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Development of a cardiac-centered frailty ontology

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Abstract

Background: A Cardiac-centered Frailty Ontology can be an important foundation for using NLP to assess patient frailty. Frailty is an important consideration when making patient treatment decisions, particularly in older adults, those with a cardiac diagnosis, or when major surgery is a consideration. Clinicians often report patient's frailty in progress notes and other documentation. Frailty is recorded in many different ways in patient records and many different validated frailty-measuring instruments are available, with little consistency across instruments. We specifically explored concepts relevant to decisions regarding cardiac interventions. We based our work on text found in a large corpus of clinical notes from the Department of Veterans Affairs (VA) national Electronic Health Record (EHR) database.

Results: The full ontology has 156 concepts, with 246 terms. It includes 86 concepts we expect to find in clinical documents, with 12 qualifier values. The remaining 58 concepts represent hierarchical groups (e.g., physical function findings). Our top-level class is clinical finding, which has children clinical history finding, instrument finding, and physical examination finding, reflecting the OGMS definition of clinical finding. Instrument finding is any score found for the existing frailty instruments. Within our ontology, we used SNOMED-CT concepts where possible. Some of the 86 concepts we expect to find in clinical documents are associated with the properties like ability interpretation. The concept ability to walk can either be able, assisted or unable. Each concept-property level pairing gets a different frailty score. Each scored concept received three scores: a frailty score, a relevance to cardiac decisions score, and a likelihood of resolving after the recommended intervention score. The ontology includes the relationship between scores from ten frailty instruments and frailty as assessed using ontology concepts. It also included rules for mapping ontology elements to instrument items for three common frailty assessment instruments. Ontology elements are used in two clinical NLP systems.

Conclusions: We developed and validated a Cardiac-centered Frailty Ontology, which is a machine-interoperable description of frailty that reflects all the areas that clinicians consider when deciding which cardiac intervention will best serve the patient as well as frailty indications generally relevant to medical decisions. The ontology owl file is available on Bioportal at http://bioportal.bioontology.org/ontologies/CCFO.

Keywords: Ontology, Frailty, Surgery, Cardiology, SNOMED-CT

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Background

Frailty and cardiac decision making

Frailty is an important patient attribute for treatment decisions in general [1–4] because assessing frailty severity predicts response to treatment and patient outcomes across many conditions [2, 5-10]. In the modern era of interventional cardiac care, patient frailty is increasingly important to decisions regarding major cardiac surgery and interventional procedures [1, 9, 11]. With the growing numbers of elderly and diabetic patients [6, 12], these decisions are common [13]. Older, frail patients with aortic valve stenosis can now be referred for coronary artery bypass graft surgery (CABG), a transcatheter valve replacement (TAVR), or medical management [14-16]. While TAVR is minimally invasive with shorter length of stay, frail patients may not necessarily benefit due to non-cardiac illnesses that limit quality of life or increase risk of procedural complications [11, 17], including increased length of stay, infection rates, and re-hospitalization. In 2015, the National Institute on Aging cited frailty assessment as a key priority in the perioperative approach to cardiac surgery [13].

Assessing frailty is done by intuitive estimates or appraisals, counting comorbid conditions, and the use of formal assessment instruments [2, 18, 19]. Frailty can include physical disability, deficits in mood, sensorium, and cognition, along with patient experience of pain or incontinence [3, 6].

The purpose of this paper is to describe the development of an ontology of frailty, paying special attention to how it relates to cardiac care decisions. Our ontology is designed to access the aspects of frailty that distinguish it from a simple count of comorbid conditions. We describe a necessary and sufficient view of patient frailty indicators apart from comorbid conditions. This ontology has been designed to allow computerized extraction of frailty information from the narrative documents patient records. Because frailty is a topic that is interpreted in many ways and measured with several instruments [5, 18-20], we built our ontology using as many term identification techniques as possible, gathering terms using existent instruments, physician interviews, and automated chart reviews. We aimed to allow cross walking between the measurement instruments. The hierarchical structure was adapted from SNOMED-CT [21] but expanded and informed by the nuances in clinical decision-making. We tested a draft version of our ontology by creating instrument-scoring rules, by using it to improve automated detection of frailty indicators in a Natural Language Processing (NLP) system, and by using it as an input feature to a system trained to predict patient mortality after a major cardiovascular procedure (MCVP) [22].

Frailty and NLP

Given that frailty information is so important, extracting it from clinical records is vital for patient care and research. The three methods of extracting information from clinical records are structured data, human chart review, or automated NLP systems. There are 3 reasons why an NLP approach is likely to be the most successful: 1) physicians do not consistently use frailty instruments, 2) there is no key, which reconciles scores across instruments, 3) they do not all use the same definition of frailty.

Clinicians collect frailty information, but not in a systematic fashion nor by consistently using frailty instruments [20]. They document narrative descriptions of frailty information that they find relevant to the specific clinical situation. It is possible that since clinicians believe they can rapidly use their clinical judgment to assess a patient's frailty when they see them [18], they do not feel the need to systematically use specific frailty instruments. Their narrative notations are considered sufficient. However, large-scale retrospective studies of patient outcomes require chart review, and if frailty is largely documented in narrative, then structured text cannot be used and the effort of wading through text in a chart causes a time bottle-neck for human reviewers.

The inconsistent use of frailty instruments would not matter for chart review based on structured information if there were a method for reconciling scores from different instruments. The method would create equivalences between the instruments. These equivalences would take as many factors into account as possible. Creating score equivalence metrics would be a task that humans would find challenging.

If clinicians all used a similar definition of frailty, humans chart review or NLP systems without ontology components would be able to locate their descriptions easily, but they do not. Clinicians' ideas about which patients are frail are influenced by both the culture within their organizational department, the decision at hand, and the wider society. For example, departmental culture may involve specific frailty tests (e.g., 6-min walk distance) and social culture may mean that frailty indicators have different thresholds (e.g., low body mass index (BMI) in Japan vs. the US [23].) Frailty indicators are also specific to each patient. A patient's level of mobility is highly dependent on prior exercise activities, desire for exercise, and the patient's personal preferences. The number of frailty instruments that have been developed evidences the variability in the conception of frailty, and therefore the complexity of the relationship between frailty and decision-making. Buta, et al. [20] identified 67 frailty instruments of which nine were cited more than 200 times.

Since current charting practices make human chart review or using structured data untenable, one could force a structured data solution by picking a single instrument and screen all patients. Picking a single instrument and screening everyone is hampered by the low specificity of current instruments [4, 20] and by lack of instrument adoption. In addition, frailty assessment varies substantially over time. Assessing all individuals over time would be necessary to understand the trajectory and implications of frailty [4], which would mean that the structured data solution would only become helpful after a significant time-interval. Systematically conducting frailty assessments at all encounters would fail to highlight decision-specific frailty issues, it would add a substantial burden to the clinician, and cost to the healthcare system.

Ontology building

Ontologies may be used in NLP projects to bridge structured data fields. Some structured data fields from clinical records across institutions or even within the same institution use different words to denote the same information (e.g., "patient name" vs. "lastname, firstname") [24–26]. Ontologies are also used for named entity recognition and decision modeling [27–29]. For example, named entity recognition can locate all mentions of disorders that patients may have as well as relevant patient demographics. Decision modeling uses either the named entities found or other inputs to access ontological elements, which contribute to creating rules or other models of decisions. Our ontology of patient frailty is designed to fulfill both purposes.

We employed the standard methodology for building ontologies including reconciling clinical text, medical literature, and existing ontologies [26, 30–32]. We chose to develop our ontology by adhering as closely as possible to realist principles. Realist principles lead to stable ontologies [33], which can be reasoned with while avoiding illogical inferences [34].

We conceptualized clinical records as textual recordings of the author's *ideas* about the patient. An existing ontological concept that corresponds to the author's ideas about the patient is *clinical finding* from the Ontology of General Medical Science (OGMS), which is defined as "A representation that is either the output of a clinical history taking or a physical examination or an image finding, or some combination thereof." [35] In contrast, the definition of a *clinical finding* used in SNOMED-CT is "observations, judgments or assessments about patients." The definition specifies that it is designed to convey "...the actual state of the body" and is inclusive of concepts with a semantic tag *disorder* (http://browser.ihtsdotools.org/). By referring directly to the patient's body and not the clinician's findings, one is

ignoring consideration of human error, cognitive biases, and other aspects that may influence patient-clinician and clinician-EHR interactions [36, 37]. However, SNOMED-CT's definition of *clinical finding* also includes concepts with the semantic tag *finding*, which "... are not separate from the observing of them," which brings them closer to the OGMS definition. We restricted ourselves to *findings*.

Integration with prior work

Two prior studies have successfully mined frailty information from rehabilitation and nursing home notes. One generated International Classification of Functioning, Disability and Health (ICF) codes and the other extracted Barthel index scores [38, 39]. Their work indicates that it is possible to locate and extract frailty-relevant terms. We expanded their work by increasing the number of frailty-related terms identified.

The UMLS Metathesaurus [40] contains a complex structure of frailty-related concepts. It is evident that SNOMED CT contributes concepts from many, if not all, of the available frailty instruments. However, the UMLS Metathesaurus is not realist due to long-standing requirements of backward compatibility [33, 41, 42]. We wanted our ontology to be interoperable with as many other ontologies as possible. We did not want to create something that entirely ignored the UMLS Metathesaurus. Recent papers have discussed realist approaches, specifically with respect to SNOMED-CT [33, 41, 43, 44]. Compatibility with SNOMED-CT can be used as a bridge to the UMLS Metathesaurus. Therefore, we incorporated SNOMED-CT concepts into the Cardiac-centered Frailty Ontology as often as possible, but did not limit ourselves to SNOMED-CT.

Objective

In this study we created a machine-interoperable description of frailty that reflects all the areas that clinicians consider when deciding which cardiac intervention will best serve the patient as well as general indications of frailty found in patient records.

Results

In this section we describe each of the four phases of ontology development (Identify other ontologies and official clinical tools, group terms into high-level classes, define attributes of classes, analyze and validate), which led to the final ontological structure.

Phase 1 – Collect terms by identifying other ontologies and official clinical tools

The research team met regularly to iteratively identify terms from a variety of sources. We reviewed 14 frailty instruments described in the methods section (below). The terms from these instruments were mapped onto the UMLS meta-thesaurus. If there was a SNOMED-CT term we used it.

In addition, we interviewed 12 clinicians (cardiologists, geriatricians, and cardiac surgeons) where we provided 7 hypothetical patient vignettes. Clinicians were asked to discuss patient frailty in relation to a decision between CABG, TAVR, and medical management. Each hypothetical patient had a different mix of frailty indicators. The full study will be described in a separate paper. The terms found in the interviews included muscle weakness, oxygen need, gait velocity, 6-min walk distance, volunteers, lower extremity strength, robust, functional status, functionality, deconditioned, acute vs. chronic findings and BMI. We identified terms at the level of concept granularity relevant to cardiac decision-making. For example, terms relating to housework (e.g., "dusting," "washing dishes," and "vacuuming") were grouped into a single concept ability to perform domestic activities because whether a patient is dusting or vacuuming is not relevant to their cardiac health.

In order to filter the terms into unique groups to aid the next step of creating hierarchies, the total set of terms underwent an initial sorting by the research team to identify explicit synonyms and concepts. The terms were found to correspond to the general categories of toileting, mental health, social functioning, working, exercise, walking, eating, general health, bathing, dressing & grooming, transfers, and modifiers (i.e., body locations and qualifiers). Using these groupings, we had an initial set of 108 unique concepts.

For those 108 concepts, we identified 198 unique concept-related terms. Terms within concepts were further expanded by SNOMED-CT synonyms and augmented by the team's previous experience with clinical documents. After the term expansion, nearly all of the concepts had either 1 or 2 terms, while one concept had 12 terms. The concept with the most terms was *Lack of energy finding*, with "lack of energy," "tired," "fatigue," "lack energy," "tiredness," "sleepiness," "drowsiness," "exhaustion," "exhaust," "wear out," "drain," and "weary."

Our clinician interviews made it clear that clinicians rely heavily on the Society of Thoracic Surgeons (STS) score in assessing patient's likelihood of surviving surgery. We examined the STS calculation and found it was more sensitive to comorbid conditions than indications of frailty. The clinicians also indicated that comorbid conditions are often included in assessments of patient frailty. Since we were interested in creating an ontology of frailty apart from comorbid conditions, we only included the concept *comorbid conditions count*, not specific conditions the patient might have.

A second term expansion was done using automated chart review. Terms found included "stagger" as indicative of the concept *impairment of balance finding*, "prosthetics," "shoes," "gripping strength," "fatigue," "weakness," "SOB," "short of breath," "dyspnea," "muscle strength," "motor strength," "decreased strength," "assist," "paralyzed," "handicap," "unassisted," "dresses," "bathes," and "stand." Some terms were removed because they commonly occurred with a meaning alternate to the one we were after. These terms included "dressing," "supine," "working," "eating," "strength," "incontinence."

After all terms had been gathered we had 246 terms associated with the 108 concepts.

Phase 2 - Group terms into high-level concepts

The identified frailty concepts were arranged in hierarchical relationship by mapping them to SNOMED-CT equivalents using SNOMED-CT concepts within the category *clinical finding*, with semantic type *finding*. The goal for this process was to restrict the SNOMED-CT mappings to as small a selection as possible, while maintaining correspondence, which means our ontology is somewhat compatible with SNOMED-CT.

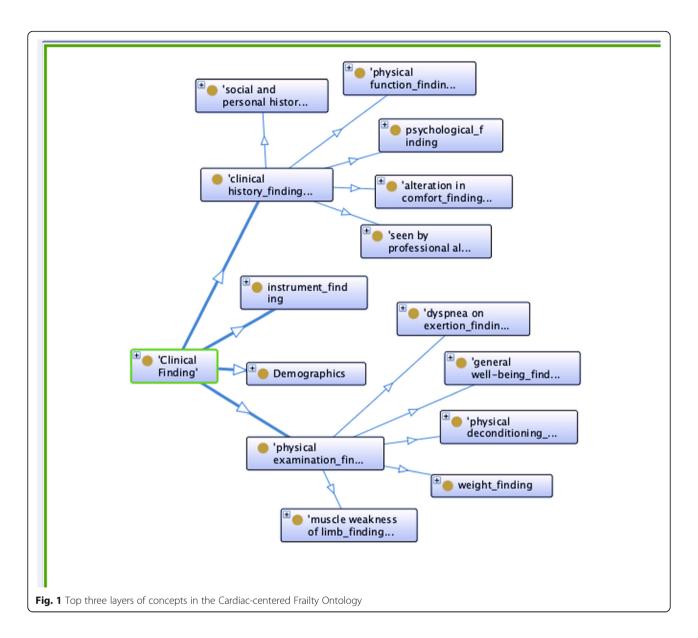
Figure 1 shows the top concepts in our Cardiac-centered Frailty Ontology. In order to create an ontology that is interoperable with SNOMED-CT, it is important that where concepts in the two ontologies share a name and an id number they are used to represent precisely the same portion of reality. Therefore, if we did not match the exact use described by SNOMED-CT, we explain why and do not use the SCTID.

As discussed in the introduction, we did not use *clinical finding* in the same way as SNOMED-CT, we restricted ourselves to the subset with semantic tag *finding*. Therefore, we did not use the SNOMED-CT ID number for the concept *clinical finding*. Our concept *clinical finding* (CCFOID:1) has children *clinical history finding* (CCFOID:11), *instrument finding* (CCFOID:12), and *physical examination finding* (CCFOID:13), reflecting the OGMS definition of clinical finding. *Instrument finding* is any score found for the existing frailty instruments already mentioned. We included classes not mapped to SNOMED-CT for *demographics* (CCFOID:14) and *qualifier values* for the properties of our concepts in our top level.

In the Cardiac-centered Frailty Ontology we included *demographics* in *clinical findings* because we are referring to demographic information collected by the clinician at a clinical visit, not to the demographic information that inheres in the patient and may change between visits.

Clinical history finding

The obvious choice for findings arising from the patient's clinical history taking is *clinical history and observation finding (finding)* (SCTID: 250171008). It turns



out that all of the children listed in the SNOMED-CT browser (http://browser.ihtsdotools.org) have semantic type finding, which is what we were after. Like clinical finding, we do not want to include all the children of clinical history and observation finding, which means we are not referring to the same portion of reality. Synonymous terms and the SNOMED-CT term identification number (SCTID) are not necessary because we are not looking for the concept to be represented in clinical documents. For these reasons, we shorten the name to clinical history finding and exclude the SCTID.

The same problem arises when we try to find a SNOMED-CT equivalent of *physical function finding* (CCFOID:112). The topic modeling, our interviews with clinicians, and existing instruments all indicate that the patient's physical abilities are a necessary

category. The closest SNOMED-CT equivalent is functional finding (finding) (SCTID: 118228005), which has among its children concepts we need, for example finding of activity of daily living (finding) (SCTID: 118233009, CCFOID:1124). However, it includes findings unrelated to physical abilities like does comply with treatment (SCTID: 386673006). Therefore, we do not use functional finding and leave physical function finding, with no SCTID.

SNOMED-CT is so exhaustive that there can be hierarchical structure that is beyond our needs. *Social and personal history finding* (SCTID: 365448001, CCFOID:115) has two intervening problem parents *finding by method* and *history finding*, which lead to *clinical finding* and not *clinical history and observation finding*. There is no indication in the documentation

about how history finding differs from clinical history and observation finding. The children of Social and personal history finding and psychological finding (SCTID: 116367006, CCFOID:113) that we are interested in also have intervening concepts. It may be better practice to just use the SCTIDs for the lowest levels. Those will be the only ones used in the NLP.

Alteration in comfort finding (SCTID: 130979001, CCFOID:111) has the intervening parent problem sensory nervous system finding (finding) (SCTID: 106147001) and neurological finding (finding) (SCTID: 102957003), which go to clinical finding (finding). The courses qualifier value (CCFOID:43) concept we include corresponds to the SNOMED clinical finding attribute clinical course. We also include a concept for seen by a professional allied to medicine finding because the snippet annotations indicated that being seen by physical therapy, occupational therapy or other allied professions indicated patient frailty.

Physical examination finding

We included in *Physical* Examination Finding (CCFOID:13) concepts that fall in the SNOMED-CT hierarchy under general findings of observation of patient (finding) (SCTID: 118222006). In SNOMED-CT general finding of observation of the patient is a child of clinical history and observation finding. Since we did not include observation in our concepts, we included this separate concept for physically observing the patient. We took a very restricted subset of the children of general findings of observation of patient (finding), hence the name change and absence of SCTID. The children we included are physical deconditioning finding (SCTID: 31031000119102, CCFOID:134), dyspnea on exertion finding (SCTID: 60845006, CCFOID:131), muscle weakness of limb finding (SCTID: 713514005, CCFOID:133), weight finding (SCTID: 107647005, CCFOID:135), and general well-being finding (SCTID: 365275006, CCFOID:132).

Physical deconditioning has no children. We included all of the children of muscle weakness of limb because they separate upper and lower limbs, which our interviews indicated is an important distinction. Weight finding has many irrelevant children including finding of color zone for Broselow Luten pediatric weight estimation (finding). We did not include these children.

For dyspnea on exertion finding, and General well-being finding, we kept the SCTID because we could map all of the children, although this is not currently part of the ontology. For the children not currently explicitly listed, we would need to determine whether they were indicative of high or low frailty. We added the concept comorbid condition count finding (CCFOID:131), which we discussed earlier, as a child of general well-being finding.

The full ontology has 156 concepts, with 246 terms. The ontology owl file is available on Bioportal at http://bioportal.bioontology.org/ontologies/CCFO. We consider CCFO a "view" into SNOMED-CT. We define "view" in accordance with the Ontology Views Project being done by the Structural Informatics Group at Washington University (http://sig.biostr.washington.edu/projects/ontviews/). In this definition a view is a new ontology that includes some portion of the viewed ontology. CCFO contains portions of SNOMED-CT. It is therefore a view of SNOMED-CT. As a view, it falls under SNOMED-CT's existing licensure (https://www.snomed.org/snomed-ct/get-snomed).

Table 1 shows the number of concepts by their number of terms. The table lists the number of terms associated with each of the 86 concepts we expect to find in clinical documents. The remaining 58 concepts, not in the table, represent hierarchical groups (e.g., *physical function findings*) and 12 *qualifier values*. Two concepts from *demographics* (CCFOID:14) have no terms (*patient age finding*, CCFOID:141 and *indeterminate sex finding*, CCFOID:1422). *Lack of energy finding* (CCFOID:132222) still has the highest number of terms.

Table 2 lists some important concepts and their associated terms. The term list is included as Additional file 1. Since terms are not synonyms for the concepts in the ontology, they are not included in the ontology itself. Terms are text that we consider indicative of the author's thoughts about the concept. Concepts themselves are portions of reality, not pieces of text.

Phase 3 - Define object properties for concepts

Concept properties were determined by rating scales used in the instruments. Activities have a frequency property that is found in the SNOMED-CT frequency qualifier value (SCTID: 272123002, CCFOID:44) restricted to high frequency qualifier value (SCTID: 27732004, CCFOID:441) and mid-frequency (SCTID: 255218000, CCFOID:442). Frequency values contrast with a value of absent finding qualifier value (SCTID: 272519000, CCFOID:42). Possible values are restricted based on the likelihood of finding specific text qualifiers. Abilities have an ability interpretation property that is found in ability interpretation qualifier value (SCTID: 371148001, CCFOID:41). These values are also restricted to able qualifier value (SCTID: 371150009, CCFOID:412), able with difficulty qualifier value (SCTID: 371157007,

Table 1 Breakdown of the number of terms per concept in the Cardiac-centered Frailty Ontology. These counts are for the 86 concepts that we expect to find in clinical documents

# terms	1	2	3	4	5	6	7	8	> 8
# concepts	24	29	7	8	6	5	0	3	4

Table 2 List of concepts central to assessing frailty and their associated terms. Terms are not synonymous with the concept or the concept name. They indicate author may have been thinking about the concept. Bolded terms were not found in the topic modeling paper. Underlined terms were added by the annotation task

Concept	Terms (not synonyms)			
ability to run finding	Difficulty running; able to run; unable to run; run			
ability to stand finding	Difficulty standing up; unable to stand uable to stand up; stand up			
able to mobilize finding	ambulate independently; steady gait; unsteady gait			
bed-ridden finding	bed-ridden; supine; stretcher			
Paralysis finding	paralysis; paralyzed			
wheelchair bound finding	wheelchair; scooter; w/c; wheel chair			
able to perform dressing activity finding	dresses; Able to dress; independent wit dressing; Needs help with dressing; Dependent for dressing; unable to dres Difficulty dressing; shoes; ties shoes;			
able to perform personal grooming activity finding	Able to wash own hair; Unable to wash own hair; Difficulty washing own hair; clean appearance; personal grooming; neatly dressed; well-groomed; well-groomed without assistance; good personal hygiene			

CCFOID:411) and unable qualifier value (SCTID: 371151008, CCFOID:413). Finally, all clinical findings have a course property from courses qualifier value (SCTID: 288524001, CCFOID:43), including chronic qualifier value (SCTID: 90734009, CCFOID:431), clinical course with short duration qualifier value (SCTID: 424572001, CCFOID:432), and sudden onset qualifier value (SCTID: 385315009, CCFOID:433).

More properties of the concepts were determined by scores of relevance to cardiac decisions and their likelihood of resolving after the recommended intervention. Three investigators and three interview participants scored 81 of the 84 concepts that we expected to find in clinical documents. Three concepts were added after scoring was complete (ability to drive a car finding CCFOID:11241, quadriceps weakness finding CCFOID:1334, and calf weakness finding CCFOID:1331).

For the 81 concepts that were scored, ability concepts were qualified with the *able qualifier values* (able/independent, with difficulty/assisted, and unable/dependent) each concept-value pair was given a separate score. Activity and mental state concepts were qualified with *frequency qualifier values* (high frequency, mid-frequency, absent) and scored seperately. Rockwood categories as described in the Dalhousie University Clinical Frailty Score [45] were averaged across the eight raters. Ratings of low, medium, or high for relevance to frailty and fix-ability where set to the majority rating for the six raters, who had clinical experience.

Only three concepts were given low relevance to frailty ratings by all six raters *calm finding* (CCFOID:113331), happy finding (CCFOID:113332), and nervous finding (CCFOID:113333) concepts from the mental state finding (CCFOID:11333) concept. Fifty-two concepts were rated as highly relevant by all six raters, nine by at least three raters. Thirteen concepts had relevancy ratings of medium by all six raters, four by at least three raters. Table 3 shows the findings for nine concepts central to the assessment of frailty. Ability to participate in leisure activities finding (CCFOID:112412) is included in Table 3 to demonstrate a cardiac intervention-specific concept.

Table 3 Scores for nine concepts central to the assessment of frailty. Rockwood scores are on a scale of 1 - very fit to 9 - terminally ill. They are averaged across raters. "Will fix" refers to clinical findings that the cardiac intervention will alleviate. Relevance is how important the concept is to decisions about cardiac interventions. L - low, m - medium, h - high

Concept	Rockwood				Relevance	
	Able	With Difficulty	Unable			
ability to run finding	1	3.33	4	М	Н	
ability to stand finding	2.67	5.11	7.44	TIED L+	Н	
able to mobilize finding	steady gait 3		unsteady gait 6	М	Н	
bed-ridden finding	Only level 8			L	Н	
Paralysis finding	Paraplegic 6		Quadriplegic 8	L	TIED M+	
wheelchair bound finding	Only level 6			М	Н	
able to perform dressing activity finding	3	4	7	L	Н	
able to perform personal grooming activity finding	1	4	7	L	Н	
ability to participate in leisure activities finding	2	3	5.5	Н	Н	

We determined which concepts are specific to cardiac intervention decisions by using the difference between ratings for will fix and relevance (in Table 3). Will fix refers to findings the cardiac intervention will alleviate; while relevance refers to findings that our reviewers indicated were relevant to frailty. We considered concepts that are highly relevant to frailty and are either highly likely to be alleviated by cardiac intervention or are associated with eventual recovery, to be especially important. For instance, bed-ridden is seen as generally relevant and not specifically relevant to cardiology. Enjoys light exercise finding (CCFOID:11231), ability to participate in leisure activities finding (CCFOID:112412), dyspnea on exertion finding (CCFOID:131), and fit and well finding (CCFOID:1323) are all rated as specifically relevant to cardiology as well as being generally relevant. Thirty-two concepts are rated as moderately specific to cardiology and 45 were given low cardiology-specific ratings.

In this section, we also looked at the mapping instrument scores found in the clinical document set to frailty scores from Rockwood categories as described in the Dalhousie University Clinical Frailty Score [45]. Table 4 lists the instruments and their scoring criteria.

Phase 4 - analyze and validate

We created implementation rules to map ontology elements to instrument questions for three common instruments. The rules for these three instruments (Barthel index, Katz ADLs, and SF-36) are listed in Table 5. Note that the mappings are not one-to-one. Some of the instrument questions were mapped to equivalent concepts. For example, both Barthel index and Katz ADLs uses the parent concepts *ability to perform personal care activities* (SCTID: 284774007) to include feeding self, dressing, grooming, toileting, and washing oneself, and *ability to transfer location* (SCTID: 714882001). By creating implementation rules, we were able to demonstrate that the Cardiac-centered Frailty Ontology covered the topics used in the instruments.

In addition, we conducted preliminary NLP analysis using the ontology. We wanted to determine if narrative text that included frailty terms also included enough information to determine whether the patient had frailty-related functional deficits. Frailty terms from the Cardiac-centered Frailty Ontology and from a prior study [46] were used. We extracted 2460 clinical record snippets centered at the frailty keyword terms. Three

Table 4 Instrument scores found in clinical document set and the scoring criteria, which allow the NLP system to use the scores to determine indication of frailty

Instrument Name	Scoring Criteria				
Activities of Daily Living (ADL) Screen	18 patient independent 6 patient very independent				
Functional Independence Measure (FIM)	7-complete independence; 6-modified independence; 5-Supervision or step-up; 4-Minimal Contact Assistance;	3-Moderate Assistance; 2-Maximal Assistance; 1-Total Assistance			
Katz index ADL	Score of $6 = \text{High}$, Patient is independent. Score of $0 = \text{Low}$, patient is very dependent.				
Barthel index	ADL: 70–100 = Independent; Less than 70 = Needs significant physical/supervisory assistance.				
Instrumental activities of daily living (IADL)	2 = without assistance, 1 = with assistance, 0 = unable				
Instrumental activities of daily living (IADL) scale (Lawton) / IADL Screen	The total score may range from 0 to 8. A lower score indicates a higher level of dependence.				
Functional Activity Questionnaire (FAQ)	Score of 5 or more indicates significant impairment in instrumental activities of daily living.				
Morse fall scale / Annual Fall Scale / MRT	> = 45: high fall risk 25–44: moderate risk 0–24: low risk				
Tinetti assessment measures	Maximum possible balance score: 16 points. Maximum possible gait score: 12 points. Maximum total score: 28 pointsScores below 19 indicate high risk for falls. Scores in the 19–24 range indicate some risk for falls.				
Braden scale	Pressure Ulcer Risk: total score 13–14 mode total score <=9 very high risk total score 15–18 mild r total score 10–12 high risk total score 19–23 no risk				

Table 5 Examples of Frailty Instruments implemented with the Cardiac-centered Frailty Ontology

Frailty Insrument: Barthell Index *Incontinence finding* (CCFOID:11271) of either kind = 0 Continence finding (CCFOID:11271) or absent qualified incontinence finding of both kinds = 2 Able to perform personal care activities finding (CCFOID:11244) for each of its children: unqualified or able = 2with difficulty = 1unable = 0Ability to transfer location finding (CCFOID:1122)(any one) = 2Absent (CCFOID:42) qualified = 0Able (CCFOID:412) qualified able to mobilize finding (CCFOID:112121) or no aid for walking finding (CCFOID:112472) = 3 Able with difficulty (CCFOID:411) qualified able to mobilize finding or walking aid use finding (any kind) = 2 wheelchair bound finding (CCFOID: 1121223) = 1 unable (CCFOID:413) qualified able to mobilize finding or bed-ridden finding (CCFOID:1121221) = 0 Able qualified able to walk upstairs finding (CCFOID:1124713) or able to walk downstairs finding (CCFOID:1124711) = 2 Unable qualified able to walk upstairs finding = 0 Barthell Index Scoring Add up the score: 20 = no disability 0 = complete disability Frailty Instrument: Katz - ADLs Count the number of: Each of the able qualified ability to perform personal care activities finding (bathing, dressing, toileting, feeding)(CCFOID:11244) = 1 *Unable* qualified = 0Any able qualified ability to transfer location finding (CCFOID:1122) = 1 Unable qualified = 0Both unqualified continence finding or absent qualified incontinence finding of either kind = 1 Unqualified *Incontinence finding* of either kind = 0 Katz - ADLs Scoring Add up the score: 6 = high functioning 0 = low functioningFrailty Instrument: SF-36 Average the following for General Health score: Questions 1, 33, 34, 35, 36 First assessment covers 5 questions, 1 score. unqualified or high frequency (CCFOID:441) qualified fit and well finding = 100 mid-frequency (CCFOID:442) qualified fit and well finding = 75 absent qualified generally unwell finding (CCFOID:1324) = 50 mid-frequency qualified generally unwell finding = 25 unqualified or high frequency qualified generally unwell finding = 0 Ouestion 2 unqualified or high frequency qualified fit and well finding with sudden onset (CCFOID:433) qualification = 100 mid-frequency qualified fit and well finding with sudden onset qualification = 75 absent qualified generally unwell finding = 50 generally unwell finding: mid-frequency qualified = 25 unqualified or high frequency = 0 Average the following for Pain score Question 21 absent qualified alteration in comfort: pain finding (CCFOID:1111) = 100 mid-frequency qualified alteration in comfort: pain finding = 50 high frequency qualified alteration in comfort: pain finding = 0 Question 22 absent qualified alteration in comfort: pain finding, with able qualified able to carry out daily routine finding (CCFOID:11245) = 100 mid-frequency qualified alteration in comfort: pain finding, with able qualified able to carry out daily routine finding = 75 high-frequency qualified alteration in comfort: pain finding, with able qualified able to carry out daily routine finding = 50 mid-frequency qualified alteration in comfort: pain finding, with difficulty qualified able to carry out daily routine finding = 25 high-frequency qualified alteration in comfort: pain finding, with unable or with difficulty qualified able to carry out daily routine finding = 0 Average the following for Physical Functioning score: Question 3 high frequency qualified enjoys vigorous exercise finding (CCFOID:11233) or able qualified ability to run finding (CCFOID:11213) = 100 mid-frequency qualified enjoys vigorous exercise finding = 50 gets no exercise finding or unable qualified ability to run finding or absent qualified enjoys vigorous exercise finding = 0 high frequency qualified enjoys moderate exercise finding (CCFOID:11232) = 100 mid-frequency qualified enjoys moderate exercise finding = 50 gets no exercise finding or unable qualified ability to run finding or absent qualified enjoys moderate exercise finding = 0 able qualified ability to perform general purpose physical activity finding (CCFOID:11243) or able qualified ability to perform shopping activities finding (CCFOID:112413) = 100

Table 5 Examples of Frailty Instruments implemented with the Cardiac-centered Frailty Ontology (Continued)

with difficulty qualified ability to perform general purpose physical activity finding or with difficulty qualified ability to perform shopping activities finding =

unable qualified ability to perform general purpose physical activity finding or unable qualified ability to perform shopping activities finding = 0 Ouestion 6 & 7

This covers 2 question (scores twice) able qualified able to walk upstairs finding = 200

with difficulty qualified able to walk upstairs finding = 100

unable qualified able to walk upstairs finding = 0

Question 8

able qualified able to kneel finding (CCFOID:11211) = 100

with difficulty qualified able to kneel finding = 50

unable qualified able to kneel finding = 0

Questions 9-11

This covers 3 question (scores three times) able qualified able to walk finding (CCFOID:112471) = 300

with difficulty qualified able to walk finding = 200

unable qualified able to walk finding = 0

Ouestion 12

able qualified ability to perform personal care activities finding = 100

with difficulty qualified ability to perform personal care activities finding = 50

unable qualified ability to perform personal care activities finding = 0

Average the following for Role Limitations due to Physical Health score:

Ouestion 13-15

This covers 3 question (scores three times) absent qualified occupational maladjustment finding (CCFOID:1154) = 300

mid-frequency qualified occupational maladjustment finding = 150

high-frequency qualified occupational maladjustment finding = 0

Question 16

able qualified able to carry out daily routine finding = 100

with difficulty qualified able to carry out daily routine finding = 50

unable qualified able to carry out daily routine finding = 0

Average the following for Role Limitations due to Emotional Problems score

Question 17-19

This covers 3 question (scores three times) absent qualified occupational maladjustment finding and any psychological finding (CCFOID:113) = 300

mid-frequency qualified occupational maladjustment finding and any psychological finding = 150

high-frequency qualified occupational maladjustment finding and any psychological finding = 0

Average the following for Energy/Fatigue score

Questions 23, 27, 29, 31 (1 score)

able qualified able to sustain energy level finding (CCFOID:132221) or absent qualified lack of energy finding or absent qualified fatigue = 100 with difficulty qualified able to sustain energy level finding or mid-frequency qualified lack of energy finding or mid-frequency qualified fatique = 50 unable qualified able to sustain energy level finding or high frequency qualified lack of energy finding or high frequency qualified fatigue = 0

Average the following for Emotional Well-Being score Ouestions 24 26

This covers 2 question (scores twice) high frequency qualified calm finding or absent qualified nervous finding or absent qualified anxiety diagnosis (CCFOID:21) = 200

mid-frequency qualified calm finding = 150

mid-frequency qualified nervous finding = 100

absent qualified calm finding = 50

high frequency qualified nervous finding or anxiety diagnosis = 0

Questions 25, 28, 30

This covers 3 question (scores three times) high frequency qualified happy finding or absent qualified sad finding (CCFOID:113334) or absent qualified

depression diagnosis (CCFOID:22) = 300

mid-frequency qualified happy finding = 225 mid-frequency qualified sad finding = 150

absent qualified happy finding = 75

high frequency qualified sad finding or depression diagnosis = 0

Average the following for Social Functioning score

Ouestion 32

able qualified ability to perform community living activities finding (CCFOID:11241) = 100

with difficulty qualified ability to perform community living activities finding = 50

unable qualified ability to perform community living activities finding = 0

absent qualified impaired social interaction finding (CCFOID:11531) = 100

mid-frequency qualified impaired social interaction finding = 50

high frequency qualified impaired social interaction finding = 0

Scores are from 0 to 100 for each section, higher score = less frail/better health

clinicians and two informatics researchers reviewed the snippets. They categorized them as: a) Yes Deficit, or b) other. We trained a classifier on the snippets using a support vector machine (SVM). The average SVM performance, using 10-fold cross validation, achieved an accuracy score of 80.5%. Since frail patients typically have multiple frailty descriptions, the accuracy was deemed to adequately indicate that the terms in our ontology could be used to focus a learning system on frailty-relevant clinical text.

Finally, in [22] we tested whether the ontology could be used to help train a system to predict mortality for heart failure patients who underwent a major cardiovascular procedure (MCVP). We collected 2-years of clinical history data for a cohort of 20,000 heart failure patients leading to the MCVP. Frailty terms were identified in the text and classified as asserted or negated (i.e., "yes deficit" or "other") using NLP. The ontology was used to map identified terms to their concepts. This study used an early draft of the ontology that had only 7 higher-level concepts: therapy, medical findings, exercise, mobility, living activity, self-care and social function. These concepts became clinical findings; seen by professional allied to medicine, physical examination finding, activity exercise pattern, ability to move, activity of daily living, eating, feeding drinking ability, and social and personal history finding, respectively. We aggregated the frailty concepts by group and selected maximum frailty score from among the concepts in each group.

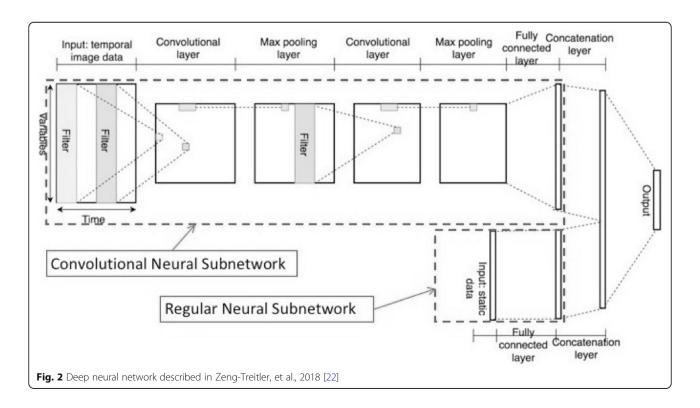
A deep neural network (DNN), pictured in Fig. 2, was trained on a visual representation of the data features, which were hospitalizations, ICD9 codes for diagnoses, medications, and the frailty score. In ten-fold cross validation, the area under the curve (AUC) for mortality prediction was 78.3% (95% CI 77.1 to 79.5%) on the test

data for the DNN model. We view this as additional validation for the ontology.

Discussion

We developed and validated the Cardiac-centered Frailty Ontology. We created our own hierarchy to allow removal of unnecessary layers, unnecessary concepts, and maintain realist design principles as much as possible. We used SNOMED-CT concepts for all of the lowest level concepts. We incorporated 14 existing instruments in our initial development. We added five more for scoring and rule sets, when analysis of a 400-document clinical document set showed these instruments were in common use [20]. We adapted the standard ontology development model [32] by using clinician interviews to identify important concepts, without the necessity of forcing clinician agreement.

Our ontology development techniques differed from the standard techniques in two ways. We used vignette-guided interviews in lieu of a subject matter expert meeting to gain consensus and used validated frailty assessment instruments in lieu of the frailty literature. Interviews allowed us to determine concepts to assess patient frailty based on specific clinical decisions. That our participants came from different institutions helped minimized institution-related medical-cultural bias in frailty assessment. By including concepts that any one group might have excluded, we retain the chance that our NLP system will find all relevant



concepts. Since the research group determined the concepts to include, we needed to be sure that we were not injecting our own opinions. Based on the amount of previous work in the area and existing SNOMED-CT concepts we included, we felt that our choices were not influenced by our opinions. We had our participants separately rate concepts for frailty severity, chance of cardiac intervention alleviating the problem, and relevance to frailty. By returning to the participants, we made explicit the extent and nature of their disagreements. We can then use this information going forward.

To address the concepts of frailty related cardiac interventions specifically; we included the concepts found in our interviews that we had not found from other sources. Quadriceps weakness finding is particularly relevant to cardiac intervention decisions because post-surgical patients cannot use their arms to help themselves stand. Surgical incisions require the upper body not be used. Therefore, if the patient cannot stand using their quadriceps alone, their post-surgical mobility is impaired, which impedes healing. Our participant ratings show that only a few concepts are specifically relevant to cardiac decisions, while around half are generally relevant, but not specifically relevant to cardiology. All four of the cardiac-specific concepts were also considered highly relevant to general assessments of frailty. Assessment of the utility of these cardiac-specific concepts in predicting patient outcomes was piloted as part of two studies to predict patient mortality [22, 47].

The Cardiac-centered Frailty Ontology reflects general frailty assessment as implemented in the frailty instruments used in its development [2, 48–58]. It includes participant ratings and our separate analysis of the interview data to create a picture of clinical decision-making with respect to cardiac interventions. Based on our orientation toward surgery-related decision-making [8–11, 13–17], we have excluded some of the specificity required to make non-surgery-related frailty decisions. We grouped all types of *lack of energy findings* together even though difference between "tiredness" and "weariness" may be important in other contexts. Once we have the NLP system functioning, it will be important to assess differences in outcome prediction using STS scores, with and without Cardiac-centered Frailty Ontology concepts.

We used a realist ontology development process [59] because it appeals to our understanding of the world, it helps ensure that the ontology is stable, and it avoids illogical inferences. By separating concept name and term list, we allow for language evolution, because the way *terms* are used changes over time. However, the portions of reality denoted by the concept and the concept name do not change.

Concepts refer to *Representations* in the minds of clinicians. These *representations* are far richer than the

terms used to indicate their presence in the mind of an author. *Representations* are multi-modal. They include memories and imaginings, relevance to goals, and other information value attributes.

We are looking for clinical *findings*, which are conclusions drawn by clinicians and recorded in narrative form [60]. Restricting ourselves to *findings* also minimizes problems with illogical inferences. Take for example the concept *enjoys light exercise finding*, the truth of this as a conclusion drawn by a clinician is unchanged by whether or not the patient "enjoys" the process of exercising or whether or not the patient actually exercises. That it is an *activity exercise pattern finding* also remains a valid inference.

Our main focus was the findings noted in narrative text documented during clinical care, i.e., clinical history findings. We recognize that comorbid conditions are relevant to frailty assessment, but there are existent tools for identifying comorbid conditions. Clinical history findings represent the frailty-specific information we are interested in automatically extracting from clinical documents. We included a very restricted subset of findings from physical examination. We tested the comprehensiveness of our coverage by creating scoring rules for the frailty assessment instruments. If concepts were missing, we would not be able to create appropriate rules. We assessed the relevance of each concept by asking participants to rate them as high, medium, or low in relevance to assessing frailty with respect to cardiac intervention decisions. A preponderance of low relevance ratings would indicate a problem. We found only three. Three quarters of the concepts were rated as highly relevant. Taken together these results indicate that we have covered the necessary and sufficient concepts related to frailty assessment.

One of the best qualities of both SNOMED-CT [21] and the UMLS Metathesaurus [40] is their exhaustive coverage of the medical domain. One would be hard pressed to find a medical concept that was not contained within them. This exhaustiveness creates problems when we try to use them in NLP applications. Simple matching to either vocabulary results in too many false positives. The Cardiac-centered Frailty Ontology creates a comprehensive picture of frailty, while limiting the concepts from SNOMED-CT to only those directly relevant. We used concepts, with semantic type *finding*, found by human review of frailty assessment instruments, physician interviews, and chart review.

Limitations

The main limitation of this work is the influence imparted on the ontology by our own ideas and biases. This limitation is shared by all ontologies. Our personal bias was minimized by the inclusion of the current accepted validated instruments on frailty. Each instrument reflects both expert consensus on the relevant concepts and empirical evidence of validity.

As is the case with all ontology development for NLP, ontologies precede NLP systems. The clinical outcome prediction NLP system used in our validation was not designed to model clinician decision-making. Without having a decision-making NLP system, it is difficult to assess whether the Cardiac-centered Frailty Ontology will facilitate all of the outcome predictions that we envision.

Conclusions

We developed and validated a Cardiac-centered Frailty Ontology. The ontology is a machine-interoperable description of frailty that reflects all the areas that clinicians consider when deciding which cardiac intervention will best serve the patient. It was designed to share as many elements as possible with SNOMED-CT to allow interoperability. It could not be simply a subset of SNOMED-CT because there was no appropriate subset for us to choose.

Methods

We used the ontology development process described in Noy, et al. [32]. This process consisted of four phases. Phase 1 used existing ontologies and official clinical tools to identify individual terms. The clinical tools we used were validated frailty instruments and automated chart review. We expanded this to term based on physician interviews. In Phase 2 we grouped terms into high-level concepts. We did this by examining concepts and hierarchies found in the existing SNOMED-CT ontology, while keeping the structure compact and realist. In Phase 3 we defined object properties for concepts. Our methodology included mapping concept attributes from scoring collected for the identified concepts and properties indicated by instrument questions. Instruments have an associated property, which indicates a mapping between instrument scores and our ontology's concept frailty scores. For Phase 4, analysis and validation, we created implementation rules for using the Cardiac-centered Frailty Ontology to reconcile scores on three common frailty instruments. Ontology structure was developed in Protégé [61], while term mappings were kept and shared in a Google sheet.

Phase 1 – Aggregate terms form other ontologies and validated clinical tools

To extract frailty concepts from existing instruments, five members of the research team reviewed the specific items from 14 instruments chosen by the number of times they were cited and expert recommendation [20]: (1) Physical Frailty Phenotype (PFP, also called CHS frailty phenotype) [2]; (2) SF-36 [48]; (3) FIM [49]; (5) Clinical Frailty Scale [50]; (6) Brief Frailty Instrument [62]; (6) the Barthell Index [51]; (7) Health Assessment Questionnaire (HAQ) [52]; (8) PSMS [53]; (9) Katz ADL [54]; (10) Duke Activity Index [55]; (11) RDRS [56]; (12) FACIT [57]; (13) NYHA [58]; (14) Deficit Accumulation Index (DAI, also called Frailty Index) [63]. Each person reviewed each individual item from each instrument. Terms from the World Health Organization's International Classification of Functioning, Disability and Health (ICF) were also included in the analysis because the instruments varied in their levels of abstraction, their scopes, and uniqueness.

At this step we added an ontology entry for comorbid condition count. Comorbid conditions are an important indicator of frailty. However, they are not the focus of our investigations. We focused on frailty-specific indicators in order to identify core frailty concepts in clinical documents.

The concept list was expanded by the findings of the interviews of cardiologist and cardiac surgeons described above.

Finally, we included terms extracted from manual note review by members of the research team with clinical experience. These reviews were in preparation for NLP topic modeling by Shao, et al., (2016). They reviewed clinical notes and social media posts [64].

Phase 2 - group terms into high-level concepts

We organized constructs in hierarchical relationship based on: 1) the results of topic modeling, 2) the basic organization of the frailty instruments, and 3) by mapping them to SNOMED-CT equivalents. We used SNOMED-CT concepts within the category *clinical finding*, with semantic type *finding*. The goal for this process was to restrict the SNOMED-CT mappings to as small a selection as possible, while maintaining correspondence with groupings from topic modeling and instruments, which means our ontology is somewhat compatible with SNOMED-CT.

Phase 3 - Define object properties for concepts

Object properties were determined in two ways, through the scales used to answer instrument questions and by scoring terms and concepts based on key decisions when making cardiac surgery decisions. For *instrument findings*, we defined properties, which related instrument scores to the Rockwood global assessment of frailty (described below) [45].

Rockwood categories are described in the Dalhousie University Clinical Frailty score, which has 9 categories [45]. The categories are (1) very fit, (2) well, (3) managing well, (4) vulnerable, (5) mildly frail, (6) moderately frail, (7) severely frail, (8) very severely frail, and (9) terminally ill [50]. Concepts in the ontology vary in how they map onto these severity categories. Some concepts have three levels of severity of impairment (able, assisted, and unable). The concept ability to walk, for example, has these three levels, where each level indicates a different Rockwood category. The scales provided for question answering in the frailty instruments indicated concept severity levels. These scales took the form of able to unable and all the time to never.

To establish the relationship of the concepts to these aspects of frailty three members of the team (BB, CW, KDH) and 3 participant cardiologists scored the concepts on three key decisions identified in the interviews: 1) Rockwood category (described below) as an indicator of ability to survive surgery, 2) relevance to cardiac decision-making as a reflection of the patient's ability to recover from surgery, and 3) the likelihood that the cardiac intervention will fix the problem.

Another outcome of our physician interviews was that clinicians consider indications of frailty within the context of cardiac decisions by assessing whether they are likely to be a result of the patient's cardiac condition and whether they are specifically relevant to cardiac decisions. The medically trained members of this group also characterized the constructs as low, medium, high for both their likelihood to be fixed by cardiac intervention and their relationship to cardiac decision-making.

For instrument scores, we created score rating for 10 commonly cited instruments that were not included in the initial concept-finding step. The initial mapping was created by author YC and verified by the remaining authors.

Phase 4 - Analyze and validate

For this phase, we created rules to implement three frailty assessment instruments using the Cardiac-centered Frailty Ontology. We mapped instrument questions and responses to Cardiac-centered Frailty Ontology concepts and properties.

We also tested the utility of the ontology in two different automated NLP systems. One system was designed to classify clinical note snippets as indicative or frailty or not. The other was designed to predict patient mortality after MCVP.

Additional file

Additional file 1: Frailty ontology concept list. (XLSX 37 kb)

Abbreviations

ADLs: Activities of daily living; BMI: Body mass index; CABG: Coronary artery bypass graft; CCFOID: Cardiac-centered frailty ontology identification number; DNN: Deep neural network; EHR: Electronic Health Record; FAQ: Functional activity questionnaire; FIM: Functional independence measure; IADL: Instrumental activities of daily living; MCVP: Major cardiovascular procedure; MRT: Morse fall scale; NLP: Natural Language Processing; OGMS: Ontology of General Medical Science; SCTID: SNOMED-CT identification number; SF-36: MOS 36-item short form health survey; STS: Society of Thoracic Surgeons; SVM: Support vector machine; TAVR: Transcatheter valve replacement; UMLS: Unified Medical Language System; VA: Veterans affairs

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Availability of data and materials

The datasets used and/or analyzed during the current study are available from the corresponding author on reasonable request. The ontology owl file is available on Bioportal at http://bioportal.bioontology.org/ontologies/CCFO.

Authors' contributions

KDH oversaw the group generating terms from assessment instruments and constructed the final ontology; CW and BEB provided oversight and ideas on all aspects of the project, and edits to the manuscript; BEB and RUS provided cardiology expertise, ideas, and edits. AT and JHG provided ideas and edits; YS, YC, and QZT provided terms gathered from two previous studies, ideas and integration with NLP. YC provided initial instrument score to frailty score mappings. All authors read and approved the final manuscript.

Ethics approval and consent to participate

This work is approved under University of Utah IRB_00096877 (Use Frailty Status to Predict Postoperative Outcomes in Elderly Patient) consent to participate was verbal (use of forms was waived).

Consent for publication

Not applicable

Competing interests

The authors declare that they have no competing interests.

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