Managing Complexity. From Documentation to Knowledge Integration and Informed Decision Findings from the Clinical Information Systems Perspective for 2018

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Summary
Objective: To summarize recent research and to propose a selection of best papers published in 2018 in the field of Clinical Information Systems (CIS).
Method: Each year a systematic process is carried out to retrieve articles for the CIS section of the IMIA Yearbook of Medical Informatics and to select a set of best papers for the section. The same query as in the last five years was used. The retrieved articles were categorized in a multi-pass review carried out by the two section editors. The final selection of candidate papers was then peer-reviewed by Yearbook editors and external reviewers. Based on the review results the best papers were then chosen at the selection meeting with the IMIA Yearbook editorial board. Text mining, and term co-occurrence mapping techniques were again used to get an overview of the content of the retrieved articles.
Results: The query was carried out in mid-January 2019, yielding a consolidated, deduplicated result set of 2,264 articles which had been published in 957 different journals. This year, we nominated twelve papers as candidates and three of them were finally selected as best papers in the CIS section. Again, the content analysis of the articles revealed the broad spectrum of topics which is covered by CIS research.
Conclusions: We could observe ongoing trends from our 2017 analysis. The patient increasingly moves in the focus of the research activities and trans-institutional aggregation of data is still an important field of work. The move to use patient and other clinical data directly for the patient and to support data driven process management, the move away from clinical documentation to patient-focused knowledge generation and support of informed decision, is gaining momentum by the application of new or already known but, due to technological advances, now applicable methodological approaches.

Keywords
Medical informatics, International Medical Informatics Association, Yearbook, Clinical Information Systems

Introduction
In our synopsis for the 2018 issue of the IMIA Yearbook of Medical Informatics [1] we concluded that, modern clinical information systems served as backbone for a very complex, trans-institutional information logistics process and that data was more and more reused for multiple purposes. Last year we found a lot of examples showing the benefits and novel approaches to tackle the challenges of such data reuse. And we found that the patient was moving in the focus of interest of CIS research. This year we observed similar results.

About the Paper Selection
The selection process in the CIS section is stable now for five years. It is described in detail in [1] and the full queries are available upon request from the corresponding authors.
The queries were carried out in mid-January 2018. This year the search result set comprised 2,264 unique papers. From these papers 2,160 were retrieved from PubMed and 104 additional publications were found in Web of Science®. The resulting articles had been published in 957 different journals. Table 1 depicts the top twenty journals with the highest numbers of resulting articles. This year we used RAYYAN, an online systematic review tool [2] to carry out the multi-pass review by the two section editors (AH, WOH). Both section editors independently reviewed all 2,264 publications. In the first step, ineligible articles were excluded based on their titles and/or abstracts (section editor 1: n=2,184; section editor 2: n=2,140). In this first step, the agreement between the two editors was n=2,089 for “exclude” and n=29 for “not exclude” (i.e. include). The remaining 146 conflicts were solved on mutual consent which resulted in nine additional inclusions. The final candidate selection from the remaining 38 publications was done based on full text review and yielded 12 candidate papers for the CIS section 2019.
This list was then reviewed by the Yearbook editors who checked if any articles had also been selected for other sections. As in the last year, no overlaps were found, and all twelve candidates were sent in the peer-review process for the Yearbook. For each paper at least five independent reviews were collected. During the selection meeting held on April 26, 2019 in Paris where section editors from all Yearbook sections participated, three papers [3–5] were finally selected as best papers for the CIS section (Table 2). A content summary of these three best CIS papers can be found in the appendix of this synopsis.

### Findings and Trends: Clinical Information Systems Research 2018

During the selection of the best papers we as section editors get a broad overview on the research field of our CIS section. Of course this overview may be biased and to avoid selective perception we again applied our more formal text mining and bibliometric network visualizing approach [6] to summarize the content of titles and abstracts of the articles in our CIS result set as in the previous years [1, 7, 8].

This year we also extracted the 21,264 authors’ keywords from all articles and present their frequency in a tag cloud (cf. figure 1). We found 7,250 different key words, of which 5,149 were only used once. Most frequent keywords were “human” (n=699) followed by “female” (n=296), “male” (n=263), “electronic health record(s)” (n=235), “adult” (n=230), “child” (n=191) and “health communication” (n=164).

Figures 2 and 3 depict the resulting co-occurrence maps of the top-100 terms from the titles and of the most relevant terms (top 60 percent, n =524) from the abstracts of the 2,264 papers of the recent CIS result set. In 2017 we discovered six different clusters of terms within the titles, last year four and this year we found five main clusters. Whereas “human” was the top authors’ keyword in the articles, the analysis of the titles revealed “system” as central hub for these five clusters.

The cluster analysis of the abstracts yielded also five clusters (cf. figure 3). The red cluster on top comprises terms connected with electronic health records (EHR) and requirements, data sharing, interoperability, architecture and integration. The blue cluster on the right side is dedicated to geographic information systems and related terms. The violet cluster on the bottom and center represents medication and adverse event related research. The yellow cluster on the bottom left as well as the green cluster on the left side can be seen as containers for context factors mentioned in the abstracts.

As in the previous years, the results of these analyses demonstrate the wide variety of the CIS domain. Again, we found a multitude of well written and highly interesting articles of first-rate quality. Among these we interestingly found a high proportion

### Table 1 Number of retrieved articles for Top-20 journals

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<thead>
<tr>
<th>Journal (Total Number of Journals = 957)</th>
<th>Number of papers</th>
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<tbody>
<tr>
<td>PLOS ONE</td>
<td>51</td>
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<tr>
<td>JOURNAL OF THE AMERICAN MEDICAL INFORMATICS ASSOCIATION</td>
<td>45</td>
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<tr>
<td>HEALTH COMMUNICATION</td>
<td>41</td>
</tr>
<tr>
<td>APPLIED CLINICAL INFORMATICS</td>
<td>36</td>
</tr>
<tr>
<td>JOURNAL OF MEDICAL INTERNET RESEARCH</td>
<td>35</td>
</tr>
<tr>
<td>INTERNATIONAL JOURNAL OF MEDICAL INFORMATICS</td>
<td>34</td>
</tr>
<tr>
<td>INTERNATIONAL JOURNAL OF ENVIRONMENTAL RESEARCH AND PUBLIC HEALTH</td>
<td>32</td>
</tr>
<tr>
<td>INTERNATIONAL JOURNAL OF CLINICAL PHARMACY</td>
<td>24</td>
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<tr>
<td>JOURNAL OF MEDICAL SYSTEMS</td>
<td>24</td>
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<tr>
<td>Bmj Open</td>
<td>23</td>
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<tr>
<td>Bmj health services research</td>
<td>22</td>
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<tr>
<td>Drug safety</td>
<td>19</td>
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<tr>
<td>Computers</td>
<td>17</td>
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<tr>
<td>JOURNAL OF HEALTH COMMUNICATION</td>
<td>15</td>
</tr>
<tr>
<td>JOURNAL OF EVALUATION IN CLINICAL PRACTICE</td>
<td>14</td>
</tr>
<tr>
<td>INFORMATICS FOR HEALTH &amp; SOCIAL CARE</td>
<td>14</td>
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<tr>
<td>ENVIRONMENTAL MONITORING AND ASSESSMENT</td>
<td>14</td>
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<tr>
<td>Bmj public health</td>
<td>14</td>
</tr>
<tr>
<td>Patient education and counseling</td>
<td>13</td>
</tr>
<tr>
<td>Bmj medical informatics and decision making</td>
<td>12</td>
</tr>
</tbody>
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### Table 2 Best paper selection of articles for the IMIA Yearbook of Medical Informatics 2019 in the section ‘Clinical Information Systems’. The articles are listed in alphabetical order of the first author’s surname.
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of articles from Chinese researchers. This is also reflected in our candidate selection where half of the twelve papers come from China or involve Chinese contributors.

From a thematic perspective we identified a growing proportion of contributions showing that the potentials and possibilities due to technological advances, increased computing power, availability of a huge amount of structured data and sophisticated algorithms now can be unlocked.

The first of the best papers in the CIS section 2019 is an example for that. Cao Xiao and colleagues developed a novel deep learning model that learns distributed patient representation from the EHR data and performs prediction for the 30-day readmissions [10]. This inspiring paper is a very well-presented example of how deep learning techniques can be applied to distill new knowledge out from existing EHR data. The approach is highly innovative, and the developed model outperforms state-of-the-art approaches in readmission prediction.

Another well-presented example of a powerful risk prediction model (1-year incident hypertension) that is based on and validated using a huge amount of patient data by Chengyin Ye and colleagues [11] can be found among the candidate papers.

An "evergreen" within the scope of CIS research is the detection of adverse events within EHR data. This year we chose a contribution by Jiebin Chu and colleagues who used neural attention networks for that [12]. Data Mining also was an important keyword during the last years in CIS research. This year we didn’t find this keyword very often (cf. figure 1). However, a very interesting article introducing a novel data or text mining technique termed as optimized swarm search-based feature selection [13] coming from Daohui Zeng and colleagues was included in the selection of candidate papers. The “newcomer” Blockchain as identified last year was also a prominent research topic in 2018. We included a contribution by Hongyu Li and colleagues on a blockchain-based data preservation system for medical data in our CIS best paper candidate pool.

The second of the actual best papers tackles an important problem within the domain of clinical information systems in a very inspiring way. Sharidan K. Parr and colleagues developed and applied a machine learning approach that leverages “noisy labels” for automatic mapping of an enormous amount of laboratory data (>6.5 billion lab test results) to LOINC codes [14]. Among the candidate papers another one by Ronald George Hauser and colleagues dealt in a very inspiring way with the standardization of laboratory test results [15].

An increasing number of research articles deals with process mining or data driven process modeling in the healthcare domain. So for example the contribution from Jingfeng Chen and colleagues on automatic extraction of typical treatment processes from electronic health record data [16].

On the other hand, clinical processes themselves can also deliver data that can be used for different analyses. Kaat De Pourcq and colleagues present a step-by-step methodology to implement an automated process-oriented performance measurement system for hospitals [17].

Besides all benefits and possibilities, the use of massive EHR data sets can bring, we have to keep in mind that reality and data not necessarily are congruent. Without careful
Fig. 2 Clustered co-occurrence map of the most relevant terms (top-100) from the titles of the 2,264 papers in the 2019 CIS query result set. Node size corresponds to the frequency of the terms (binary count, once per paper). Edges indicate co-occurrence and distance of nodes corresponds to the association strength of the terms within the titles (only top 200 out of 455 edges are shown). Colors represent the five different clusters. The network was created with VOSviewer [9].

consideration of the context in which these data are produced many pitfalls and sources of bias may be overlooked. Denis Agniel and colleagues, with their very worth reading candidate paper, remind us of this eternal truth. Another core aspect that may not be overlooked within the CIS domain are the users. Pantelis Natsiavas and colleagues propose a comprehensive user requirements methodology for secure and interoperable health data exchange [18]. This candidate paper is worth reading as it also provides a comprehensive set of user requirements and presents sets of barriers as well as facilitators for health IT solutions.

Last but not least we want to highlight the third of the best papers in the CIS section 2019. In their article which received top ratings from the IMIA Yearbook reviewers Johannes C. Eichstaedt and colleagues show that the content shared by consenting users on Facebook can predict a future diagnosis of depression [3]. This study suggests that social media content may point clinicians to specific symptoms of depression. It sheds a new light on privacy and anonymity with regard to health information and vividly indicates that health related information can not only be retrieved from “core” medical and health data, but also from other user provided and publicly available data, such as social media content.
As every year, at the very end of our review of findings and trends for the clinical information systems section, we want to recommend a reading of this year’s survey article in the CIS section [19].

**Conclusions and Outlook**

We could observe ongoing trends from our 2017 analysis that the patient increasingly moves in the focus of the research activities and that the trans-institutional aggregation of data is still an important field of work. Common to these efforts is the creation of benefits for the patients. A key differentiator from what we observed in the last years are the methods applied to reach this goal. As already been described in the findings we
found a great number of papers – amongst them the best papers for 2018 – that were tackling problems using machine learning approaches combining them with relatively large and sometimes immense clinical data sets or so far uncommon data sets outside the core clinical domain. Apart from these ongoing developments we have also found inspiring work that deals with data driven management of processes and the use of blockchain technology to support data aggregation beyond institutional boundaries. The move to use patient data directly for the patient, to move away from clinical documentation to patient-focused knowledge generation and support of informed decision, is gaining momentum by the application of new or already known but, due to technological advances, now applicable methodological approaches.

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References

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Facebook language predicts depression in medical records
Proc Natl Acad Sci U S A 2018;115(44):11203-8

The paper deals with depression, one of the most prevalent mental illnesses. The authors state that each year around 7%-26% of the US population experience a depression or depression related symptoms. The number of patients that receive minimally adequate treatment is according to their investigation at most 49%. Therefore, the authors conclude that such high rates of underdiagnosis and undertreatment suggest that existing procedures for screening and identifying depressed patients are inadequate. The authors introduce a novel approach by using Facebook language data from a sample of consenting patients to detect/predict depression from that data. The study involved the analysis of the Facebook posts of 114 patients that prior had a diagnosis of depression. For each of the patients with such a diagnose, five random patients without a diagnose of depression in the same period of time were added to the analysis to simulate the prevalence of the disease. The authors built a prediction model using the textual content of the Facebook posts, post length, frequency of posting, temporal posting patterns, and demographics. A 10-fold cross-validation was applied to avoid overfitting. To yield interpretable and fine-grained language variables, 200 topics were extracted using latent dirichlet allocation. The results suggest that the closer in time the Facebook data are to the documentation of depression in the EMR, the better their predictive power. Within 6 months preceding the documentation of depression an accuracy of 0.72(AUC) was achieved. The authors conclude that Facebook language-based prediction models perform similarly to screening surveys in identifying patients with depression when using diagnostic codes in the EMR to identify diagnoses of depression.

Parr SK, Shotwell MS, Jeffery AD, Lasko TA, Matheny ME
Automated mapping of laboratory tests to LOINC codes using noisy labels in a national health record system database
J Am Med Inform Assoc 2018;25(10):1292-300

Aggregated data from multiple data-sources can be a valuable source for different fields of research and other domains such as public health or the creation of clinical evidence. A very common problem related to the use of data from different sources is their different coding and the use of non-standardized terminology. An important aspect in the assessment of patient outcomes are laboratory findings. In this context Logical Observation Identifiers Names and Codes (LOINC) can be seen as an important standard. However, mapping laboratory findings to LOINC codes manually can be time consuming. As electronic health records (EHR) are a rich source of data accumulated through routine clinical care the authors aim to develop a machine learning algorithm to automate mapping of unlabeled data and reclassification of incorrect mappings within labeled data. For this purpose, inpatient and outpatient laboratory data (6.6 billion laboratory results) from 130 Veterans Affairs (VA) hospitals was collected. The dataset used for training contained an unknown number of labelling errors and was not manually cleaned (noisy labelling approach). They implemented logistic regression, a random forest multiclass classifier, and a 1-versus-rest ensemble of binary random forest classifiers. All models were refined with a 5-fold cross-validation. Although the mapping results of the models are not in all cases convincing, the authors investigated important reasons for this matter. In addition, they were able to prove that their results are similar in accuracy to the best reported automated methods for laboratory test mapping. This is a remarkable result as the model was built using noisy data. The approach is one of the first that can be applied fully automated with no need of manual intervention.

Xiao C, Ma T, Dieng AB, Blei DM, Wang F
Readmission prediction via deep contextual embedding of clinical concepts

Hospital readmission is a critical figure in the assessment of the quality of a treatment process. Indeed, many hospital readmissions are avoidable and pose a certain risk for the patient. Although it is not an easy task to predict hospital readmission as it is not only related to the disease and the treatment but also a complex set of risk factors which are interrelated. The current paper aims at presenting a hybrid deep learning model structure that combines topic modeling and Recurrent Neural Network (RNN) to distill the complex knowledge hidden in those contexts and perform accurate readmission prediction. The proposed “CONTENT” model covers both the global and local contexts within the patient journey from an EHR through a hybrid Topic Recurrent Neural Network (TopicRNN) model. It transforms patients’ complicated event structures into deep clinical concept embedding, which can be viewed as a novel form of patient representation encoding the patient clinical conditions from both long and short terms. In order to build that model the EHR data (including disease, lab findings and medication codes) from a cohort of 5,393 patients with congestive heart failure was used. The model outputs a context vector for each patient, which characterizes his/her overall condition. The proposed model outperforms existing methods in readmission prediction (e.g. 0.6103 ± 0.0130 vs. second best 0.5998 ± 0.0124 in terms of ROC-AUC). The derived patient representations were further utilized for patient phenotyping. The subgroups allow a better understanding of different readmission risks in the cohort.