

21 **Keywords:** clinical pathways, electronic health records(EHR), matching rates, evidence-based
22 approach, total laparoscopic hysterectomy(TLH), rotator cuff tears(RCTs)
23

24 **Abstract**

25 **Objective**

26 A clinical pathway is one of the tools used to support clinical decision making that provides a
27 standardized care process in a specific context. The objective of this research was to develop a method
28 for building data-driven clinical pathways using electronic health record data.

29

30 **Materials and methods**

31 We proposed a matching rate-based clinical pathway mining algorithm that produces the optimal set of
32 clinical orders for each clinical stage by employing matching rates. To validate the approach, we utilized
33 two different datasets of deidentified inpatient records directly related to *total laparoscopic*
34 *hysterectomy* (TLH) and *rotator cuff tears* (RCTs) from a hospital in South Korea. The derived data-
35 driven clinical pathways were evaluated with knowledge-based models by health professionals using a
36 delta analysis.

37

38 **Results**

39 Two different data-driven clinical pathways, i.e., TLH and RCTs, were produced by applying the
40 matching rate-based clinical pathway mining algorithm. We identified that there were significant
41 differences in clinical orders between the data-driven and knowledge-based models. Additionally, the
42 data-driven clinical pathways based on our algorithm outperformed the models by clinical experts, with
43 average matching rates of 82.02% and 79.66%, respectively.

44

45 **Conclusion**

46 The proposed algorithm will be helpful for supporting clinical decisions and directly applicable in
47 medical practices.

48

49 **1. Introduction**

50 A clinical pathway is a tool that delivers structured clinical services on the basis of evidence-based
51 healthcare in a specified medical context (e.g., for diseases, diagnostics, and surgeries) [1-8]. It has been
52 developed with the aim of standardizing and optimizing care processes to minimize the undesired
53 practice variability and manage clinical outcomes, e.g., length of stays or rehospitalization [1-8]. As
54 such, the application of clinical pathways has received attention since it shortens the length of hospital
55 stays, lowers costs, reduces complications and lowers mortality [9-12].

56 Typically, a clinical pathway is organized in a day-by-day format, composed of clinical orders that
57 contain clinical services (e.g., prescriptions and treatments) from medical practitioners [4,13].
58 Additionally, each clinical pathway has a specified length of hospital stay, and clinical stages are defined
59 such as *regular*, *preoperation*, *postoperation*, and *discharge* [4,13]. Therefore, solid theoretical
60 backgrounds are required to develop clinical pathways, and typically, most medical sites have developed
61 clinical pathways based on the knowledge of domain experts, e.g., health professionals [14,15].

62 The process of developing knowledge-based clinical pathways by discussing with medical professionals
63 is ideal, and clinical pathways have been developed on the basis of this approach for frequently
64 occurring significant diseases [14,15]. However, it was challenging to develop clinical pathways for all
65 clinical contexts using the knowledge-based approach due to the following two reasons: (i) time and
66 efforts are limited since health professionals engaged in the development of clinical pathways are
67 overloaded with medical activities, and (ii) healthcare processes are dynamic and complicated since
68 clinical operations, medicines, and therapies are developed continuously.

69 As an alternative approach and support for the knowledge-based approach, data-driven methods have
70 been developed using the data from electronic health records (EHR) [4,7,16-23]. Specifically, data-
71 centric techniques, e.g., data mining and process mining, have been utilized to develop more realistic
72 clinical pathways [4,7,16-23].

73 These approaches have resolved the limitations of knowledge-based clinical pathways addressed above,

74 however, unfortunately, it still has a couple of challenges to apply them as follows immediately: (i) they
75 only focus on deriving a coarse-grained clinical pathway (i.e., at the activity level) with the traditional
76 process mining approaches, in other words, they do not provide any detailed the steps (i.e., clinical
77 orders) for a specific timeframe [16,20,23], (ii) they focus on finding clinical pathway patterns or a
78 summarized model instead of providing how to develop the standardized clinical order set for a specific
79 surgery or diagnosis [4,7,21,38].

80 This paper proposes an approach to produce a realistic clinical pathway at the order level, i.e., a
81 matching rate-based clinical pathway mining algorithm. The proposed approach starts with our prior
82 research suggesting a method to compare clinical pathways and the relevant log data with a quantitative
83 approach, i.e., matching rates (see the details in Section 3) [13]. The method aims to derive the optimal
84 set of clinical orders for specific timeframes, i.e., clinical pathways, that maximizes the matching rate
85 using patient data collected in EHR. Specifically, we utilized two different sets of deidentified inpatient
86 records directly related to *total laparoscopic hysterectomy* (TLH) and *rotator cuff tears* (RCTs)
87 collected in a tertiary hospital in South Korea. Based on the proposed approach, we derived two different
88 clinical pathways for each clinical context and compared them with the knowledge-based models
89 created by health professionals to validate our work.

90

91 **2. Related works**

92 The primary discipline associated with our work, i.e., the development of data-driven clinical pathways,
93 is process mining. It aims at deriving process-related knowledgeable insights from event logs and has
94 been widely applied in a healthcare environment [26,29]. The use of process mining in healthcare has
95 been more focused on process discovery, and there have been approaches for automatically extracting
96 clinical workflow process models in different medical fields [29]. Mans et al. [18] presented the
97 discovered spaghetti-like process model for the gynecological oncology healthcare process, while
98 Rebuge and Ferreira [30] proposed a method to generate emergency process models using process

99 mining with clustering techniques. Also, there have already been attempted to discover clinical process
100 models for outpatients, inpatients, and surgery [29].

101 Similar to these works, some researchers have strived to develop data-driven clinical pathways, with
102 the focus on identifying frequent patterns and process models, using process discovery techniques
103 pertaining to process mining [16,22,23]. Lakshmanan et al. [22] proposed a method for mining a clinical
104 pathway mining approach based on clinical outcomes by merging process mining with frequent pattern
105 mining. Also, Xu et al. [16] presented a series of steps to generate topic-based activity clusters using
106 Latent Dirichlet Allocation (LDA) and derive a process model using fuzzy mining. These approaches
107 have contributed to developing a high-level abstraction clinical pathway, i.e., an activity-level process
108 model. However, these approaches do not include the details of how to make clinical orders required
109 by a specific stage or date; thus, a low-level abstraction clinical pathway, i.e., the order-level, is required
110 for practical use.

111 In this regard, to overcome this limitation, several approaches have devoted to creating the order-level
112 clinical pathway using process mining [4,7,21,38]. Huang et al. [4] suggested a mining approach to
113 derive summarized clinical pathways given minimum support threshold and event logs, while Iwata et
114 al. [21] suggested the similarity-based visualization approach that provides a compressed model based
115 on the probabilistic threshold from users. Huang et al. [7] focused on finding clinical pathway patterns,
116 not the homogenized model. Also, Huang et al. [38] proposed a method to detect anomalies in clinical
117 pathway patterns. To recap, those methods were effective in analyzing clinical pathways such as
118 identifying patterns and deriving summarized information, but they had a limitation that is not
119 applicable to find a more standardized clinical pathway presented in this study.

120 The other related discipline to our research is healthcare data mining, which has more focused on
121 discovering clinical order sets necessitated for clinical decision support tools [31-35]. More in detail,
122 they developed several approaches to provide the patient-personalized clinical order sets using Hidden
123 Markov Model [31], K-means clustering [32,33], recommendation systems [34], and frequent itemset
124 mining and association rule mining [35]. These approaches are quite similar to the data-driven clinical

125 pathway mining in that both approaches produce a series of order sets as output. However, they devote
126 to identifying patient-personalized clinical order sets with the aim of reducing cognitive click costs of
127 experts and providing personalized healthcare [34,35], while clinical pathway mining focuses on
128 deriving comprehensive order sets for a specific clinical context in an unfavorable economic scenario
129 with the aims of minimizing variations of the clinical results. Thus, it is required to suggest a different
130 method for clinical pathway development.

131

132 **3. Materials and Methods**

133 **3.1. Clinical pathways**

134 This section introduces the features and overall structure of clinical pathways developed in this research.
135 To determine the format of clinical pathways, we have employed the existing works of literature and
136 discussion with health professionals. In this regard, we aimed at developing the most standardized
137 clinical order set for a specific surgery or diagnosis; in other words, our models did not cover the
138 diversity, i.e., conditional branching in models, caused from the comorbidities, e.g., characteristics or
139 history of patients. To this end, an alternative method was considered as building branch CPs by
140 distinguishing from the parent CP, if needed to create the model for a specific comorbidity. In this regard,
141 we utilized a simple two-step algorithm [37]; (i) the statistical analysis is performed to identify the
142 relationship between patient characteristics and clinical outcomes (e.g., length of stays, readmission
143 rates, and matching rates) and (ii) if exists, the proposed approach in this study is applied to two patient
144 cohorts based on patient characteristics (i.e., holding and non-holding), and then a branch CP is defined
145 in the case that the derived two order sets have a clear difference.

146 Regarding the structure of the clinical pathways, it has the specified length, i.e., duration, for a particular
147 surgery or diagnosis, and a set of clinical orders is constructed for each day, i.e., a day-by-day format.
148 Also, relative dates based on surgery dates (e.g., *OP day*, *1 day before*, and *1 day after*) were utilized
149 instead of the actual dates on admission (e.g., *day 1* and *day 2*) since the difference in medical orders is

150 clear based on the time of the surgery. Furthermore, clinical pathways hold clinical stages such as
151 *regular, pre-operation, post-operation, and discharge* for each day, and only unique orders can be
152 included for a specific stage and date. Thus, it does not take into account that a single order appears
153 multiple times in a particular stage and date. This is because historical data may enclose biased and
154 suboptimal behaviors including overutilization of orders [36]. Lastly, with the same reason, only
155 medication and test orders are included in clinical pathways based on the opinion of health professionals
156 that they do not cause this issue.

157

158 **3.2. Data**

159 Two different sets of deidentified inpatient records related to total laparoscopic hysterectomy and rotator
160 cuff tears were collected from the Seoul National University Bundang Hospital, a tertiary hospital in
161 South Korea. TLH is one of the surgeries performed in obstetrics and gynecology [24], and we collected
162 records from 520 inpatients who received the surgery from January 2012 to May 2014. RCT is a
163 common disease managed in orthopedics [25], and data from 360 inpatients between June 2014 and
164 2015 were extracted to develop a data-driven clinical pathway.

165 These records included patient information, hospitalization, operation, diagnosis, and orders.
166 Specifically, concerning the clinical orders, i.e., basic units of clinical pathways, 18115 and 14862
167 events were extracted, respectively. Additionally, each event included order attributes such as stages
168 (e.g., preoperation, operation, and postoperation) and types (e.g., medicines and tests). Table 1 provides
169 detailed information about the collected data.

170 The present study was approved (IRB No. B-1609/361-105) by the Institutional Review Board of the
171 Seoul National University Bundang Hospital, which waived patients' informed consent. All deidentified
172 EHR data were then provided to the researchers for this study.

173 **Table 1. Types and attributes of the collected data.**

Type	Attribute
Patient Information	Patient ID, Age, Sex, Drinking, Smoking, Allergy, Disease history, Operation history, Drug history, Family disease history, Diabetes, Hypertension, Hyperlipidaemia, Cardiovascular, Cerebrovascular
Hospitalization	Hospitalization ID, Patient ID, Admission date, Discharge date, Admission type, Discharge schedule type, Assigned physician ID, Department code, Department name
Operation	Operation ID, Patient ID, Hospitalization ID, Operation date, Operation code, Operation name
Diagnosis	Diagnosis ID, Patient ID, Hospitalization ID, Diagnosis code, Diagnosis code classification, Diagnosis name, Physician ID, Department code, Department name, Diagnosis date
Order	Order ID, Patient ID, Hospitalization ID, Order code, Order name, Order date, PRN status, Order stage, Order type, Order interruption classification code, Order interruption date

174 Hospitalization ID: a unique ID for identification of inpatients.

175

176 **3.3. Data re-engineering**

177 In a clinical pathway, the same orders can be repeatedly involved in different dates or stages; thus, a
 178 single order can be published multiple times according to its usage. Thus, to distinguish the same
 179 clinical order in building a clinical pathway, we performed a data re-engineering approach. More in
 180 detail, we determined the clinical orders as the combination of order names, relative dates based on
 181 surgery dates, and order types (i.e., *'order_date_type'*). Here, regarding the relative dates based on
 182 surgery dates, the day of the operation was calculated as 0, the next day as 1, and the one day before
 183 surgery as -1, while the order types were defined as regular, pre-operation, post-operation, operation,
 184 and discharge. Thus, if order A was used as a normal order on the day of surgery, the code would be
 185 expressed as *'A_0_regular'*.

186

187 3.4. Order-level matching rate

188 Prior to explain our algorithm for deriving clinical pathways, we first explain our prior research that
189 compares clinical pathways and the relevant log data with a quantitative approach [13]. More
190 specifically, it generates a numerical value of comparison between clinical orders from clinical
191 pathways and logs. The order-level matching rate has employed conformance checking in process
192 mining discipline, and it signifies the extent to which the log is related to the possible behaviors in the
193 process model. Here, we have defined the matching rate by replacing with the clinical pathway. Formula
194 (1) provides the proposed matching rate.

195

$$196 \quad \text{Matching rate} = \frac{1}{2} \left(1 - \frac{M_{CP}}{N_{CP}} \right) + \frac{1}{2} \left(1 - \frac{R_{Log}}{N_{Log}} \right) \quad (1) [13]$$

197 - M_{CP} : The number of orders included in the clinical pathway but not shown in the log data

198 - N_{CP} : The number of orders included in the clinical pathway

199 - R_{Log} : The number of orders included in the log data but not shown in the clinical pathway

200 - N_{Log} : The number of orders included in the log data

201

202 As shown in this formula, the matching rate is composed of the application rate of orders in the clinical
203 pathway (i.e., $\frac{1}{2} \left(1 - \frac{M_{CP}}{N_{CP}} \right)$) and the matched ratio of orders in the log data (i.e., $\frac{1}{2} \left(1 - \frac{R_{Log}}{N_{Log}} \right)$). Thus, it
204 covers both how the orders included in the clinical pathway are applied to the patients and how the
205 orders used by the patients are different from the clinical pathway.

206 Table 2 provides an example of how to measure the matching rate with the clinical pathway and log
207 data. In this example, the clinical pathway is composed of 12 clinical orders (i.e., T1, T2, T3, Te1, Te2,
208 Te3, M1, M2, M3, I1, I2, and I3), while the different order codes for four patients are contained. As
209 provided in the Table, P1's orders are the same as those of the clinical pathway. Thus, M_{CP} and R_{Log} are
210 0, and the matching rate becomes 1.00. In the case of P2, compared to those in the clinical pathway,

211 some orders, e.g., Te3, I2, and I3, are missing (i.e., expressed as “-”). As such, M_{CP} and N_{Log} are 3 and
 212 9, respectively, and the matching rate becomes 0.88. Compared to P2, the orders for P3 include some
 213 orders not defined in the clinical pathway, e.g., T4, I4, and I5. Thus, N_{Log} and R_{Log} are 15 and 3,
 214 respectively, and these values decrease the matching rate to 0.90. Lastly, P4 lacks some of the required
 215 orders and the addition of orders not defined in the clinical pathway. As such, the matching rate becomes
 216 the lowest value among the four patients.

217

218 **Table 2. An example of how to measure the matching rate.**

<i>CP</i>	<i>T1</i>	<i>T2</i>	<i>T3</i>	<i>Te1</i>	<i>Te2</i>	<i>Te3</i>	<i>M1</i>	<i>M2</i>	<i>M3</i>	<i>I1</i>	<i>I2</i>	<i>I3</i>	<i>N_{CP}</i>	<i>M_{CP}</i>	<i>N_{Log}</i>	<i>R_{Log}</i>	<i>Matching rate</i>			
P1	T1	T2	T3	Te1	Te2	Te3	M1	M2	M3	I1	I2	I3	12	0	12	0	1.00			
P2	T1	T2	T3	Te1	Te2	-	M1	M2	M3	I1	-	-	12	3	9	0	0.88			
P3	T1	T2	T3	T4	Te1	Te2	Te3	M1	M2	M3	I1	I2	I3	I4	I5	12	0	15	3	0.90
P4	T1	-	T3	T4	Te1	Te2	-	M1	-	M3	I1	I2	-	I4		12	4	10	2	0.73

219

220 3.5. Matching rate-based clinical pathway mining algorithm

221 The matching rate-based clinical pathway mining algorithm is outlined in Figure 1. Our algorithm takes
 222 log data (i.e., L) as an input and produces the clinical pathway (i.e., CP) and its matching rate (i.e., m).
 223 In the proposed algorithm, the first step is to define and initialize the required variables (e.g., m) and
 224 sets (e.g., CP , O_L , and AR_O). Then, as shown in lines 3-7, a unique clinical order set (i.e., O_L) is prepared
 225 and included in the log data. Next, we calculate the application rates for clinical orders involved in O_L .
 226 In the following steps (i.e., line 9-12), the application rate is defined as the number of applied clinical
 227 orders (i.e., $\sum_{0 < k \leq p} \sum_{0 < l \leq n} \begin{cases} 1 & \text{if } o_{k,l} = o_j \\ 0 & \text{otherwise} \end{cases}$) divided by the number of patients (i.e., p). As a result of
 228 this step, the list of application rates for each clinical order (i.e., AR_O) is prepared. Next, all clinical
 229 orders are sorted in descending order based on their application rates, and O_L' is generated (line 13).
 230 Lines 14-22 provide an iterative approach to find the optimal clinical pathway. In the sorted clinical
 231 order set O_L' , we first select a clinical order that has the highest application rate and is included in the
 232 clinical pathway set CP . If the application rates of different orders are the same, then the order is

233 randomly selected. Then, we measure the average value of matching rates (i.e., m_i) using
234 $\frac{1}{2} \left(1 - \frac{\sum_{0 < k \leq p} M_{k,CP}}{\sum_{0 < k \leq p} N_{k,CP}} \right) + \frac{1}{2} \left(1 - \frac{\sum_{0 < k \leq p} R_{k,Log}}{\sum_{0 < k \leq p} N_{k,Log}} \right)$. Here, let $N_{k,CP}$ denote the number of clinical orders in the
235 clinical pathway for patient k . $M_{k,CP}$, $N_{k,Log}$, and $R_{k,Log}$ are defined in a similar fashion, e.g., $R_{k,Log}$ is the
236 number of orders included in the data of patient k but is not shown in the clinical pathway. After that,
237 the measured m_i is compared with the current maximum value m . If m_i is larger than m , the selected
238 order is unchanged in CP . In the opposite case, however, it is removed from the clinical pathway. By
239 conducting this step iteratively, the proper set of clinical orders is determined and becomes the optimal
240 clinical pathway.

241

Algorithm 1 Matching rate-based clinical pathway mining

Input

A log data L

Output

A clinical pathway (i.e., a list of clinical orders) CP

A matching rate of the clinical pathway m

1. $CP, O_L, AR_O \leftarrow \{\}$
2. $m \leftarrow 0$
3. **for all** o_i in L **do**
4. **if** o_i does not exist in O_L **then**
5. $O_L \leftarrow O_L \cup o_i$
6. **end if**
7. **end for**
8. Let n be the number of unique clinical orders in O_L and p be the total number of patients in L
9. **for all** o_j in O_L **do**
10. Calculate the application rate $ar_j = \frac{\sum_{0 < k \leq p} \sum_{0 < l \leq n} \begin{cases} 1 & \text{if } o_{k,l} = o_j \\ 0 & \text{otherwise} \end{cases}}{p}$
11. $AR_O \leftarrow AR_O \cup ar_j$
12. **end for**
13. Sort O_L by AR_O in the descending order to get the $O_L' = \{o_1', o_2', \dots, o_n'\}$
14. **for all** o_j' in O_L' **do**
15. $CP \leftarrow CP \cup o_j'$
16. Calculate the average of matching rates $m_i = \frac{1}{2} \left(1 - \frac{\sum_{0 < k \leq p} M_{k,CP}}{\sum_{0 < k \leq p} N_{k,CP}} \right) + \frac{1}{2} \left(1 - \frac{\sum_{0 < k \leq p} R_{k,Log}}{\sum_{0 < k \leq p} N_{k,Log}} \right)$
17. **if** $m_i > m$ **then**
18. $m \leftarrow m_i$
19. **else**
20. $CP \leftarrow CP \otimes o_j'$
21. **end if**
22. **end for**

23. *return CP, m*

242

243 **Figure 1. Matching rate-based clinical pathway mining algorithm**

244 We provide the procedure of the suggested matching rate-based clinical pathway mining algorithm with
245 the running example in Table 2. In the table, the log data and the clinical orders of four patients are
246 included. Based on the log, we first identify the set of unique clinical orders, i.e., $O_L = \{T1, T2, T3, T4,$
247 $Te1, Te2, Te3, M1, M2, M3, I1, I2, I3, I4, I5\}$. Then, the application rates for clinical orders are
248 calculated, e.g., the values of T1 and T2 are 1.0 and 0.75, respectively. Among them, a clinical order
249 with the highest application rate, i.e., T1, is added into the clinical pathway CP , and the average
250 matching rate is calculated as 0.55. In the second iteration, T3 is added in the clinical pathway, i.e., CP
251 $= \{T1, T3\}$. Then, the average matching rate becomes higher than the value from the first iteration. By
252 conducting this process continuously, we can obtain the highest matching rate as 0.88 and the relevant
253 clinical pathway, i.e., $CP = \{T1, T3, Te1, Te2, M1, M3, I1, T2, Te3, M2, I2, T4\}$.

254

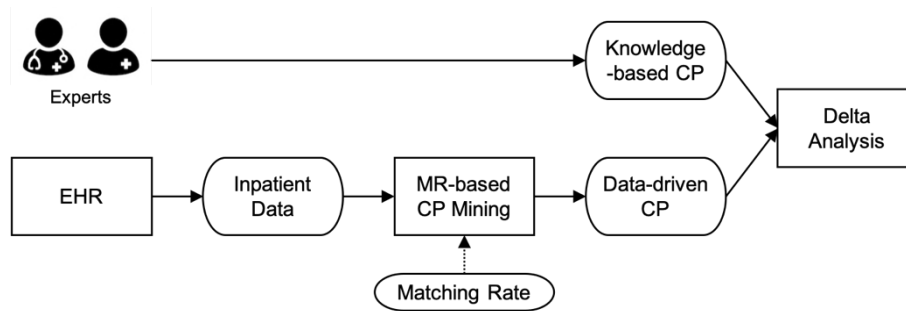
255 **3.6. Evaluation**

256 In this section, we explain how to evaluate the derived clinical pathways by comparing with knowledge-
257 based models from health professionals. In this research, knowledge-based clinical pathways have been
258 developed through the CP task force team (TFT) committee consisting of clinical departments, nursing
259 departments, pharmacy departments, insurance review teams, medical information teams, and
260 management innovation teams. If a target is selected according to the consensus of the clinical
261 department, the initial CP is manually designed based on existing order sets and then assumed to be
262 subject to CP TFT. After identifying the appropriate medication, antibiotics appropriateness, and
263 insurance cutbacks by the CP TFT committee, CP is developed in the EHR system with final approval
264 of the committee. In such a process, the committee meets regularly once a month periodically monitors
265 the results of utilizing the developed CP, receives feedback, and updates the order sets.

266 Figure 2 provides a schema for deriving clinical pathways using the matching rate-based clinical

267 pathway mining algorithm for TLH and evaluating the derived model. We used the delta analysis that
268 identifies the difference with a qualitative approach by domain experts [26]. Thus, we thoroughly
269 compared the difference between the knowledge-based models from domain experts and the data-driven
270 models.

271



272

273 **Figure 2. A schema for deriving clinical pathways and evaluation of them.**

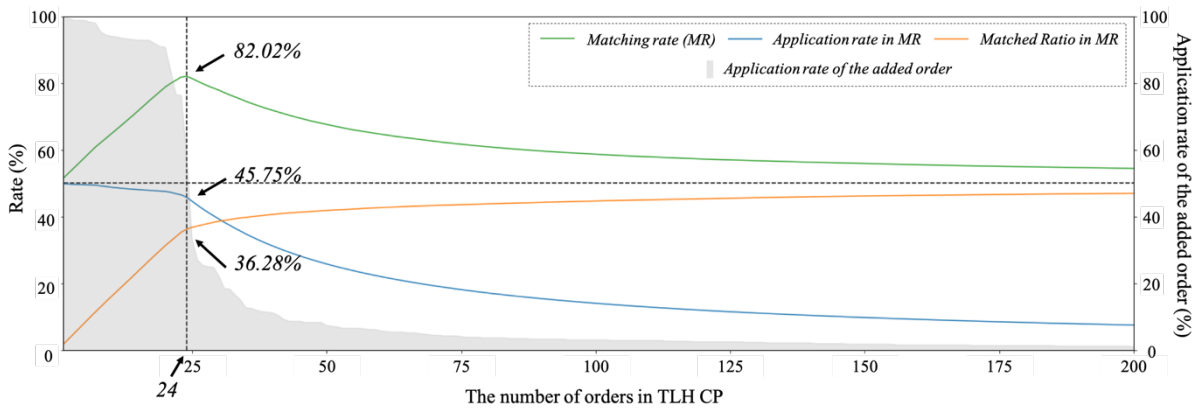
274

275 **4. Results**

276 **4.1. Matching rate-based CP mining algorithm results**

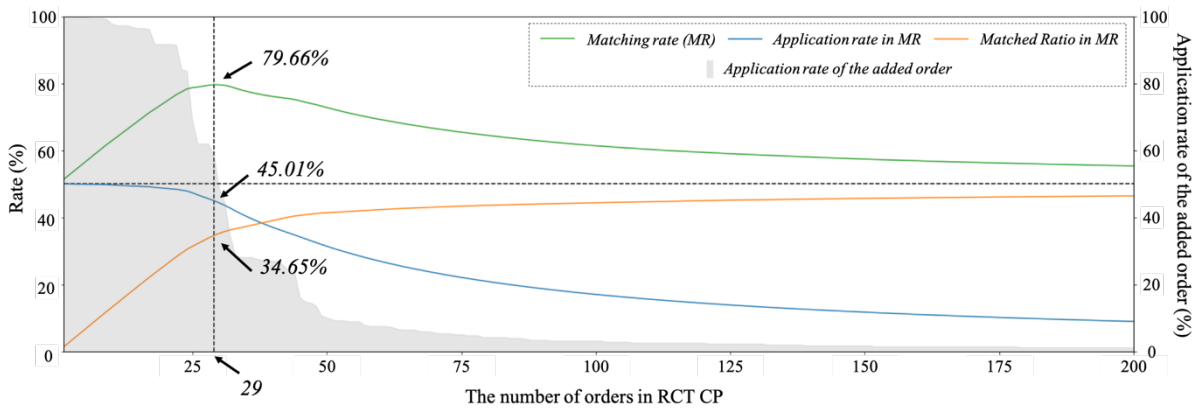
277 This subsection explains the results for deriving data-driven clinical pathways using the proposed
278 matching rate-based clinical pathway mining algorithm. As a result of applying our algorithm, two
279 different sets of clinical orders that maximize the matching rate were prepared for the TLH and RCT
280 clinical pathways, respectively. Figure 3 provides a graphical representation of the process for deriving
281 clinical pathways.

282



283

284 (a) The result for deriving the TLH clinical pathway



285

286 (b) The result for deriving the RCT clinical pathway

287 Figure 3. The results for deriving the clinical pathways

288

289 As provided in Figure 3a, in the process of deriving the TLH clinical pathway, a clinical order with the
 290 highest application rate, i.e., *Acetylcysteine*, was first included in the clinical pathway (application rate:
 291 99.62%). Since its application rate was almost 1.0, the application rate of orders in the clinical pathway
 292 $\left(\frac{1}{2}\left(1 - \frac{M_{CP}}{N_{CP}}\right)\right)$, i.e., the left side of the matching rate, was close to 50.0% (more precisely, 49.81%).
 293 However, the other side of the matching rate, i.e., the matched ratio of orders in the log data
 294 $\left(\frac{1}{2}\left(1 - \frac{R_{Log}}{N_{Log}}\right)\right)$, was almost 0 (more precisely, 1.64%) since the one-order clinical pathway is not
 295 sufficient to reflect the whole log data. As a result, the matching rate became 51.45% by adding the two
 296 values. Then, we included an additional order with the second highest application rate of 99.42% in the

297 clinical pathway. This decreased the average application rate (49.76%) and increased the matched ratio
298 of orders (3.28%). As such, the matching rate was also increased by 53.04%. Through the repetition of
299 this process, we identified that 24 clinical orders maximized the matching rate as 82.02% and became
300 the optimal set for the clinical pathway; the average application rate and the matched ratio of orders
301 were 45.75% and 36.28%, respectively.

302 Adding one more clinical order with the 25th highest application rate in the clinical pathway, the
303 matching rate was decreased by 81.37% because the average application rate had a greater decrease
304 than the increase in the matched ratio. If we include all 1259 orders in the clinical pathway, the average
305 matched ratio of clinical orders becomes the perfect score of 50%. Even if it has the maximum value,
306 however, the average matching rate becomes at most 51.31% since the application rate has the lowest
307 value of 1.31%.

308 Similar to the procedure of deriving the TLH clinical pathway, we discovered the RCT clinical pathway.
309 Figure 3b provides the results for deriving the RCT model. As a result of applying the proposed
310 algorithm, it was determined that a clinical pathway with the top 29 clinical orders within the application
311 rate had the highest matching rate of 79.66%; the average application rate and the matched ratio of
312 orders were 45.01% and 34.65%, respectively.

313

314 **4.2. Comparison with knowledge-based clinical pathways**

315 **4.2.1. The statistical comparative analysis results**

316 From the derived data-driven clinical pathway, we performed a comparative analysis with the
317 knowledge-based model to show the outperformance of the proposed algorithm. Table 3 provides the
318 statistical comparative analysis results for the TLH and RCT clinical pathways. Concerning the TLH
319 clinical pathway, the model from our algorithm was relatively more straightforward than the existing
320 model. In detail, one order was newly added in the data-driven clinical pathway, while four existing
321 orders were deleted. Both the average and median matching rates of the derived model were increased

322 by approximately 5% compared to those of the existing model. Additionally, as a result of conducting
 323 the statistical hypothesis testing, e.g., a *t-test* [27], it was determined that the matching rate of the data-
 324 driven clinical pathway was significantly higher than that of the knowledge-based model (*p-value* <
 325 0.001). Regarding the RCT clinical pathway, there was a dramatic reduction in the number of clinical
 326 orders in the derived model. In the newly developed clinical pathway, 33 existing orders were removed,
 327 while 5 orders were newly included. Regarding the matching rate, the average and median from the
 328 model applied in the proposed algorithm were increased by approximately 23% and 24%, respectively,
 329 compared to those from the knowledge-based model. Furthermore, similar to the TLH case, the
 330 statistical result showed that the data-driven clinical pathway significantly outperforms the model from
 331 domain experts (*p-value* < 0.001).

332 **Table 3. The statistical comparative results for the TLH and RCT clinical pathways**

	TLH		RCT	
	Knowledge-based	Data-Driven	Knowledge-based	Data-Driven
Number of orders	27	24	57	29
Added	-	1	-	5
Removed	-	4	-	33
Average of matching rates (%)	76.97	82.02	56.64	79.66
95% CI of matching rates (%)	(75.89-78.05)	(80.89-83.16)	(55.82-57.47)	(78.52-80.79)
T-test (<i>p-value</i>)	-	<0.001	-	<0.001
Median of matching rates (%)	80.03	85.12	57.12	81.17
SD of matching rates (%)	12.51	13.21	7.93	10.90

333

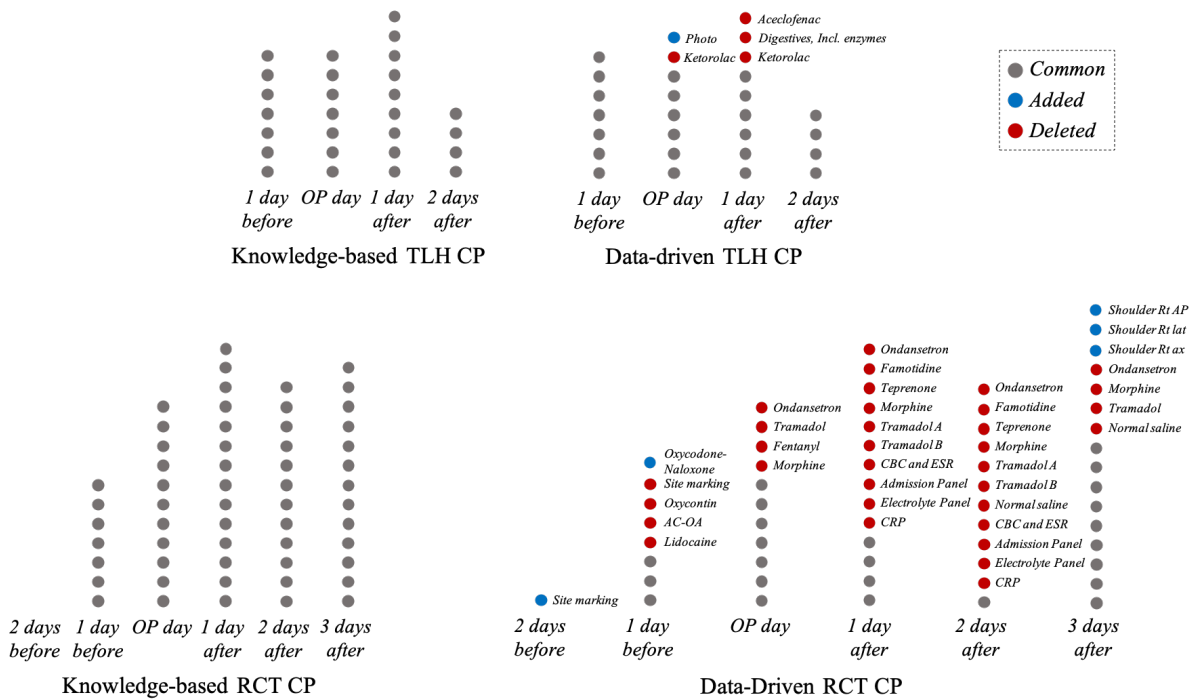
334 **4.2.2. The delta analysis results**

335 We performed the order-level delta analysis to identify the difference between clinical pathways within
 336 an order level, and Figure 4 provides the results of the analysis. Regarding the TLH clinical pathway,
 337 two models from the domain experts and log data were composed of 27 and 24 clinical orders over four
 338 days, respectively. Specifically, *photography* was newly added to the data-driven model, while four
 339 medicine orders (e.g., *ketorolac*, *aceclofenac*, and *multienzymes*) were deleted on the operation day and
 340 one day after. Regarding the RCT clinical pathway, there was a significant difference between the newly

341 developed and the existing model. In contrast to the existing clinical pathway being a five-day schedule,
 342 the data-driven clinical pathway was developed over six days. To this end, *site marking* was moved
 343 from one day before to two days before the operation day. Additionally, *oxycodone-naloxone* and three
 344 orders for *shoulder Rt* were added one day before and three days after the operation, respectively.
 345 Furthermore, 33 clinical orders, including *ondansetron*, *tramadol*, *fentanyl*, and *morphine*, were
 346 removed from the clinical pathway.

347

348



349

350 **Figure 4. The order-level comparative analysis**

351 We performed further comparative analysis on the basis of dates (e.g., 1 day before, OP day, and 1 day
 352 after), order stages (e.g., regular, pre-operation, post-operation, and discharge) and order types (e.g.,
 353 medications and tests) to identify the details of the difference between two models.

354 The results of the detailed delta analysis for the TLH clinical pathway is provided in Table 4. Comparing
 355 two different TLH models, it was identified that most of the clinical orders are included in the
 356 medication type and that there is no significant difference, i.e., commonly utilized orders in both models.

357 Regarding the deleted orders in the data-driven model, it was recommended to remove *Ketorolac*
 358 included in the post-operation of the operation day and the regular stage of the 1 day after. Also, the
 359 model suggested deleting two medication orders in the discharge stage of the 1 day after since there
 360 were not enough patients who leave out the hospitals on the next day after the surgery. As far as the test
 361 type orders were concerned, both models included two orders, e.g., *CBC* and *Urine analysis*, in the
 362 regular stage of the 1 day after, and the supplement of a single order, e.g., *Photography*, in the OP day
 363 was recommended in the data-driven model.

364

365 **Table 4. The detailed delta analysis results for the TLH clinical pathway**

Day	Stage	Medication			Test		
		Common	Added	Deleted	Common	Added	Deleted
1 day before	Regular	Electrolytes, Thioglycolic, Bisacodyl, Sodium Phosphate, Multienzymes, Magnesium Citrate, Cefotetan					
OP Day	Regular				Photo		
	Pre-OP	Electrolytes, Cafotetan, Tissue exam.					
	Post-OP	Acetylcystein, Carbohydrates, Famotidine		Ketorolac			
1 day after	Regular	Acetylcystein, Famotidine, Carbohydrates, Cefotetan		Ketorolac	CBC, Urine Analysis		
	Discharge			Aceclofenac, Multienzymes			
2 days after	Regular	Aceclofenac, Multienzymes					
	Discharge	Aceclofenac, Multienzymes					

366

367 As a result of the detailed delta analysis for the RCT clinical pathway (presented in Table 5), medication
 368 type orders occupied the majority of the models, and all stages hold them. Comparing two different
 369 RCT models, there were three significant differences; (i) deleting a series of medication orders (e.g.,
 370 *Ondansetron*, *Tramadol*, *Morphine*, *Fentanyl*, and *Famotidine*) engaged in the post-operation and
 371 regular stages from the OP day to 3 days after, (ii) removing some medication (e.g., *Tramadol* &

372 *Paracetamol and Teprenone*) and test (e.g., *Admission Panel, CBC & ESR, CRP, and Electrolyte Panel*)
 373 orders connected to the discharge stage at 1 day and 2 days after the surgery, and (iii) adding three test
 374 orders (e.g., *Shoulder Rt AP, Shoulder Rt lat, and Shoulder Rt ax*) in the discharge stage at the last day.

375

376 **Table 5. The detailed delta analysis results for the RCT clinical pathway**

Day	Stage	Medication			Test		
		Common	Added	Deleted	Common	Added	Deleted
2 days before	Regular					Site Marking	
1 day before	Regular	Acetaminophen, Pregabalin, Celecoxib	O-N	Lidocaine, Oxycontin			AC-OA, Site Marking
OP Day	Pre-OP	Cefazolin, Electrolytes					
	Post-OP	Palonosetron, Sodium Chloride, EwC, Famotidine, Cefazolin		Ondansetron, Tramadol, Morphine, Fentanyl			
1 day after	Regular	Palonosetron, Sodium Chloride, Cefazolin, T&P		Famotidine, Ondansetron, Tramadol, Morphine			
	Discharge			T&P, Teprenone			AP, CBC & ESR, CRP, EP
2 days after	Regular	T&P		Famotidine, Ondansetron, Tramadol, Morphine			
	Discharge			T&P, Teprenone			AP, CBC & ESR, CRP, EP
3 days after	Regular	Teprenone, Afloqualone, T&P		Ondansetron, Tramadol, Morphine, Sodium Chloride			
	Discharge	Teprenone, T&P			AP, CBC & ESR, CRP, EP	Shoulder Rt AP, Shoulder Rt lat, Shoulder Rt ax	

377 (O-N: Oxycodone-Naloxone, EwC: Electrolytes with Carbohydrates, T&P: Tramadol & Paracetamol, AP:
 378 Admission Panel, EP: Electrolyte Panel)

379

380 4.3. Organizational relevance

381 The data-driven CPs were reviewed and commented by domain experts including the obstetrics and
 382 orthopedic clinicians. As far as the TLH clinical pathway was concerned, the clinical order that needed
 383 to be added was revealed after visual inspection of the resected pathologic tissue after the operation.

384 Since it should be decided whether or not the order was issued according to the surgical result, expert
385 commented to exclude it from the CP. Additionally, The four medication orders recommended for
386 removing (eg, ketorolac, aceclofenac, and digestives, incl. Enzymes) were analgesic and digestive
387 system orders to prescribe to patients with pain and dyspepsia, which can occur frequently after surgery.
388 As a result, it was concluded that we need to improve the system to make it possible to provide those
389 orders only to the necessary patients rather than issuing them regularly.

390 As far as the RCT clinical pathway was concerned, the US Extremity site marking test was added two
391 days before the operation, which provoked the change in the total schedule of the clinical pathway.
392 Additionally, we recognized that three orders for X-ray photography were regularly implemented before
393 the patients were discharged. In addition, it was discovered that 33 orders recommended to be removed
394 were used to avoid bleeding, prevent infection, relieve pain, and improve digestion. As with the TLH
395 clinical pathway, we concluded that those orders should not be commonly applied to all patients.

396 Overall, both clinical departments agreed that the results of this study could be well accepted and
397 reflected in the clinical setting. The orders derived from the data-driven CPs provide reliable results,
398 but the final decision should be made in a semi-automatic manner after expert reviews according to the
399 various context of the patient or medical practice patterns of the healthcare organization. Also,
400 considering these results, the hospital prepared the revised clinical pathway by modifying clinical orders
401 and changing the schedule, and we finally arrived at a conclusion that it is necessary to develop a new
402 system that can recommend the orders appropriately according to the patients' symptoms or test results.

403 **5. Discussion**

404 An existing study presented four criteria to define a new clinical pathway: (i) a structured
405 multidisciplinary plan of care, (ii) translating guidelines into local structures, (iii) detailed steps in a
406 course of care in a plan, and (iv) aiming to standardized care for a specific population [28]. Based on
407 these criteria, we evaluated whether our algorithm is suitable for deriving a clinical pathway. First, the
408 output of the algorithms, i.e., a set of clinical orders, are the clinical activities that members in multiple
409 disciplines are intimately involved in. In particular, this study utilized data from inpatients who have

410 undergone TLH and RCT surgery performed with multidisciplinary care; the same is true for the derived
411 models from data. Also, our algorithm takes EHR data, i.e., evidence, and provides a structured and
412 detailed order plan for a specific clinical context (e.g., TLH or RCT). Therefore, we argue that clinical
413 pathways from our algorithm address the full four criteria.

414 This research makes a significant contribution of automatically developing clinical pathways based on
415 the collected data in electronic health records. Hospitals generally cannot build clinical pathways due
416 to an insufficient workforce, time, and costs. The proposed algorithm will enable the preparation of
417 more accurate clinical pathways without any human intervention. In other words, this paper is of value
418 because it is useful for supporting decision making with an evidence-based approach.

419 The proposed algorithm produces a fine-grained model (i.e., at the order-level), not a coarse-grained
420 model (i.e., at the activity-level). Thus, it is directly applicable to medical practices. Additionally, it is
421 extensible where various clinical pathways can be developed. For example, we can develop a clinical
422 pathway that considers operations and patient characteristics together, e.g., a TLH-diabetic-female
423 clinical pathway.

424 Despite these contributions, this study has several limitations. First, this research highly depends on
425 data variability. Specifically, it would be difficult to identify clinical pathways based on data from
426 internal medicine departments that have a high variability of patient behaviors. To overcome this
427 challenge, future studies should develop a method to determine in advance whether a clinical pathway
428 needs to be created and an approach for building a clinical pathway considering both data and domain
429 knowledge. *Second, this research does not cover distinguishing of infrequent events from random noise
430 since our primary research goal is to derive a standardized order set. However, it may require including
431 a couple of orders for specific patients who have the same comorbidity with the first-diagnosis. To deal
432 with this, future works should present a method for analyzing the need for clinical pathway
433 segmentation by analyzing the relationship between patient characteristics and their clinical orders and
434 developing the relevant branch CPs.* Additionally, the analysis results presented in this paper were only
435 from a single hospital. Thus, this study may lack generalizability since clinical pathways and their data

436 can differ among hospitals. Future studies should perform more case studies using data from multiple
437 hospitals.

438

439 **6. Conclusion**

440 This paper suggested a matching rate-based clinical pathway mining algorithm to automatically develop
441 clinical pathways based on collected data in electronic health records. The practical applications at a
442 real site resulted in a rise in matching rates of two different clinical pathways, i.e., the TLH and RCT
443 clinical pathways. This research will be helpful in supporting clinical decision making and can be
444 directly applied in medical practices.

445

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451

452 **Statement on conflicts of interest**

453 None.

454

455 **Summary table**

456 What was already known on the topic

457 - Clinical pathways deliver structured clinical services by evidence-based healthcare in a
458 specified medical context.

459 - Knowledge-based models by health professionals are prominent; however, nowadays, data-

460 driven clinical pathways have been implemented with the data from electronic health records.

461 What this study added to our knowledge

462 - Our algorithm, i.e., matching rate-based clinical pathway mining algorithm, can automatically
463 develop the fine-grained clinical pathway at the clinical order level.

464 - In the real-life cases of TLH and RCTs, we identified that data-driven clinical pathways using
465 our algorithm could complement the deficiencies of the knowledge-based models. Therefore,
466 this research will help support clinical decision making and can be directly applied in medical
467 practices.

468

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