Comparison of the Effects of Magnetically Controlled Growing Rod and Traditional Growing Rod Techniques on the Sagittal Plane in the Treatment of Early-Onset Scoliosis

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Objective: Comparing the effects of magnetically controlled growing rod (MCGR) and traditional growing rod (TGR) techniques on the sagittal plane in the treatment of early-onset scoliosis (EOS).

Methods: Twelve patients were operated using dual MCGR technique in one center, while 15 patients were operated using dual TGR technique for EOS in another center. Patients’ demographic characteristics, complications and radiological measurements such as cobb angle, thoracic kyphosis, lumbar lordosis, T1–S1 range (mm), proximal junctional angle, distal junctional angle, sagittal balance, coronal balance, pelvic incidence, sacral slope and pelvic tilt were assessed and compared in preoperative, postoperative and last follow-up period.

Results: Age and sex distributions were similar in both groups. The mean number of lengthening in the MCGR group was 12 (8–15) and 4.8 (3–7) in the TGR group. Two techniques were shown to be effective in controlling the curvature and in the increase of T1–S1 distance. In TGR group, four patients had rod fractures, six patients had screw pull-out and four patients had an infection, whereas three patients had screw pull-out and one patient had infection complications in the MCGR group.

Conclusion: There was no significant difference between the two groups in terms of cobb angle, coronal and sagittal balance and sagittal pelvic parameters. MCGR can cause hypokyphosis and proximal junctional kyphosis in a minimum 2-year follow-up period. The implant-related complications were less in the MCGR group. However, larger case groups and longer follow-up periods are required for the better understanding of the superiority of one method on other in terms of complications.

Key Words: Early-onset scoliosis ∙ Growing rod ∙ Magnetic growing rods ∙ Spinopelvic parameters ∙ Proximal junctional angle.

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INTRODUCTION

Early onset scoliosis (EOS) is defined as curvature of the spine in children seen before 10 years of age which might be depending on various etiologies. Definitive surgical treatments with a large fusion performed on young children with severe spinal deformity may result in severe pulmonary insufficiency, short body stature and growth retardation. In order to prevent these problems, physicians are in search of a treatment modality that allows the thorax to grow and at the same time correcting the deformity. Many techniques have been proposed to prevent early spinal fusion. The goal of these methods is to keep spinal deformity under control and to provide the optimum space for the lungs, until the child reaches skeletal maturity.

Traditional growing rod (TGR) technique has been shown to be effective in achieving body lengthening and maturation while spinal growth continues in early-onset scoliosis. However, there are concerns that TGR use may cause recurrent surgical intervention and associated complications, as well as leading to worsened sagittal plane deformity of posterior-based distraction systems and increasing the risk of junction problems. As a result of these problems encountered with the conventional methods, a less invasive method which do not require repetitive surgical interventions and has a lower risk of autofusion development, magnetically controlled growing rod (MCGR) systems have been developed.

Little attention has been paid to the sagittal plane and spinopelvic parameters, focusing on existing GR studies, coronal plane deformity, spine height and lung capacity. The importance of spinopelvic parameters on the patient’s clinical results is well known in the literature and, as a result, is becoming increasingly important in the spinal deformity research. As the changes in the sagittal plane began to gain importance in spinal deformity surgery, studies on alterations in sagittal plane after TGR and MCGR were performed. However, a web-based search using databases PubMed and Google scholar regarding the comparison of the effects of TGR and MCGR on sagittal plane did not yield a positive result. Therefore, we aimed to contribute to the literature data by comparing the effects of TGR and MCGR on sagittal plane in treatment of EOS.

MATERIALS AND METHODS

The study was approved in advance by the Institutional Review Board of University of Kyrenia (RY-2019-10) and all patients signed an informed consent form. We evaluated patients who were operated in two centers due to EOS with dual TGR or dual MCGR techniques, retrospectively. Inclusion criteria in our study were determined as having a major curvature of 30 degrees or more with at least 3 previous lengthening procedures, previous experience of an extended fusion surgery or an appointment for surgery and fulfilling the follow-up period of at least two years after the first operation. The etiology of all 27 patients included in our study was idiopathic EOS. Twelve patients from one center were operated with dual MCGR technique, while 15 patients were operated with dual TGR technique in another center.

Surgical techniques and distraction interventions

The patients were placed in the prone position on a spinal frame under general anesthesia. Neuromonitorisation was used in all surgical procedures. After confirming the levels where the proximal and distal vertebrae pedicle screws would be implanted under fluoroscopy control, two separate subperiosteal dissections were carried out in these regions and a pedicle screw was implanted by preserving the facet joint structures. Two or three levels of spine were instrumented in the upper and lower segment. The number of instrumented level in upper and lower segment depends on the surgeon’s preference based on patient’s weight and the degree of curvature. Sublaminar hooks were not used for any of the patients. Then, appropriate kyphosis and lordosis were given from the upper part of the rod and the lower part of the rod by protecting the actuator (5.5 mm rod thickness). The magnetic rod was submuscularly placed between the proximal and distal screws, and fixed onto the pedicle screws for each side. In order to avoid junctional kyphosis and sagittal plane decompensation, the proximal interspinous ligaments were secured.

In TGR technique, lengthening was performed periodically.
through a posterior small midline incision over the connectors, which was routinely checked with 6–10 months intervals. After reaching the connectors, the connector screws that were connected to the proximal rod were loosened and a regular distractor was placed between the connector and the rod holder attached on the proximal rod for a gentle distraction. The connector screws were then tightened. In the MCGR technique, the magnetic rod was lengthened at 12-week intervals. In outpatient clinic conditions, the actuator portion of the magnetic rod was found with the help of a magnet while the patient was in the prone position, and the skin level was marked with a marker. The rod was lengthened through the actuator portion with the help of remote control by gently applying traction from the legs and under the armpit. Four mm lengthening was obtained in each session. Distraction numbers of both groups were noted.

Radiologic assessment

Patients’ radiological measurements such as cobb angle, thoracic kyphosis (TK), lumbar lordosis (LL), T1–S1 range (mm), proximal junctional angle (PIA), distal junctional angle (DJA), sagittal balance, coronal balance, pelvic incidence, sacral slope and pelvic tilt were assessed and compared in preoperative, postoperative and last follow-up period (before definitive spinal fusion surgery). The proximal junctional kyphosis was evaluated on the upper endplate of the vertebra which is at the two levels upper position from the last instrumented vertebra, and on the lower endplate of the instrumented vertebra which is at the top level. Distal junctional kyphosis was calculated from the upper endplate of the lowest instrumented vertebra and the lower endplate of the one-down vertebra. Spinal and pelvic parameters were measured with Surgimap 2.2.15.1 version (Nemaris Inc., New York, NY, USA). Two authors (S.E. and B.P.) assessed all images and measured all angles independently. The images were blinded and randomized. Where there was disagreement between the examiners, re-evaluation was performed until a consensus was reached.

Statistical analysis

The analyses were performed by using the SPSS software (Statistical Package for the Social Sciences, version 20.0; SPSS Inc., Chicago, IL, USA). Descriptive statistics were expressed as numbers and percentages for categorical variables. Descriptive statistics; mean, the standard deviation were given for numerical variables. The difference in categorical variables be-

| Parameter                        | All participants (n=27) | MCGR (n=12) | TGR (n=15) | p-value |
|----------------------------------|-------------------------|-------------|------------|---------|
| Age (years)                      | 8.6 (6–11)              | 8.7 (7–11)  | 8.5 (6–11) | 0.282   |
| Gender                           | 18 F–9 M                | 7 F–5 M     | 11 F–4 M   | 0.448   |
| Number of lengthening            | 8.0 (3–15)              | 12 (8–15)   | 4.8 (3–7)  | <0.001  |
| Mean follow up (months)          | 38.7±8.8 (24–57)        | 36±6.6 (24–45) | 40±8±10 (24–57) | 0.186  |
| Thoracic kyphosis (º)            | 49.2±14.3               | 48.5±16.3   | 49.8±13.1  | 0.922   |
| Lumbar lordosis (º)              | −49.1±10.8              | −43.9±10.4  | −53.2±9.6  | 0.033   |
| Proximal junctional angle (º)    | 8.6±4.8                 | 8.9±4.9     | 8.4±4.9    | 0.806   |
| Distal junctional angle (º)      | −13.7±10.4              | −12.4±13.5  | −14.8±7.3  | 0.788   |
| Pelvic incidence (º)             | 42.5±10.4               | 39.5±9.1    | 44.8±11.1  | 0.096   |
| Sacral slope (º)                 | 34.1±8.1                | 29.9±5.9    | 37.4±8.2   | 0.032   |
| Pelvic tilt (º)                  | 8.5±8.2                 | 9.6±8.8     | 7.6±7.9    | 0.494   |
| T1–S1 range (mm)                 | 296±38                  | 280±48      | 309±21     | 0.056   |
| Cobb angle (º)                   | 52.6±12.5               | 52.7±15     | 52.6±10.6  | 0.574   |
| Sagittal balance (mm)            | −15.7±27.9              | −21.6±15.4  | −10.3±34.8 | 0.558   |
| Coronal balance (mm)             | 19±10.6                 | 24±8.8      | 14.9±10.5  | 0.015   |

Values are presented as mean±standard deviation (range) or number (range). MCGR : magnetically control growing rod, TGR : traditional growing rod, F : female, M : male
between groups was evaluated using chi-square analysis. Mann Whitney U test was used for the comparisons between the two independent groups when the numerical variables were not normally distributed. Changes in findings between the preoperative, postoperative and the last follow-up examinations were tested using repeated measures using analysis of variance with a Greenhouse-Geisser correction. Spearman’s correlation coefficient was used for evaluation of relationships between parameters. Statistical significance was accepted as \( p < 0.05 \).

### RESULTS

Age and gender distributions were similar in both groups. The baseline data and preoperative values of the patients are given in Table 1. The preoperative, early postoperative and final control radiological data of the patients who were operated with MCGR technique are given in Table 2 (Figs. 1 and 2), and those operated with TGR technique are given in Table 3 (Figs.

### Table 2. MCGR radiological results in preoperative, postoperative and last follow-up period

| Parameter                      | Preoperative | Postoperative | Last follow-up | \( p \)-value |
|--------------------------------|--------------|---------------|----------------|--------------|
| Thoracic kyphosis (º)         | 48.5±16.3    | 27.8±13.0     | 23.1±6.7       | <0.001       |
| Lumbar lordosis (º)           | −43.9±10.4   | −32.7±13.7    | −39.0±10.8     | 0.071        |
| Proximal junctional angle (º) | 8.9±4.9      | 11.7±7.8      | 16.5±7.4       | 0.001        |
| Distal junctional angle (º)   | −12.4±13.5   | −8.5±13.5     | −10.3±16.5     | 0.678        |
| Pelvic incidence (º)          | 39.5±9.1     | 38.9±13.1     | 42.4±8.3       | 0.319        |
| Sacral slope (º)              | 29.9±5.9     | 33.9±7.2      | 35.9±8.7       | 0.04         |
| Pelvic tilt (º)               | 9.6±8.8      | 5.0±14.4      | 6.5±9.0        | 0.208        |
| T1–S1 range (mm)              | 280.2±48.8   | 309.8±46.7    | 352.1±51.0     | <0.001       |
| Cobb angle (º)                | 52.7±15.0    | 27.8±13.0     | 32.2±15.4      | <0.001       |
| Sagittal balance (mm)         | −21.6±15.4   | −13.7±36.8    | −4.7±31.2      | 0.027        |
| Coronal balance (mm)          | 24.0±8.8     | 10.3±8.0      | 13.0±8.5       | 0.001        |

Values are presented as mean±standard deviation. MCGR: magnetically controlled growing rod.

**Fig. 1.** Preoperative AP/lateral X-ray of patient treated with MCGR technique. AP: anteroposterior, MCGR: magnetically controlled growing rod.

**Fig. 2.** Last follow-up AP/lateral X-ray of patient treated with MCGR technique. AP: anteroposterior, MCGR: magnetically controlled growing rod.
The comparison of the radiological findings of the two groups on the postoperative and final control visits is given in Table 4.

No correlation was observed between TK, PJA, and DJA in the postoperative period and last postoperative visit in both patient groups. Correlation was not detected between postoperative LL and DJA in both patients groups. In patients who underwent MCGR, the number of lengthening was positively correlated with postoperative PJA and negatively correlated with postoperative DJA ($r=0.674, p=0.016; r=-0.766, p=0.004$, respectively). There was a negative correlation detected between PJA and DJA in the postoperative period in patients who underwent MCGR ($r=-0.687, p=0.014$).

Table 3. TGR radiological results in preoperative, postoperative and last follow-up period

| Parameter                              | Preoperative | Postoperative | Last follow-up | $p$-value |
|----------------------------------------|--------------|---------------|----------------|-----------|
| Thoracic kyphosis (º)                  | 49.8±13.1    | 38.4±12.0     | 39.2±11.7      | 0.004     |
| Lumbar lordosis (º)                    | -53.2±9.6    | -43.2±12.1    | -47.0±11.9     | 0.009     |
| Proximal junctional angle (º)          | 8.4±4.9      | 6.5±5.4       | 10.8±6.4       | 0.166     |
| Distal junctional angle (º)            | -14.8±7.3    | -14.1±11.5    | -15.8±8.8      | 0.591     |
| Pelvic incidence (º)                   | 44.8±11.1    | 43.0±11.7     | 45.0±10.7      | 0.455     |
| Sacral slope (º)                       | 37.4±8.2     | 34.1±6.3      | 35.2±7.5       | 0.183     |
| Pelvic tilt (º)                        | 7.6±7.9      | 9.1±7.3       | 9.6±7.7        | 0.467     |
| T1–S1 range (mm)                       | 309.4±21.5   | 333.0±27.3    | 362.2±27.7     | <0.001    |
| Cobb angle (º)                         | 52.6±10.6    | 30.4±10.2     | 26.5±12.5      | <0.001    |
| Sagittal dege (mm)                     | -10.3±34.8   | 0.1±33.5      | -3.0±40.6      | 0.628     |
| Coronal dege (mm)                      | 14.9±10.5    | 9.2±7.0       | 8.2±6.7        | 0.067     |

Values are presented as mean±standard deviation. TGR : traditional growing rod.

In patients who underwent TGR, the number of lengthening was detected positively correlated with postoperative and last postoperative PJA ($r=0.683, p=0.010; r=0.687, p=0.005$, respectively), and negatively correlated with postoperative and last postoperative DJA ($r=-0.818, p<0.001; r=-0.542, p=0.037$, respectively). In patients who underwent TGR operation, a negative correlation was detected between PJA and DJA in the postoperative period and last postoperative period ($r=-0.528, p=0.043; r=0.520, p=0.047$).
Superficial wound infection in one patient and screw pull out complication in three patients were encountered in the MCGR group. Superficial wound infection in three patients, deep tissue infection in one patient, 6-times rod breakages in four patients and 7-times screw pull out complications in six patients were occurred in TGR group.

**DISCUSSION**

In the TGR application, the requirement of periodic lengthening in the operation room every 6 to 9 months, and the requirement of a total of 6 to 18 scheduled surgical procedures from the beginning of treatment are the disadvantages of this technique. In addition, there is an increased risk of complications such as surgical wound complications, the termination of the growth by stiff spine development (autofusion), deteriorated chest development, and thoracic insufficiency syndrome. In MCGR application, it is thought that the increase in the frequency of lengthening may reduce the development of the autofusion by meeting the spine growth more closely. There is an opinion that, with the more frequent lengthening in MCGR, which is possible as outpatient, the rods would more closely match up with spinal growth and possibly stress may be reduced throughout the implants. In our study, in the TGR technique, rod breakage in four patients (27%), screw pullout complication in six patients (40%) were encountered, while screw pullout complication in only three patients (25%) were encountered in patients who underwent surgery with MCGR technique.

Akbarnia et al. and Shah et al. have observed a decrease in TK after the first implantation, however, they have observed an increase in the last follow-up. Similarly, LL has initially decreased, and increased at the latest follow-up. In their study comparing TGR and MCGR, Akbarnia et al. reported a decrease in postoperative value of TK compared to the preoperative value in both groups, and a tendency to increase in the last follow up. In our study, in line with the literature, we have observed a decrease in the early postop period in TK and LL values, however, in the last follow up, there was an increase in the patient group treated with TGR, while the decrease was continuing in the patients treated with MCGR. Since the kyphosis could not be given to the magnetic rod actuator part and this part coincided with the patient’s kyphosis region, we think that hypokyphosis might have been developed in the MCGR group.

In order to prevent proximal junctional kyphosis in patients whom they applied TGR, Akbarnia and Emans suggested that rods should be countered in the shape of kyphosis and interspinal ligaments should be kept as strong as possible, upper level should be limited at T2 level and the upper level should rarely be reached, and mentioned that in the presence of thoracic hyperkyphosis, the excessive correction of the rods may lead to implant failure after surgery. In their study in which they have applied MCGR and evaluated the proximal junctional kyphosis, Inaparthy et al. have emphasized that male, syndromic, hyperkyphotic and younger patients should be closely followed-up in terms of proximal junctional kyphosis.

| Table 4. Comparasion of MGRS and TGR radiological results in postoperative and last follow-up period |
|---------------------------------------------------------------|
| Postoperative | MCGR (n=12) | TGR (n=15) | p-value |
| Thoracic kyphosis (º) | 27.8±13.0 | 38.4±12.0 | 0.001 |
| Lumbar lordosis (º) | −32.7±13.7 | −43.2±12.1 | 0.032 |
| Proximal junctional angle (º) | 11.7±7.8 | 8.6±5.4 | 0.378 |
| Distal junctional angle (º) | −8.5±13.5 | −14.1±11.5 | 0.340 |
| Pelvic incidence (º) | 38.9±13.1 | 43.0±11.7 | 0.497 |
| Sacral slope (º) | 33.9±7.2 | 34.1±6.3 | 0.825 |
| Pelvic tilt (º) | 5.0±14.4 | 9.1±7.3 | 0.478 |
| T1–SI range (mm) | 309.8±46.7 | 333±27.3 | 0.139 |
| Cobb angle (º) | 27.8±13.0 | 30.4±10.2 | 0.420 |
| Sagittal balance (mm) | −13.7±36.8 | 0.1±33.5 | 0.366 |
| Coronal balance (mm) | 10.3±8.0 | 9.2±7.0 | 0.732 |

| Last follow-up | MCGR (n=12) | TGR (n=15) | p-value |
| Thoracic kyphosis (º) | 23.1±6.7 | 39.2±15.5 | 0.03 |
| Lumbar lordosis (º) | −39.0±10.8 | −47.0±11.9 | 0.087 |
| Proximal junctional angle (º) | 16.5±7.4 | 10.8±6.4 | 0.056 |
| Distal junctional angle (º) | −10.3±16.5 | −15.8±8.8 | 0.825 |
| Pelvic incidence (º) | 42.8±8.3 | 45.0±10.7 | 0.303 |
| Sacral slope (º) | 35.9±8.7 | 35.2±7.5 | 0.941 |
| Pelvic tilt (º) | 6.5±9.0 | 9.6±7.7 | 0.463 |
| T1–SI range (mm) | 352.1±51 | 362.2±27.7 | <0.999 |
| Cobb angle (º) | 32.2±15.4 | 26.5±12.5 | 0.366 |
| Sagittal balance (mm) | −4.7±31.2 | −3.0±40.6 | 0.282 |
| Coronal balance (mm) | 13.0±8.5 | 8.2±6.7 | 0.083 |

Values are presented as mean±standard deviation. MCGR : mayyetik control growing rod, TGR : traditional growing rod.
In our study, we found that the increase in PJA was higher in the MCGR group in the last follow-up, although it did not reach statistical significance. It is evident that the PJA values increased as the TK values of the patients decreased. Although there was an increase in PJA values in the early postoperative period and last postoperative period in patients who underwent TGR operation, there was no correlation between TK and PJA. Since inadequate kyphosis was given to the actuator part in the MCGR group, patients have less kyphosis angle than the TGR group in postoperative and future postoperative follow-up. We think that this situation may be effective in the increase of PJA in MCGR group in the follow-ups. Besides, we think that further lengthening in the MCGR group compared to the TGR group may be the reason for distractive forces to increase more in the adjacent segments and higher increase rate in PJA. It should be kept in mind that repeated distraction below the existing upper thoracic kyphosis may worsen the upper deformity. Due to the small number of patients, we could not subgroup patients according to preoperative kyphosis values and this is one of the limitations of our study. Another limitation of our study was that the PJK operative kyphosis values and this is one of the limitations of our study. Another limitation of our study was that the PJK values were higher in the MCGR patient group at the last follow-up compared to the TGR patient group, but we could not find a statistically significant value due to the insufficient number of patients and follow-up time.

Lowe et al. have reported that preoperative thoracolumbar kyphosis was a risk factor for postoperative DJK development. Shah et al. have indicated that DJA showed the same pattern as LL and correlated with the change in LL and that DJA closely reflected the change in LL. In our study, we observed that both of our groups had the same correlative pattern between LL and DJA, however, this change was not statistically significant.

Some studies have shown that coronal and sagittal plane deformity can develop after initial surgery in both single and dual GR techniques. In a study, which included 67 patients and examined the effect of serial lengthening on sagittal balance (SB) in TGR, it was reported that the balance tended to come to neutral in the early postoperative period and it was found to be statistically significant, however, this significance was reported to be lost in future follow-up period. We observe in our study that the SB and coronal balance change tended to come to neutral in the MCGR group but may not provide information about the superiority of methods to each other due to small number of patients in the groups and short follow-up periods.

Pelvic incidence (PI) is defined independently of position and has been shown to play an important role in maintaining sagittal alignment. The presence of abnormal PI is a risk factor for sagittal imbalance and failure following adult scoliosis surgery. In their recent study, Atici et al. stated that pelvic parameters (PI, pelvic tilt, and sacral slope) remained unchanged in patients with EOS treated with GR. Shah et al. have found a 5-degree increase in PI compared to preoperative value, however, they have reported that this was not statistically significant. In our study, although we observed a statistically significant increase in the sacral slope in our follow-up in the MCGR group, we did not find any significant changes in other spinopelvic parameters and spinopelvic parameters in the TGR patient group. As a result, we have not determined statistically significant difference between the MCGR group and the TGR group in terms of spinal and spinopelvic parameters in the short-term follow-up.

Sankar et al. have shown that there was an inconsistency between the expected and the obtained net T1–S1 length in the case of repeated lengthening. They attributed this to the hardening of the spine due to autofusion. The effects of this phenomenon on the sagittal profile are still unclear. We provided statistically significant lengthening of T1–S1 distance in both groups in the follow-up period. When we compared the surgical technique, we could not find a statistically significant difference between the groups. Both methods are effective and successful in providing increase in length. In both methods, it should be considered that the planned and actual lengthening will not be the same.

Limitations of our study were; limited number of patients, being retrospective in the nature, short-term follow-up periods and performance of the two procedures by two different surgical teams and clinics.

Studies on TGR procedures have shown that wound and implant complication rates are as high as 58%. In their multicentric study, Choi et al. have shown that MCGR results in a lower rate of infections compared to TGR (11% in TGR versus 4% in MCGR). Rod breakage and screw pull out are frequently observed complications with TGR use and it was seen that the rod type or corset usage did not prevent these problems. It is thought that this is a problem regarding instrumentation, which cannot be overcome by any type of
implant, and is caused by spanning unfused spinal segments\(^2\). On the other hand, the MCGR is considered not eliminating implant-related common complications such as rod breakage or screw pull out\(^2\). The absence of lengthening after some lengthening procedures is specific to MCGR, however, does not always indicate a problem with the device\(^2\). Also in our study, implant-related complications were higher in the TGR group. However, planning and follow-up of the treatment (TGR or MCGR) according to the normal pediatric spinal sagittal parameters may help in preventing the complications.

**CONCLUSION**

When evaluating MCGR and TGR techniques in their own, we can obtain significant changes in Cobb angle, and T1–S1 values when we look at preoperative, early postoperative and last follow up values. In the evaluation of sagittal pelvic parameters, we observed the lack of significant difference between the two groups. MCGR can cause hypokyphosis and PJK in a minimum 2-year follow-up period. In order to prevent proximal junctional kyphosis, magnetic rods with different kyphotic degrees might be manufactured by the company and compatible magnetic rods with pelvic parameter and lumbar lordosis can be placed. The implant-related complications were less in the MCGR group. However, larger case groups and longer follow-up periods, may aid in the understanding the superiority of one method to other in terms of complications.

**CONFLICTS OF INTEREST**

No potential conflict of interest relevant to this article was reported.

**INFORMED CONSENT**

Informed consent was obtained from all individual participants included in this study.

**AUTHOR CONTRIBUTIONS**

Conceptualization : SE  
Data curation : ONÖ  
Formal analysis : YA  
Funding acquisition : ÇÖ  
Methodology : BP  
Project administration : SE  
Visualization : BP, ONÖ  
Writing - original draft : SE, BP  
Writing - review & editing : A, ÇÖ

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