of hospitalization (admission and discharge) taking into consideration the access / lack of access of these patients to thrombolytic therapy.

Materials and methods
We conducted an observational, prospective, analytical study of 110 patients with acute ischemic stroke, hospitalized within two months in the Neurology Clinic of Sibiu (March-April 2019). The patients were divided into two groups according to the therapy applied (patients with or without thrombolysis). NIHSS has been evaluated in all patients on the first day of the admission and at the discharge date. The primary objective was to establish correlations between the recovery degree of patients and the type of applied therapy. The secondary objective was to determine the pattern of neurological damage in patients included in study and to achieve correlations between the extent of neurological damage, type of therapy applied and the degree of patients’ early recovery.

Results
All 110 patients with acute ischemic stroke (mean age 71,42 +/- 20 years) hospitalized during the study period were included. In respect to the type of therapy, the patients were divided into two subgroups with specific characteristics presented in the synoptic table below.
Table 1. Patients’ demographic and clinical characteristics according to applied therapy

<table>
<thead>
<tr>
<th>Items</th>
<th>Total</th>
<th>r-TPA treated</th>
<th>Non r-TPA treated</th>
</tr>
</thead>
<tbody>
<tr>
<td>Patients</td>
<td>110 (100)</td>
<td>19 (17.3)</td>
<td>91 (82.7)</td>
</tr>
<tr>
<td>Mean age: y</td>
<td>71.9</td>
<td>71.94</td>
<td>71.91</td>
</tr>
<tr>
<td>Female</td>
<td>50 (45.4)</td>
<td>8 (42.1)</td>
<td>42 (46.1)</td>
</tr>
<tr>
<td>Male</td>
<td>60 (54.5)</td>
<td>11 (57.9)</td>
<td>49 (53.8)</td>
</tr>
<tr>
<td>Mean H days: n</td>
<td></td>
<td></td>
<td></td>
</tr>
<tr>
<td>In H deaths</td>
<td>12 (10.9)</td>
<td>3 (15.8)</td>
<td>9 (9.9)</td>
</tr>
<tr>
<td>Mean NIHSS</td>
<td></td>
<td></td>
<td></td>
</tr>
<tr>
<td>Admission: p</td>
<td>8.62</td>
<td>12.84</td>
<td>7.73</td>
</tr>
<tr>
<td>Discharge: p</td>
<td>4.39</td>
<td>4.06</td>
<td>4.45</td>
</tr>
<tr>
<td>Vascular risk factors: n (%)</td>
<td></td>
<td></td>
<td></td>
</tr>
<tr>
<td>Hypertension</td>
<td>58 (52.7)</td>
<td>7 (6.4)</td>
<td>51 (46.4)</td>
</tr>
<tr>
<td>Diabetes mellitus</td>
<td>26 (23.6)</td>
<td>4 (3.6)</td>
<td>22 (20)</td>
</tr>
<tr>
<td>Hyperlipidemia</td>
<td>28 (25.4)</td>
<td>7 (6.4)</td>
<td>21 (19.1)</td>
</tr>
<tr>
<td>Atrial fibrillation</td>
<td>28 (25.4)</td>
<td>7 (6.4)</td>
<td>21 (19.1)</td>
</tr>
<tr>
<td>Stroke</td>
<td>26 (23.6)</td>
<td>1 (0.9)</td>
<td>25 (22.7)</td>
</tr>
</tbody>
</table>

NIHSS: NIH stroke scale, H days: hospitalization period, In H death: in hospital deaths. Values are number (%) unless otherwise stated.

In our clinic, the mean hospitalization period of all patients in this study was 12.02 days. For the whole study group, the mean value of NIHSS was 8.62 points at admission and 4.39 points at discharge.

From a global perspective, during the hospitalization in our clinic, the degree of early recovery of neurological deficits was 49.07% (see fig. 1).

The percentage of patients who underwent thrombolytic therapy was 17.27%. The mean value of NIHSS at admission and at discharge (A/D) were 12.84/4.06 points for the patient with thrombolytic therapy and 7.73/4.45 points for the patients without thrombolytic therapy (see fig. 2).

The mean value of in-hospital neurological recovery was superior for r-TPA treated patients (8.78 points in NIHSS score reduction) versus non r-TPA treated patients (3.28 points in NIHSS score reduction). The degree of early recovery of neurological deficits was 68.4% for first subgroup of patients and 42.4% for second subgroup. As expected, the highest grade of early recovery was observed in patients with thrombolytic therapy.

The data from our study revealed that facial and arm motor deficit with language functions were more affected (maximal scores), compared with cognitive and sensory functions (minimal scores). (see fig. 3)
Studying the pattern of neurological damage in the first and in the last day of hospitalization in each patient from this study group, correlated with the access at r-TPA therapy, made us observe specific neurological patterns (see next radar type charts).

Recovery of the motor deficit (lower/upper limbs and language) and verbal response was significantly greater in patients with thrombolytic therapy compared with the recovery of patients who did not received thrombolytic therapy. Recovery of sensitive and sensorial deficits has not varied significantly in the two therapeutic subgroups.

**Discussion**

Studies in humans and animal models show that most recovery from impairment occurs in the first 1-3 months after stroke as a result of both spontaneous recovery as well as increased responsiveness to enriched environments and training. Improvement from impairment is attributable to a short-lived "sensitive period" of post-stroke plasticity defined by unique genetic, physiological, and structural events (6).

Overall, trials of rehabilitation in the first 2 weeks after stroke are scarce. In the realm of very early mobilization, one large and one small trial found potential harm from mobilizing patients within the first 24 h after stroke, and only one small trial found benefit in doing so (7,8).

However, thrombolytic treatment applied to patients who were admitted in the hospital in the first 4.5 hours window, significantly reduced the neurological deficit measured by the NIHSS from 12.84 to 4.06 points, averaging 8.78 points. Although the patients who underwent thrombolytic therapy had at the beginning a higher neurological damage than the other ones (12.84 vs. 7.73), at discharge their neurological status was better than that of those without thrombolytic therapy (4.06 vs. 4.45).

The NIHSS predicts post-acute care disposition among stroke patients. Predicting disposition on the first day of admission may facilitate the time-consuming and costly process of securing a bed at rehabilitation and perhaps decrease unnecessary length of stay in acute care settings (4,8,9).

After the initial acute stroke therapy, applying the same rehabilitation methods in acute care settings, the hospitalization duration of patients who underwent thrombolysis was reduced by 8.33% compared to non-thrombolized (11.4 vs 12.2). Much more important was the extent of early recovery of neurological deficits (68.4% vs. 42.4%). A high degree of early recovery from neurological deficits reduces the need for expensive care in the rehabilitation department (7,10).

The degree of early recovery of non-thrombolized patients in this study (42.4%) was comparable to the degree of early recovery (45.12%) obtained in a similar study conducted in our clinic 5 years ago, when we did not have the option of thrombolytic therapy for patients with acute ischemic stroke. The severity at onset of ischemic stroke in thrombolized patients in the current study was comparable to that in patients in the previous study (mean NIHSS at admission of 12.84 points vs. 12.17 points) (11,12,13).
Conclusions
Early recovery of motor deficits is dependent on the type of therapy applied.

The degree of early recovery was superior for patients with thrombolytic therapy versus other conventional therapies (68.4% vs 42.4%). The thrombolytic therapy reduced the average hospital care period (in acute care settings) with 8.33% for patients arriving at the hospital in a timely manner.

Continued efforts are needed to improve the response time of the population and emergency transport services to the hospital at the occurrence of an acute ischemic stroke, but also an increase in the number of emergency rooms and acute stroke units in which the application of thrombolytic therapy in optimal conditions is available.

Declaration of conflict of interests/Conflict of Interest Statement
The authors declare that there is no conflict of interest regarding the publication of this article.

Informed consent
Informed consent was obtained from all patients included in this study.

References: