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An Ontology-based Framework for Authoring and Executing Clinical Practice Guidelines for Clinical Decision Support Systems

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ABSTRACT

Clinical Practice Guidelines (CPG) are used by healthcare practitioners to standardise clinical 15 practice and to provide evidence-based healthcare. However, due to the paper-based nature 16 of CPG they are under-utilised at the point-of-care. In this paper we present our CPG-based 17 Clinical Decision Support System (CDSS) development framework - CPG-EX. This offers the functionality to (i) model a CPG in a computer-interpretable format; and (ii) execute the 18 modelled CPG based on patient data to deliver CPG-mediated recommendations in line with 19 the patient's conditions. We have taken a Semantic Web approach and employ ontologies 20 to model the CPG knowledge and proof engines to execute the CPG. CPG-EX comprises three different ontologies, namely CPG ontology, Domain ontology and Patient ontology, 21 that interact at a semantic level to represent the entire disease-specific knowledge. We have 22 developed a forward-chaining CPG execution engine that executes the set of CPG decision-23 rules using JENA (a semantic web framework for JAVA) reasoning to provide patient-specific CPG-mediated recommendations. We also implemented an automated justification tree gen-24 eration module that provides the inference trace for the solution in order to assist practitioners 25 in understanding the rationale for the proposed recommendations. A working prototype of our 26 CPG-based CDSS was constructed using an international guideline for ordering radiological investigations. This was tested using a number of real-life clinical cases and both the recom-27 mendations and their justifications were validated by medical practitioners. 28

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31 INTRODUCTION

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Despite the proliferation of disease-specific Clinical Practice Guidelines (CPG) their
utilisation in healthcare settings, especially at the point-of-care, is low¹. Barriers to
the poor utilisation of CPGs in the care delivery process are manifold and can be
broadly categorised along three dimensions:
Operational issues that concern the accessibility and usability of the paper-

- 38 based CPGs in a timely manner at the point-of-care
- 39 40

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Behavioural issues that involve the perception of health professionals towards
 the notion that CPGs tends to restrict or constrain their practices of patient
 care as per their judgment

Technical issues that involve the development of practical computer-based methods to both model the disease-specific knowledge encapsulated within
 the CPG and execute the modelled CPG to provide CPG-mediated and patient-specific recommendations for healthcare professionals.

Lately, there has been an upsurge in efforts to optimise the use of CPGs, especially support and standardise clinical decisions by healthcare professionals. A popular approach, at the operational and technical levels, towards the operationalisation of CPGs is to model them in a computer-interpretable format² and then incorporate the modelled CPGs within computer-based Clinical Decision Support Systems (CDSS)^{3,4}. The disease-specific CDSS can then be deployed within a healthcare setting to:

- Execute the CPG at the point of care
- Guide healthcare practitioners to make evidence based decisions, actions and
 recommendations
- Standardise the delivery of care at a particular healthcare setting
- 19 Collect all necessary and relevant patient data.

The development of CPG-based CDSS is a challenging activity as it involves (i) 20 Computerising the paper-based CPG into a computer interpretable and executable 21 format. This involves the modelling of the disease-specific knowledge inherent 22 within a CPG and representing it in a semantically unambiguous formalism. The 23 modelling exercise identifies the key concepts, the relationships between the con-24 25 cepts, the decisional elements and the consequences of these decisions, the data elements; (ii) Specifying a functional and executable workflow of the CPG that entails 26 interactions between the different CPG elements and the corresponding actions and 27 interventions; (iii) Abstracting the clinical decision logic inherent within a CPG in 28 terms of medically salient and executable logic-based decision rules; (iv) Executing 29 the computerised (or modelled) CPG based on both acquired and inferred patient 30 information, to recommend patient-specific recommendations; and (v) Justifying 31 the recommendations suggested by the CDSS to establish a degree of 'trust' in the 32 33 output of the CDSS.

Typically, the development of CDSS involves a tedious and challenging knowl-34 edge engineering process that is geared towards the accumulation of disease-specific 35 knowledge from expert health professionals. CPGs are a validated source of disease-36 specific knowledge that is based on best evidence and designed to assist clinical 37 decision-making⁵. We propose that CPGs can serve as a validated, evidence-driven 38 39 knowledge-base for CDSS (targeting a specific disease), thereby circumventing the knowledge engineering problem faced during the development of CDSS. 40 In this paper we present our CPG-based CDSS development framework, named 41

42 CPG-EX, that offers the functionality to (i) *model* a CPG in a computer-interpretable

format; and (ii) execute the modeled CPG based on patient data to deliver CPG-1 2 mediated recommendations in line with the patient's conditions. For CPG modeling 3 we take a model-centric approach and represent the CPG knowledge using ontologies. For CPG execution we use logic-based reasoning to select the relevant CPG 4 recommendations based on given patient data. We propose to leverage the Semantic 5 Web approach to develop the CPG modeling and execution functionalities of CPG-6 EX. In this paper, we present a prototype of our CPG-EX framework to model and 7 illustrate it in practice using an international guideline devised to aid clinicians in 8 ordering radiology investigations. 9 10 FUNCTIONAL DESIGN OF CPG-EX 11 12 We believe that Semantic Web technology offers an interesting approach to both 13 model and execute CPGs, and in turn develop CPG-based CDSS. The Semantic 14 Web purports the semantic modelling and markup of knowledge in terms of formal 15 definitions of domain concepts, explicit representation of relationships between 16 concepts and any logical constraints between the concepts and relationships⁶. The 17 semantically modelled knowledge can then be reasoned over using proof engines 18 employing logic-based reasoning methods to infer 'trusted' solutions. Therefore, the 19 design of CPG-EX is guided by Semantic Web technologies. The two main functions 20 of CPG-EX are: 21 22 • CPG modelling that allows the transformation of a paper-based CPG into a

- 23 formal representation that can be executed by computer-based CDSS
- *CPG execution* that allows operationalisation of the modelled CPG to derive
 patient-specific CPG-based recommendations
- 26
- 27 CPG Modelling

CPG modelling entails the representation of a paper-based CPG in terms of a for-28 mal and expressive knowledge-model that provides (i) an in-depth understanding 29 of the clinical procedures, addressed by the guideline; and (ii) a precise and unam-30 biguous description of the guideline. CPG modelling is pursued through two main 31 approaches: (i) Document-Centric approach that entails the mark-up of the CPG, 32 33 as per a document modelling language such as XML (extensible markup language), to generate a semi-formal model of the CPG. Guideline Element Model (GEM) is 34 a prominent document-centric CPG mark-up language² that characterises the CPG 35 using over 100 different mark-up tags; (ii) Model-Centric approach aims to gener-36 ate a knowledge-model of the CPG, using a formal model description language, that 37 entails classes, relationships between classes and decision rules that operate over 38 instances of the classes and relationships. Model-based approaches provide a seman-39 tically rich expressivity of the CPG knowledge and are hence preferred whenever 40 CPG execution is also desired. Typical model-based CPG representations include 41

42 GLIF⁸, GUIDE⁹, and Proforma¹⁰.

For CPG modelling we follow the model-centric approach and in line with the 1 2 Semantic Web framework we model all the domain-specific knowledge using specialised ontologies. Ontology-based CPG modelling is pursued by other CPG rep-3 resentations formalisms such as GLIF⁸, HELEN³, SAGE⁴, EON¹¹ and PROforma¹⁰. 4 To capture the entire knowledge resources needed for the CDSS to provide 5 CPG-based recommendations we developed three independent, yet conceptually 6 7 and functionally interoperable, ontologies ¹² using Protege¹³: I. CPG Ontology that models the computerised structure of the CPG. The 8 rationale for the CPG ontology is driven by our belief that to model the 9 knowledge components of a CPG, we first need to understand (and model) 10 the underlying knowledge representation structure of the CPG; hence the 11 need to develop the CPG ontology. We model the CPG using the Guideline 12 Element Model (GEM) structure, and therefore our CPG ontology is based 13 14 on the GEM DTD (Document Type Definition)^Z. II. Domain Ontology that models the medical knowledge encapsulated within 15 the CPG. The Domain Ontology represents both the concepts described in 16 a CPG and the relationships between these concepts. The domain ontology 17 not only standardises the domain concepts but also captures the decision 18 19 logic inherent within the descriptions of the CPG and allows for writing logical decision-rules that relate patient/disease-specific conditions to cor-20 21 responding actions/recommendations. In this project, the domain ontology models the EU Radiation Protection 22 23 118 Referral Guideline for Imaging (RPG)¹⁴. Radiological investigations are 24 routinely used by clinicians to aid patient diagnosis and management. How-25 ever investigations are frequently ordered inappropriately and in addition 26 the most suitable investigation is often not ordered e.g. a CT (Computed Tomography) scan may be ordered when a plain X-Ray is sufficient. The RPG 27 28 provides guidance on whether a radiological investigation is likely to provide useful information for a specific clinical problem, the most suitable imaging 29 modality to be used, the radiation dose associated with an individual imaging 30 modality and the evidence for any recommendations given. The RPG Domain

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Table 1. Example of a Recommendation from EU Radiation Protection 118
 35 - Referral Guideline for Imaging

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37 38	Clinical Problem	Investigation (Dosage)	Recommendation (Grade)	Comments
39 40 41 42	Headache: Chronic (A7)	XR skull, sinus, C spine (l) CT (ll) or MRI (0)	Not indicated routinely (B)	Radiology of little use in the absence of focal signs or symptoms. Some exceptions for specialists or if evidence of raised intracranial pressure, posterior fossa or other signs

Ontology¹⁵ is mainly divided into three categories of concepts: Clinical Prob-1 2 lems, Investigations and Recommendations. All distinct clinical problems are 3 represented as instances of respective clinical problem classes. These classes are arranged as sub-classes under the class Clinical Problem. Investigations 4 are radiological procedures reported in the RPG and are represented similar 5 to the Clinical Problems shown in Table 1. For a given clinical problem one or 6 7 more investigations may be recommended. Recommendations are treatments based on investigations and clinical problems. Each recommendation is rep-8 resented along with the evidence grade to support the recommendation. An 9 indicated recommendation signifies that the investigation is likely to provide 10 information that can contribute to the diagnosis and treatment of the patient. 11 Conversely, a not indicated routinely recommendation emphasises that the 12 investigation is extremely unlikely to provide any information that will assist 13 the clinician in patient management 14

- *III. Patient Ontology* that models the patient in terms of various health informa tion parameters that may constitute the longitudinal medical record of the
 patient. The instances of the patient ontology determine the patient's health
 profile which is subsequently used to 'fire' the necessary decision-rules to
 yield patient-specific recommendations based on the CPG.
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21 CPG Execution

22 CPG execution involves the selection of relevant and correct recommendations from 23 the modelled CPG based on patient-data. CPG execution is pursed through a CPG execution engine that employs a variety of execution methods including: (i) logic-24 25 based reasoning; (ii) workflow engineering and (iii) graph based algorithms. In line with the Semantic Web framework we pursue CPG execution using proof-engines 26 that employ logic-based reasoning over the knowledge represented through the vari-27 ous ontologies. We use the JENA reasoning engine for executing CPG decision logic 28 on patient cases, and generating CPG-mediate recommendations. 29 30

31 ARCHITECTURE OF CPG-EX

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33 Architecturally, CPG-EX is divided into two main modules:

- 34 CPG Encoding Module
 - CPG Rule Authoring and Execution Module

The CPG Encoding Module is used to encode the text-based CPG into the CPG ontology, and to annotate the decision variables and logic structures inherent within the CPG. Both the CPG Ontology and the Domain Ontology are used for CPG encoding. The CPG Rule Authoring and Execution Module is used to define the decision logic rules in a CPG and to execute them based on given patient clinical data. A novel feature of our CPG Execution Module is that it provides an automated justification trace of inferred recommendations. This is to inform practitioners, who



¹⁴ **Figure 1.** *System Design of our Clinical Decision Support System (CDSS)*

15 CPG= Clinical Practice Guideline, DTD= Document Type Definition, GEM =Guideline Element
 16 Model.

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operate on the recommendations, the rationale for the recommendations. Figure 1
 shows the architecture of our Semantic Web based CPG-EX.

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22 CPG Encoding Module

The literature highlights a number of CPG encoding formalism that although are able to encode a CPG yet are unable to execute the encoded CPG in conjunction with a patient's case. Given that we aim to execute the CPG we developed a CPG encoding tool that transforms the CPG in an electronic format that can be executed by our CPG execution (reasoning based) engine. The CPG encoding module handles (a) CPG and Domain ontologies in order to model the CPG; and (b) Rule authoring to write decision rules capturing the CPG decision logic.

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31 Domain Ontology

The Domain Ontology for the RPG¹⁵ is mainly divided into three categories of concepts:

- Clinical Problems
- Investigations
- 36 Recommendations

All distinct clinical problems are represented as instances of respective clinical
 problem classes. These classes are arranged as sub-classes under the class Clinical

39 Problem. Investigations are radiological procedures in RPG and represented in RPG

40 Domain Ontology similar to the Clinical Problems. For a given clinical problem,

41 one or more investigations are recommended. Recommendations are treatments

42 based on investigations and clinical problems. Each recommendation is represented



13 **Figure 2.** Class Hierarchy in the RPG Ontology

14 MRI = Magnetic Resonance Imaging. Nm = Nuclear Medicine, US = Ultrasound, XR= X-ray. 15

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along with the grade of its evidence. An indicated recommendation is most likely 17 to contribute to the diagnosis and treatment, while an investigation that is not rou-18 tinely recommended emphasises that the investigation is unlikely to provide any 19 information that will aid management. Figure 2 shows a fragment of the Domain 20 21 Ontology.

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23 CPG Ontology

We developed the CPG Ontology based on the GEM DTD² using an Ontology Edi-24 25 tor Protégé¹³. The main CPG knowledge is represented in the Knowledge Component 26

27



40 **Figure 3.** *Merging CPG and Domain Ontology*

⁴¹ CPG = Clinical Practice Guideline, KC= Knowledge Component, RPG = Radiology Practice Guide-

⁴² line

class in the CPG Ontology as they describe the procedural, conditional or imperative 1 2 knowledge. We defined the *Recommendation* class, within the knowledge compo-3 nent, to describe the recommended actions (see Figure 3). The recommendations are classified as being either imperative or conditional and represented as classes 4 5 within the Recommendation class. Imperative recommendations found in the CPG are applicable to the entire eligible population. The conditional recommendations 6 7 describe the clinical conditions/scenarios that demand specific actions. These clinical conditions and actions are represented by decision.variable and action.variable 8 classes, respectively. The decision.variable and action.variable classes model the 9 structure and type of a decision or action step in the CPG. We also made use of the 10 property logic in conditional/imperative recommendations to define the decision 11 logic of conditional/imperative recommendations based on the conditions for the 12 various actions. 13 14 In our work, we merged two inter-related ontologies for representing CPG knowledge (Figure 3). The first ontology, i.e. the CPG Ontology, models the structural 15 knowledge of the CPG, whereas the second ontology, Domain Ontology, represents 16 the underling CPG knowledge in terms of medical concepts and their relationship. 17 We established semantic mapping between the atomic entities of the candidate 18 ontologies. Two such atomic entities were: (i) the decision.variable from the CPG 19 Ontology; and (ii) a property from the Domain Ontology that explains that deci-20 sion.variable. This mapping of the ontology nodes (a decision.variable instance and 21 its related property in the Domain Ontology), in the two ontologies was achieved 22 23 through a new property named variable.name that served as the bridge between the two ontologies. The property variable.name belongs to the decision.variable class, 24 25 which represents the structure and type of a *decision.variable* in the encoded CPG, whereas the property variable.name merges each decision.variable instance with a 26 27 28 29 Decision Variable: Variable Name:

30	isRecommended> []	dv1hasClinicalProblem
31	hasRadiationDose> [Radia	dv2hasSolution
32	hasClinicalProblem> [ClinicalProblem> [Clinical	dv3hasRecommendationGra
33	narrower> []	dv6hasInvestigationMethod
34	relation> [string] hasRecommendationGrade	a1isRecommended
35	hasSolution> [Investigatio	
36	cui> []	
37	semantic_type> [] hasInvestigationMethod>	
38	age> [int]	
39	betterThan> [] variable.name> [Property]	
40	moreSensitiveThan> ∏ ▼	

Figure 4. Annotation of Decision Variables and Actions based on RPG Domain
Ontology

1 property from the Domain Ontology that describes the operational details of the

2 decision.variable. Each decision.variable instance with a property variable.name is

3 annotated with a property from the Domain Ontology.

Figure 4 demonstrates the semantic annotation of decision variables $dv1 \dots dv6$ and action variable a1 to represent the inherent logic of the recommendation shown in Table 1. For example, decision variable dv1 is annotated with a property *has*-*ClinicalProblem* in the RPG Domain Ontology, where the property value describes various clinical problems in a RPG test-case. Similarly $dv2 \dots dv6$ are annotated and describe one or more investigations along with their recommendations and recommendation grades based on clinical problems. Action variable a1 is annotated with a property *isRecommended*, where the property value represents the list of investigations along with their methods and recommendations with along their grades.

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14 Patient Ontology

15 The patient ontology mirrors an Electronic Patient Record and models a patient

16 in terms of various health information and patient clinical situations. The patient

17 ontology allowed us to generate standardised descriptions of a patient, which in turn

18 serve as patient *instances* used to execute the decision logic of the CPG to derive

19 CPG-mediated recommendations/actions.

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21 Rule Authoring Sub-Module

22 Functionally, the Rule Authoring Sub-Module uses an encoded CPG (modelled by

23 CPG Encoding module), and allows the user (a medical practitioners) to define 24 the CPG's logical constructs as logic-based CPG rules using our 'simple' CPG Rule

25 Syntax.

The Rule Authoring sub-module is designed to encapsulate the clinical decision making logic inherent within a CPG in terms of logical rules. Upon completion of the rule authoring process, we apply a rule transformation algorithm to transform

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31
                    Logic := IF Decision Variable List THEN Action Variable
                       Action_Variable
32
                    Decision_Variable_List := \not\subset
                                         | dv Rel Node, dv Rel Node, ...., dv Rel Node
33
                    Action Variable := a Rel Node
34
                    Rel := < | <= | > | >= | =
                    Node := ? // variable
| 'a literal' // a plai
35
                                            // a plain string literal
36
                          Algebra
                          [dv_1 \ dv_2 \ \dots \ dv_n]
37
                    Algebra := Value
38
                                | Value + Value
39
                                Value - Value
                               Value * Value
40
                    Value := dv | number // dv must already been declared before
41
```

42 **Figure 5.** CPG Rule Syntax

Rule := bare-rule . [bare-rule] [ruleName : bare-rule] bare-rule := term, ... term -> hterm, ... hterm // forward rule term, ... term <- term, ... term // backward rule</pre> hterm := term [bare-rule] := (node, node, node) // triple pattern
| (node, node, functor) // extended triple pattern term | builtin(node, ... node) //invoke procedural primitive functor := functorName(node, ... node) // structured literal // e.g. http://foo.com/eq node := uri-ref // e.g. rdf:type
// variable prefix:localname Т ?varname 'a literal' // a plain string literal 'lex'^^typeURI // a typed literal l number // e.g. 42 or 25.5

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Figure 6. JENA Rule Syntax

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16 the CPG rules into the JENA syntax so that they can be executed by the Execution

Sub-Module that leverages the JENA inference engine¹⁶. The CPG and JENA rule
 syntax are discussed below.

Figure 5 shows the CPG syntax we have used to write rules in the *logic* element of CPG Ontology. Each rule is a *forward rule*, which has a list of decision variables (premises) and an action variable (conclusion) of the rule, followed by *IF* and *THEN*, respectively. In the decision variable list, each variable *dv* is an equality or inequality relation with either i) a variable, ii) a string, iii) a list of (already declared) decision variables or iv) an algebraic (binary) formula.

Figure 6 illustrates an informal description of the simplified JENA (text) rule syntax¹⁶. The "," separators are optional. The *functor* in an extended triple pattern is used to create and access structured literal values.

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29 *Example A*: Below is an example of a rule written in JENA rule syntax. In it R is a 30 rule written in CPG rule syntax to represent the recommendation shown in Table 31 1, where the annotations dv1 - dv6 are used according to the concepts shown in 32 Figure 4.

R = IF dv1=?, dv2=?, dv3=notIndicatedRoutinely, dv4=?, dv5=?, dv6=? THEN
a1=[dv2 dv6 dv3 dv4]

Finally, the above CPG rule *R* is transformed into JENA syntax via rule transformation algorithm and shown as follows.

- 37 **Transform (***R***)** =
- 38 [conditional1: (?X2 rpg:hasClinicalProblem ?X1) , (?X1 rpg:hasSolution

```
39 ?X3) , (?X3 rpg:hasRecommendation rpg:notIndicatedRoutinely) , (?X3
```

- 40 rpg:hasRecommendationGrade ?X4) , (?X3 rpg:hasInvestigationDetails
- 41 ?X5), (?X5 rpg:hasInvestigationMethod ?X6) -> (?X2 rpg:isRecommended
- 42 List(?X3 ?X6 rpg:notIndicatedRoutinely ?X4))]

🝰 Query Edito	r	
Patient:	Ann	7
, diona		-
Property:	Query	
Results:	Ann isRecommended a7_1, XRSkullSinusCSpine, notIndicatedRoutinely, grade	9B 🔺
	Ann isRecommended a7_3, general_MRJ, notIndicatedRoutinely, gradeB	_
	Ann type Patient	
	Ann haeClinicalDrohlom clinicalDrohlom67	
	Derive	
Derivation:	Premises:	^
	Ann hasClinicalProblem clinicalProblemA7 clinicalProblemA7 hasSolution a7 1	
	a71 hasRecommendation notindicatedRoutinely	1
	a7 1 hasinvestigationDetails InvestigationDetail 014	=
	InvestigationDetail 014 hastrivestigationMethod XRSkullSinusCSpine Conclusion:	
	Ann IsRecommended a7 1, XRSkullSinusCSpine, notIndicatedRoutine	y, gradeB
	<	>

Figure 7. *Derivation Trace for Recommendations for a hypothetical patient Ann*

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17 CPG Execution Module

18 The CPG Execution Module is designed to execute the CPG decision-rules, using 19 patient data, to derive CPG-based recommendations. To achieve this functionality, 20 we built two sub-modules namely, Execution Sub-Module and Justification Trace

21 Sub-Module.

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23 Execution Sub-Module

The Execution Sub-Module invokes the JENA inference engine¹⁶ to execute a CPG in order to infer recommendations based on patient profiles. We model instances from the Domain Ontology, CPG Ontology and Patient Ontology as RDF graphs, which serve as the knowledge base for JENA. The JENA inference engine uses both the knowledge base and the CPG rule-set in a backward reasoning mode to infer CPG-mediated recommendations based on the given patient scenario, encoded clinical knowledge in the Domain Ontology and CPG Ontology.

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32 Justification Trace Sub-Module

33 The Justification Trace Sub-Module generates a justification trace of the rule 34 execution to assist medical practitioners in understanding the logic behind the 35 inferred recommendations. The justification derivation includes the linear rep-36 resentation of premises (facts) under which the JENA rules are satisfied and the conclusions based on those rules. The justification trace initiates with a derived 37 patient recommendation (derived facts) from the JENA model (knowledge base) 38 39 and generates facts which served as premises for deriving the patient recommendation, recursively. The process terminates, if all the premises are ground instances 40 (known facts). Figure 7 shows the justification trace for the earlier presented RPG 41 42 test-case.

1 CPG-EX IN ACTION

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3 How the CPG-EX works in practice will now be briefly described. The sequence of4 actions are:

- (i) A text-based CPG is input and pre-defined Domain Ontology and thePatient Ontology are loaded
- (ii) The CPG is encoded in terms of the CPG Ontology and the Domain Ontol ogy
- 9 (iii) The CPG rules are transformed to JENA rule syntax and passed to the JENA
 10 reasoning system
- (iv) The execution engine reasons with the given patient instance and the CPG
 knowledge to infer CPG-mediated recommendations
- (v) A justification trace of the inferred recommendations is generated to estab lish the physician's 'trust' in the recommendations offered by CPG-EX

The CPG-EX interface (Figure 8) is composed of three panels. On the left panel, 15 the user can load a text-based CPG. The middle panel displays the CPG Ontology 16 structure to enable the semantic annotation of the CPG using the CPG Ontology. 17 The CPG Ontology panel further provides the following features to the user: (i) The 18 Duplicate Button to duplicate selected tag/element in the CPG Ontology structure; 19 (ii) The Save Button to save the CPG Ontology structure into a file in RDF/XML 20 (Resource Description Framework/extensible markup) format; (iii) The Run Button 21 to execute the CPG using a given patient instance; and (iv) The Query Button to 22 23 generate a justification trace for a specific recommendation.

The right panel is used for assigning instances to each tag in the CPG Ontology. 24 It consists of Ontology Instances text box, Variable Name List and a Decision Variable 25 List. The Variable Name List displays all the properties stored in the Domain Ontol-26 ogy. The Variable Name List becomes active only when a variable.name tag (from 27 CPG ontology) is selected. The user can select a variable name from the Variable 28 Name List and assign it to a decision variable. The Decision Variable List shows the 29 list of annotated decision variables associated with their variable names (as shown 30 in Figure 4). The Query Button initiates a Query Window (as shown in Figure 7). 31 The Query Window enables a user to query patient information and inferred rec-32 ommendations. Furthermore, it allows the user to view the derivation trace for a 33 specific inferred recommendation. 34

CPG Authoring involves extracting textual information of the text-based CPG 35 (shown in the left panel of CPG-EX) and annotating them based on the classes and 36 their properties in the CPG Ontology (shown in the middle panel of CPG-EX). 37 The annotated text is assigned to the Ontology Instance Text Box (see right panel of 38 CPG-EX) based on the classes/properties in the CPG Ontology. Rule Authoring is 39 performed by defining decision rules in the logic tag of CPG ontology as follows: 40 Step 1: Select decision variables from the Decision Variable List, which repre-41 42 sents the body (premises) of the rule and followed by IF;



Figure 8. *The interface for the CPG-EX, showing the CPG text, CPG ontology and the CPG ontology instances*

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Step 2: Select the action variable from the Decision Variable List, which represents the head (conclusion) of the rule and followed by *THEN*;

Step 3: For each decision variable and action variable in the rule, an equal ity/inequality relation can be defined with either a variable, a value, a binary
 algebraic formula, another decision variable or list of decision variables (see
 Example A and Figure 4).

The rule authoring steps for the CPG rule described in Example A can be traced and the outcome of each step is as follows:

27 Step # 1 & Step # 2: IF dv1, dv2, dv3, dv4, dv5, dv6 THEN a1

28 Step # 3: IF dv1=?, dv2=?,dv3=notIndicatedRoutinely, dv4=?, dv5=?, dv6=?
 29 THEN a1=[dv2 dv6 dv3 dv4]

Finally, we tested CPG-EX using the EU Radiation Protection 118 – Referral Guideline for Imaging. We used the pre-defined RPG Ontology¹⁵, the Patient Ontology and the CPG Ontology to generate a series of CPG-mediated recommendations for the patient Ann (see Example A) as shown in Figure 7.

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35 DISCUSSION

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Computerisation of CPGs provides interesting opportunities to develop CDSS that
 provide evidence-guided recommendations. One advantage is that the CPG serves as

- 39 a validated knowledge resource and allows CDSS developers to avoid the perennial
- 40 knowledge engineering problem. By design, CPG follow a decision logic that is struc-
- 41 tured in an algorithmic format that can be used to generate explicit symbolic clini-
- 42 cal decision-support rules to suggest CPG-guided clinical recommendations. The

semantic web supports a logic-based framework that allows the semantic modelling 1 2 of medical knowledge that can be used to provide a variety of knowledge-mediated 3 services. In this paper, we have demonstrated the applicability of the semantic web to model CPG and leverage the CPG knowledge to develop ontology-based CDSS. We 4 5 have presented a unique approach that features the integration of multiple ontologies to develop a CDSS. We demonstrated the integration of two ontologies, each 6 representing the form and function of a CPG-the Domain Ontology describing 7 the CPG function and the CPG Ontology representing the CPG structure. We have 8 9 developed a simple CPG rule syntax that can be followed by medical practitioners to 10 write clinical decision rules based on the logic inherent with a CPG. The CPG rules are then transformed into a much complex rule syntax to enable their execution in a 11 powerful inferencing engine to infer recommendations and other information based 12 on patient profiles. Our CDSS approach is quite generic and can be extended to other 13 domains, provided the availability of a domain ontology. We tested our CPG-EX 14 15 with a number of real-life clinical cases and both the recommendations and their justifications were validated by medical practitioners. 16 17 18 REFERENCES 19 20 1 Russel IT, Grimshaw JM. Implementing clinical practice guidelines: can guidelines be used to improve clinical practice? Effective Health Care 1994; 8: 1-12. 21 2 Peleg M, Tu SW, Bury J, et al. Comparing computer-interpretable guideline models: a case-22 study approach. J Am Med Inform Assoc 2003; 10: 52-68. 23 3 Skonetzk Si, Gausepohl HJ,van der Haak M, Knaebel S, Linderkamp O, Wetter T. HELEN, a 24 modular framework for representing and implementing clinical practice guidelines. Meth-25 ods Inf Med. 2004; 43: 413-26. 4 Tu SW, Musen MA, Shankar R, et al. Modeling guidelines for integration into clinical work-26 flow. Medinfo 2004: 174-78. 27 5 Field MJ, Lohr KN (eds). Clinical Practice Guidelines: Directions for a New Program, Institute 28 of Medicine. Washington, DC: National Academy Press, 1990. 29 6 Berners-Lee T, Hendler J, Lassila O. The semantic web. Scientific American 2001; 284: 30 34-43. 31 7 Shiffman RN, Karras BT, Agrawal A, Chen R, Marenco L, Nath S. GEM: A proposal for a 32 more comprehensive guideline document model using xml. J Am Med Informatics Assoc, 33 2000; 7: 488-98. 8 Peleg M, Boxwala AA, Ogunyemi O, et al. GLIF3: The evolution of a guideline representation 34 format. AMIA Symposium 2000: 645-49. 35 9 Ciccarese P, Caffi E, Boiocchi L, Quaglini S, Stefanelli M. A guideline management system. 36 Medinfo 2004: 28-32. 37 10 Fox J, Johns N, Rahmanzadeh A. Disseminating medical knowledge: the proforma approach. 38 Artif Intell Med 1998; 14: 157-82. 39 11 Musen MA, Tu SW, Das A, Shahar Y. EON: A component-based approach to automation of protocol-directed therapy. JAMIA 1996; 3: 376-88. 40 12 Herre H, Heller B. Semantic foundations of medical information systems based on top-level 41 ontologies. Journal of Knowledge-Based Systems 2006; 19: 107-12. 42

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