Factors Affecting Postoperative Complications and Outcomes of Cervical Spondylotic Myelopathy with Cerebral Palsy: A Retrospective Analysis

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Objective: Cervical surgery in patients with cervical spondylotic myelopathy (CSM) and cerebral palsy (CP) is challenging owing to the complexities of the deformity. We assessed factors affecting postoperative complications and outcomes after CSM surgery in patients with CP.

Methods: Thirty-five consecutive patients with CP and CSM who underwent cervical operations between January 2006 and January 2014 were matched to 35 non-cerebral palsy (NCP) control patients. Postoperative complications and radiologic outcomes were compared between the groups. In the CP group, the Japanese Orthopaedic Association score; Oswestry neck disability index; modified Barthel index; and values for the grip and pinch, Box and Block, and Jebsen-Taylor hand function tests were obtained pre-and postoperatively and compared between those with and without postoperative complications.

Results: Sixteen patients (16/35%) in the CP group and seven (7/35%) in the NCP group (p=0.021) had postoperative complications. Adjacent segment degeneration (p=0.021), postoperative motor weakness (p=0.037), and revisions (p=0.003) were significantly more frequent in the CP group than in the NCP group; however, instrument-related complications were not significantly higher in the CP group (7/35 vs. 5/35, p=0.280). The number of preoperative fixed cervical deformities were significantly higher in CP with postoperative complications (5/16 vs. 1/19, p=0.037). In the CP group, clinical outcomes were almost similar between those with and without postoperative complications.

Conclusion: The occurrence of complications during the follow-up period was high in patients with CP. However, postoperative complications did not significantly affect clinical outcomes.

Key Words: Cervical spondylotic myelopathy · Cerebral palsy · Postoperative complications · Kyphosis · Pseudoarthrosis.
INTRODUCTION

Patients with cerebral palsy (CP) demonstrate involuntary and repetitive neck movements and are more likely to develop cervical spondylotic myelopathy (CSM)\(^1\,^2\,^3\). Because CSM progresses in the 30–40-year age range, individuals with CP often develop various neurological defects later in life. The primary pathologic factors that lead to serious disability include compression against neural elements caused by canal stenosis from excessive spondylotic changes and severe dynamic instability of the spine induced by sustained involuntary movements and malalignment of the cervical spine\(^4\). Typically, conservative treatments are ineffective; patients often require surgical treatment\(^2\). The most important surgical objectives for such patients are adequate decompression of the spinal cord and nerve roots, stabilization of the cervical spine, and alignment correction\(^5\,^6\,^7\). Recent advances in medical technology have led to the development of improved internal fixation methods, which promote stronger initial mechanical stability resulting from anterior plating or posterior screw fixation\(^7\). CSM surgery remains challenging, owing to the complexity of the deformity\(^8\,^9\). Additionally, involuntary neck movements after surgical treatment can worsen neurological symptoms or further complications associated with surgical implants, frequently leading to poor long-term prognoses\(^10\).

We retrospectively investigated the prevalence of various postoperative complications and outcomes resulting from CSM surgery in patients with CP and compared the results with those of non-cerebral palsy (NCP) patients with similar surgical treatments.

MATERIALS AND METHODS

All procedures were performed in accordance with the ethical standards of the Institutional Review Board of Severance Hospital, Yonsei University College of Medicine (approval number: 2016-1447-001). The requirement for informed consent was waived because of the retrospective design of this study.

Patients

Between January 2006 and January 2014, we retrospectively identified 77 patients with CP diagnosed in the Department of Rehabilitation Medicine and Neurosurgery and CSM who underwent surgeries using cervical anterior (anterior cervical disectomy and fusion or anterior cervical corpectomy and fusion), posterior (posterior decompressive laminectomy and lateral mass screw or pedicle screw fixation), or combined (anterior and posterior) approach in our institute due to intractable pain or progressive weakness. The surgical method and implantation levels were determined by evaluating the extent of the correlation between radiological findings and clinical symptoms. Inclusion criteria were as follows: age >18 years with CP and CSM; subaxial cervical spine lesions; intractable pain or progressive weakness and baseline modified Japanese Orthopaedic Association (mJOA) score ≤18; follow-up period >2 years; and complete medical record of clinical and radiologic outcomes in the follow-up period. Exclusion criteria were as follows: infections, autoimmune state, tumors, or other pathologic conditions; follow-up period <2 years; and lack of clinical and radiologic outcome data because of poor image quality or follow-up loss. We excluded 42 patients with a follow-up period <2 years or missing clinical and radiologic outcome data. A total of 35 consecutive patients with CP and CSM who underwent cervical surgeries were ultimately included. Regardless of the type of CP before surgery, patients who did not have complications wore the postoperative orthosis for up to 3 months. Patients with complications that were found within 3 months postoperatively wore the orthosis for an additional 2–3 months depending on their condition.

Patient demographics included cervical spine alignment (C0–2 and C2–7 Cobb angle, C2–7 sagittal vertical axis [SVA], chin brow vertical angle [CBVA], and T1 slope [T1S] minus the C2–7 Cobb angle) and postoperative complications. We paired the participants with 35 age-, sex-, and operation level-matched NCP patients who also underwent cervical surgeries owing to CSM.

Data collection

Relevant patient data, including disease history, physical examination findings, and radiological outcomes, were obtained from medical records.

Radiologic outcomes

Plain radiographs, including anterior-posterior and lateral views, were obtained before and 2 years postoperatively, to assess cervical alignment. The C0–2 Cobb angle (neutral posi-
tion), C2–7 Cobb angle (cervical lordosis, CL) (neutral, flexion, and extension positions), T1S, C2–7 SVA, and CBVA were determined. The C0–2 Cobb angle was measured as the angle between the McGregor line and lower endplate of the C2 vertebra (positive values indicate lordosis and negative values indicate kyphosis between the occiput and C2). CL was defined as the angle between two crossed perpendicular lines extending parallel to the inferior endplate of C2 and C7 on a standing lateral radiograph of the cervical spine. The C2–7 SVA was defined as the distance between a plumb line dropped from the center of C2 (or dens) and the posterosuperior aspect of C7. The T1S was measured as the angle between a horizontal plane and a line parallel to the superior T1 endplate. We also evaluated T1S minus CL (T1S–CL), which is a cervical analog to the pelvic incidence minus lumbar lordosis mismatch.14 The C2–7 range of motion (ROM) was calculated by subtracting the CL on flexion from that on extension. The CBVA was defined as the angle between a line from the brow to the chin to the vertical.

Postoperative complications
Postoperative complications, including neurologic deteriorations, adjacent segment disease, instrument-related complications, and revision frequency, were recorded until the last follow-up (at least 2 years after surgery). We also investigated each complication according to the follow-up period.

Clinical outcomes
All clinical outcome assessments, including the mJOA score (range, 0–21; 0=maximal neurological deficits, 21=no neurological deficits),34 Oswestry neck disability index (NDI) questionnaire (10 questions addressing pain intensity),22 personal care, lifting, reading, headaches, concentration, working, driving, sleeping, and recreation, with higher scores indicating worse outcomes), and the modified Barthel index (MBI; range, 0–100; higher scores indicate greater functional independence), were performed preoperatively and 24 months postoperatively in the CP group. Clinical outcomes were also evaluated by a rehabilitation physician using the following tests: grip and pinch (grasping power), Box and Block (performance while carrying a box and block for 1 minute), and Jebsen-Taylor hand function (time to perform certain tasks, including writing, turning cards, picking up small objects, eating, stacking checkers, and carrying light/heavy cans).

Statistical analysis
Data are reported as means±standard deviations. Group differences in demographic and radiologic parameters were evaluated using the chi-square or Fisher’s exact test, as appropriate. The paired t-test was used to evaluate differences in clinical parameters between before and 2 years after surgery. Statistical analyses were performed using IBM SPSS Statistics for Windows, version 19 (IBM Corp., Armonk, NY, USA). p<0.05 was considered significant.

RESULTS
Patient demographics
The demographic and clinical characteristics of the patients (35 in each group) are presented in Table 1. In the CP group, there were 21 patients with dyskinetic CP, five with spastic CP, and nine with mixed CP. The average height (p=0.006), weight (p<0.001), body mass index (p=0.001), and bone mineral density (p<0.001) were significantly lower, and the mean follow-up duration was significantly longer (p=0.001) in the CP group than in the NCP group. Osteoporosis (p=0.007) and fixed cervical kyphosis (p=0.003) were significantly more frequent in the CP group than in the NCP group. There were no significant differences in age (p=0.610), sex ratio (p=1.000), comorbidities (hypertension, p=0.690; diabetes, p=0.551), and surgical approach (p=0.075) between the groups.

Radiologic outcomes
Preoperative and 2-year postoperative cervical sagittal alignment parameters are summarized according to groups in Table 2. During the preoperative period, the CL on flexion was significantly lower in the CP group than in the NCP group (p=0.008). The T1S (p=0.035) and C2–7 ROM (p=0.011) were significantly higher in the CP group than in the NCP group; however, the C2–7 SVA (p=0.245), CBVA (p=0.893), and T1S minus the C2–C7 Cobb angle (p=0.180) did not differ significantly. In the postoperative period, the C2–7 Cobb angle on flexion was the only parameter that significantly differed between the groups (p=0.019). There was no significant difference in radiologic parameters between the groups before and after surgery.
Postoperative complications

Postoperative complications are summarized in Table 3. There were 16 patients (45.7%) with postoperative complications in the CP group and seven (20.0%) in the NCP group (p=0.021). Although the number of overall neurologic deterioration events did not significantly differ between the groups (p=0.324), motor weakness was significantly more frequent in the CP group than in the NCP group (p=0.037). The groups did not significantly differ in overall instrument-related complications, including pseudoarthrosis, screw loosening, and screw fracture (p=0.280); however, adjacent segment degeneration (ASD; p=0.018) and revision events (p=0.003) were significantly more frequent in the CP group than in the NCP group. Among those with postoperative complications, only the incidence of neurological deterioration significantly differed between groups (p=0.027).

In addition, the occurrence of neurologic deterioration was significantly high in the CP group between 2 and 5 years after
Clinical and radiologic outcomes in the CP group

A summary of demographics and clinical outcomes in the CP group is clustered and presented according to the presence or absence of complications in Tables 4 and 5. In the complication group, the number of fixed cervical deformity was significantly higher than in those without complications ($p=0.037$). The types of CP were not significantly different between the groups with and without complications ($p=0.785$). The pre-($p=0.049$) and postoperative JOA scores ($p=0.042$) were significantly less in those with postoperative complications than in those without postoperative complications; however, the JOA recovery rate did not significantly differ ($p=0.892$). No other clinical outcomes significantly differed between those with and without postoperative complications. The preoperative and postoperative radiologic parameters also did not differ significantly (Table 6).

### Table 3. Distribution of postoperative complications

|                     | CP (n=35) | NCP (n=35) | $p$-value |
|---------------------|-----------|------------|-----------|
| Overall complications | 16 (45.7) | 7 (20.0) | 0.021*    |
| Neurologic deteriorations | 7 (20.0) | 4 (11.5) | 0.324     |
| Pain, sensory change without weakness | 1 (2.8) | 3 (8.5) | 0.293     |
| Motor weakness | 6 (17.1) | 1 (2.8) | 0.037*    |
| ASD | 7 (20.0) | 1 (2.8) | 0.018*    |
| Instruments related complications | 7 (20.0) | 5 (14.3) | 0.280     |
| Pseudoarthrosis, non-fusion | 2 (5.6) | 2 (5.6) | 1.000     |
| Screw loosening or fracture | 5 (14.3) | 3 (8.5) | 0.450     |
| Revision | 6 (17.1) | 0 (0.0) | 0.003*    |
| ASD | 3 (8.5) | 0 (0.0) | -         |
| Motor weakness | 6 (17.1) | 1 (2.8) | -         |
| Instrument related complications | 1 (2.8) | 0 (0.0) | -         |

### Table 4. Baseline characteristics according to the presence or absence of complications in the cerebral palsy group

|                     | YC (n=16) | NC (n=19) | $p$-value |
|---------------------|-----------|-----------|-----------|
| Age (years) | 37.95±7.06 | 42.80±11.83 | 0.140     |
| Sex, male | 12 | 9 | 0.096     |
| Height (cm) | 159.8±7.4 | 162.1±10.1 | 0.447     |
| Weight (Kg) | 55.5±14.3 | 51.5±8.3 | 0.337     |
| BMI (Kg/m$^2$) | 21.7±5.3 | 19.6±2.7 | 0.176     |
| BMD | -1.43±1.01 | -1.10±1.13 | 0.386     |
| Osteoporosis | 2 | 3 | 0.781     |
| HTN | 1 | 2 | 0.649     |
| DM | 0 | 1 | 0.264     |
| Type of CP | 0.785 | 9 | 12 | 0.785 |
| Dyskinetic, athetoid | 9 | 12 | 0.785 |
| Spastic, diplegic | 3 | 2 | 0.785 |
| Mixed | 4 | 5 | 0.785 |
| Preoperative history | 1 | 0 | 0.206     |
| Fixed cervical deformity | 5 | 1 | 0.037*    |
| Perioperative Botox injection | 12 | 16 | 0.498     |

Values are presented as mean±standard deviation or number. *Indicates statistical significance ($p<0.05$). YC : yes complication, NC : no complication, BMI : body mass index, BMD : bone mineral density, HTN : hypertension, DM : diabetes mellitus, CP : cerebral palsy.
Surgical Outcomes of CSM with CP

| Clinical outcome | YC (n=16) | NC (n=19) | p-value |
|------------------|-----------|-----------|---------|
| Pre-op JOA      | 11.38±4.60 | 13.95±2.76 | 0.049* |
| Post-op JOA score | 13.50±3.67 | 15.74±2.58 | 0.042* |
| JOA recovery rate (%)* | 25.30±21.07 | 26.32±22.61 | 0.892 |
| Pre-op NDI      | 39.50±20.64 | 37.95±17.97 | 0.813 |
| Post-op NDI     | 26.75±16.96 | 28.05±16.71 | 0.821 |
| Pre-op MBI      | 42.94±23.22 | 57.53±24.89 | 0.084 |
| Post-op MBI     | 62.19±19.49 | 72.11±20.50 | 0.154 |

Grip and pinch test (kg)

| Pre-op grip        | 18.01±9.41 | 18.17±7.99 | 0.956 |
| Post-op grip       | 15.06±9.23 | 16.11±8.99 | 0.736 |

Post-op tip

| 2.62±1.85 | 2.74±1.78 | 0.842 |
| 2.57±1.70 | 2.22±1.47 | 0.515 |

Pre-op lateral

| 3.88±1.85 | 4.15±2.18 | 0.696 |
| 3.79±1.99 | 3.62±1.88 | 0.795 |

Pre-op palmar

| 3.42±1.66 | 3.52±2.18 | 0.888 |
| 4.01±1.58 | 3.68±2.56 | 0.665 |

Box & Block test

| Pre-op            | 26.38±16.09 | 23.57±15.98 | 0.734 |
| Post-op           | 23.50±12.89 | 24.69±16.40 | 0.689 |

Jebsen-Taylor hand function test (seconds)

| Writing, pre-op  | 54.67±48.68 | 45.81±27.70 | 0.505 |
| Writing, post-op | 43.61±29.31 | 39.36±25.81 | 0.652 |
| Card turning, pre-op | 19.10±12.30 | 17.02±11.16 | 0.604 |
| Card turning, post-op | 14.28±8.42 | 13.13±9.02 | 0.701 |

Moving light objects, pre-op

| 12.35±10.82 | 12.74±11.83 | 0.919 |

Moving light objects, post-op

| 9.44±3.96 | 9.24±6.55 | 0.917 |

Moving heavy objects, pre-op

| 9.90±7.37 | 10.43±7.75 | 0.839 |

Moving heavy objects, post-op

| 9.40±7.06 | 9.02±6.69 | 0.873 |

Stacking checkers, pre-op

| 22.09±14.37 | 19.43±13.04 | 0.575 |

Stacking checkers, post-op

| 19.22±14.93 | 15.85±13.80 | 0.493 |

Simulated feeding, pre-op

| 23.07±14.73 | 27.14±18.72 | 0.487 |

Simulated feeding, post-op

| 23.29±16.75 | 21.36±14.72 | 0.719 |

Placing small objects, pre-op

| 36.15±42.43 | 24.85±18.50 | 0.301 |

Placing small objects, post-op

| 26.20±19.94 | 18.46±11.15 | 0.158 |

Table 5. Clinical outcomes according to the presence or absence of complications in the cerebral palsy group

Table 6. Radiologic outcomes according to the presence or absence of complications in the cerebral palsy group

| YC (n=16) | NC (n=19) | p-value |
|-----------|-----------|---------|
| C0–C2 angle, neutral | 43.76±20.33 | 38.01±13.61 | 0.326 |
| C2–C7 angle, neutral | 2.45±24.48 | 5.77±24.45 | 0.691 |
| C2–C7 angle, flexion | -31.68±15.46 | -30.44±19.08 | 0.836 |
| C2–C7 angle, extension | 19.41±22.37 | 24.78±21.22 | 0.472 |
| T1 slope | 28.15±13.56 | 24.88±14.14 | 0.493 |
| T1 slope–C27 angle | 25.70±18.21 | 19.11±16.45 | 0.269 |
| C2–7 SVA (cm) | 28.29±26.97 | 24.18±19.01 | 0.601 |
| C2–7 ROM | 51.09±29.12 | 55.23±20.02 | 0.624 |
| CBVA | -3.93±14.00 | -0.08±17.81 | 0.488 |

Post-op radiologic parameters

Values are presented as mean±standard deviation. YC : yes complication, NC : no complication, Pre-op : preoperative, SVA : sagittal vertical axis, ROM : range of motion, CBVA : chin brow vertical angle, Post-op : postoperative

DISCUSSION

CP is a movement disorder caused by a non-progressive abnormality appearing in the perinatal period. Lack of control and/or spasticity of truncal muscles frequently lead to spinal degenerative diseases or deformities, such as CSM and scoliosis. Early degeneration onset and malalignment of the spine have been reported in patients with CSM and CP. Owing to repetitive and unusual movements associated with this condition, the pathophysiology of CSM accelerates in patients with CP, and inexplicable changes or deterioration of neurological function requiring surgical treatments can occur. CSM surgery for patients with CP remains challenging because of peri-operative instrumentation failure caused by the patient’s repetitive involuntary neck movements, complicated cervical spine deformities, and comorbidities. Several studies have reported
postoperative complications in patients with CSM without CP or scoliosis with CP\textsuperscript{24,33}. However, there exist only a few reports regarding postoperative complications in patients with CSM and CP. Current and accurate information regarding major complications following cervical surgeries for significant CSM in patients with CP is important for weighing the costs and benefits of corrective surgery for this population.

In previous studies, complication rates in NCP patients who underwent cervical surgery due to CSM were 9–30\%\textsuperscript{17,23,26,31}. Yaszay et al.\textsuperscript{20} reported a rate of 36\% for postoperative major complications (e.g., wound infections, pulmonary issues, instrument-related complications) in patients with CP with $\geq 2$ years of follow-up after spinal surgery, with a spine-related reoperation rate of 14.0\%. Samdani et al.\textsuperscript{24} reported a 39\% complication rate in 127 patients with CP who underwent spinal surgery. In our study, the overall postoperative complication rate for CSM surgery was significantly higher in patients with CP than in NCP patients. Additionally, there were significant group differences in terms of postoperative motor weakness, ASD, and revision events. There may be several reasons for these differences. Excessive involuntary neck motion induced by CP, in association with vertebral slippage anteriorly or posteriorly after cervical fusion, could exaggerate segmental and kyphotic instability. Moreover, these involuntary movements, as well as the abnormal cervical spinal alignment due to an imbalance in cervical muscle tone and compression of neural elements by severe cervical spondylotic changes, can cause rapid and additional degenerative changes and subsequent neurological deficits\textsuperscript{12}. In our study, six out of 35 patients in the CP group developed motor weakness. In three patients, motor weakness was caused by ASD, and in the other three patients, it was caused by progressive kyphosis. Some studies have reported that cervical ASD occurs in approximately 3\% of NCP patients. However, Azuma reported that ASD occurred in 30\% of patients with CP with a follow-up period $>10$ years after cervical fusion for CSM. Likewise, in our study, ASD occurrence was significantly more common in the CP group than in the NCP group. However, instrument-related complications, such as pseudoarthrosis and screw fractures or loosening, did not significantly differ between CP and NCP groups. We hypothesize that improvement in surgical instrument and principles, leading to strong initial mechanical stability as a result of anterior plating or posterior screw fixation, contributed to these favorable results. Additionally, our results indicate that complications such as instrument failure or non-fusion may not be considered as strong as once thought when considering cervical surgery for CSM in patients with CP.

In our study, risk factors for postoperative complications in CSM with CP included fixed cervical deformity. This result was likely because among patients with CP, surgical invasiveness, need for surgical release, and utilization of osteotomies significantly increased in those with fixed cervical kyphosis than in those with semi-rigid or flexible kyphosis. In patients with CP, fixed cervical kyphosis is often accompanied by severe degenerative changes; thus, surgical treatment of such patients is challenging and requires invasive surgery to correct and stabilize the deformity while decompressing neural elements and restoring sagittal alignment\textsuperscript{11,13,15}. Hence, patients with CSM and CP tend to undergo staged operations or combined approaches. Scheer et al.\textsuperscript{25} reported significant differences in complication rates for different approaches (anterior approach, 27.3\%; posterior approach, 68.4\%; combined approach, 79.3\%). In the present study, many patients in the CP group required posterior or combined approaches (posterior with an anterior approach or anterior with a posterior approach), but there were no significant differences in surgical approaches between the CP and NCP groups. This is one of the reasons for the high postoperative complication rate.

Apart from the incidence of complications, the clinical outcome is another important factor used to assess the postoperative prognosis. Demura et al.\textsuperscript{19} showed that the 10-year Barthel index and JOA score after cervical surgery significantly improved (by 36\% and 31\%, respectively) in 14 patients with CSM and CP. Additionally, Watanabe et al.\textsuperscript{26} reported that the JOA score improved from 8.3 points preoperatively to 10.9 points by the final follow-up (mean, 58 months) in patients with cervical disorder-associated CP who underwent posterior or fusion using cervical pedicle screws; however, one-third of the patients were unable to walk at the final follow-up owing to myelopathy progression. Other studies have shown that postoperative clinical outcomes for patients with cervical myelopathy and CP are relatively poor owing to unfavorable conditions (severe cervical spinal deformity and intervertebral instability associated with involuntary neck movements\textsuperscript{27,32}). We compared patients with CSM and CP, according to the presence or absence of postoperative complications, to determine the trends in outcomes before and after surgery. We found that JOA scores before and after surgery significantly differed

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between those with and without complications; however, the JOA recovery rate, as well as NDI and MBI scores, did not significantly differ. Some studies have reported the imitations of JOA and Barthel index for the evaluation of patients with involuntary movements, as in CP\textsuperscript{4,9}. Therefore, additional indicators were needed for an accurate evaluation of this patient group\textsuperscript{10}. Accordingly, we assessed the CP group using the functional indicators used in the field of rehabilitation medicine specializing in the evaluation of upper limb function for multilateral assessments (grip and pinch, Box and Block, and Jebsen-Taylor hand function tests). However, no significant differences were found between those with and without postoperative complications. Based on our results, it is not necessary to delay or cancel surgical treatments for patients with CSM and CP due to vague concerns of perioperative complication. Instead, careful consideration of the surgical methods and levels is required, and multidisciplinary consultations should be considered in the treatment of these patients.

Our study has a few key limitations. First, this was a retrospective, uncontrolled drawback study with small sample size. Moreover, the average follow-up period was longer in the CP group than in the NCP group, which was likely to cause selection bias in terms of the complication rate. Second, the rehabilitation medicine clinical scales were only applied to the CP group, and we did not conduct a multifaceted comparison of the clinical courses between the CP and NCP groups. Third, the instrumented level was included while matching the control group (NCP), but there was a fundamental difference in the surgical strategies between the CP and NCP groups. Additionally, the optimal prevention strategy for postoperative complications remains unclear; thus, additional studies are required. Third, the statistical analysis was limited, as this study was designed using a simple matching method for age, sex, and surgical range.

CONCLUSION

The incidence of overall complications, particularly ASD, postoperative motor weakness, and revision events, was higher in the CP group than in the NCP group during the follow-up period (at least 2 years). Risk factors for postoperative complications of CSM with CP included fixed cervical deformity. Instrument-related complications were not significantly different between groups. In the CP group, there were no differences in clinical outcomes between those with and without postoperative complications. These results suggest that postoperative complications may not be considered as a concern when deciding on surgical treatment for CSM in patients with CP.

CONFLICTS OF INTEREST

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

INFORMED CONSENT

This type of study does not require informed consent.

AUTHOR CONTRIBUTIONS

Conceptualization : HCK, YH, SRC
Data curation : HCK, JHH, SRC
Formal analysis : HCK, YH, HJ
Methodology : HCK, HJ, YHJ, SP, SBA
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