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## Using The Unified Medical Language System to Expand the Operative Stress Score – First Use Case

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## Abstract

**Background:** The Unified Medical Language System (UMLS) maps relationships between and within >100 biomedical vocabularies, including *Current Procedural Terminology* (CPT) codes, creating a powerful knowledge resource which can accelerate clinical research.

**Methods:** We used synonymy and concepts relating hierarchical structure of CPT codes within the UMLS, (1) guiding surgical experts in expanding the Operative Stress Score (OSS) from 565

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originally rated CPT codes to additional, 1,853 related procedures; (2) establishing validity of the association between the added OSS ratings and 30-day outcomes in VASQIP (2015–2018).

**Results:** The UMLS Metathesaurus and Semantic Network was converted into an interactive graph database (https://github.com/dbmi-pitt/UMLS-Graph) delineating ontology relatedness. From this UMLS-graph, the CPT hierarchy was queried obtaining all paths from each code to the hierarchical apex. Of 1,853 added ratings, 43% and 76% were siblings and cousins of original OSS CPT codes. Of 857,577 VASQIP cases (mean age, 64±11years; 91% male; 75% white), 786,122 (92%) and 71,455 (8%) were rated in the original and added OSS. Compared to original, added OSS cases included more females (14% vs 9%) and frail patients (25% vs 19%) undergoing high stress procedures (11% vs 8%; all P<.001). Postoperative mortality consistently increased with OSS. Very low stress procedures had <0.5% (original, 0.4% [95%CI, 0.4–0.5%] vs added, 0.9% [95%CI, 0.6–1.2%]) and very high 3.8% (original, 3.5% [95%CI, 3.0–4.0%] vs added, 5.8% [95%CI, 4.6–7.3%]) mortality rates.

**Conclusions:** The synonymy and concepts relating biomedical data within the UMLS can be abstracted and efficiently used to expand the utility of existing clinical research tools.

### Keywords

Computational biology; Current procedural terminology; Frailty; Unified medical language system; medical informatics; medical information computing; ontologies; surgical procedures

The Unified Medical Language System (UMLS) is a powerful resource using a common structure and disciplined curation to achieve interoperability between diverse biomedical structured vocabularies (e.g., Systematized Nomenclature of Medicine Clinical Terms [SNOMEDCT], *Current Procedural Terminology* [CPT], *International Classification of Diseases* [ICD], etc.) and unstructured data (e.g., free text notes of patient encounters) generated through the documentation of routine patient care coded into subject areas with a defined set of properties and relationships with other categories known as ontologies. Begun by the National Library of Medicine in 1986, the UMLS has grown to include the concepts, codes, and terms among >100 vocabularies<sup>1</sup> with associated software that exploits the relationships among digital information to find conceptual connections within and across the otherwise completely disparate vocabularies.<sup>2</sup> UMLS connects vocabularies through concept synonymy, vitally important for efficient patient care and biomedical research.

The breadth, structure, and complexity of data within the UMLS has enormous potential to aid clinical research.<sup>3</sup> However, these strengths also create barriers to use. The intricate networks of data are difficult to conceptualize and requires advanced data management skills to manipulate,<sup>4</sup> confining use primarily to biomedical informatics experts and is still typically applied within only specific ontologies rather than for its structure and broad intraand inter-vocabulary conceptual consistency.

Just as a keen knowledge of anatomy is required for surgeons to perform an operation. We have defined key terms in Table 1 and designed a unique graph schema (i.e., network) connecting relationships between concepts as well as terms within and between vocabularies in the UMLS to facilitate its use by clinical and translational researchers.<sup>5</sup> As such, the clinical investigator can overcome previously frustrating limitations in administrative

datasets for tools specific to one vocabulary (e.g., CPT) but not another (e.g., ICD). Further, our tool facilitates the capacity of the UMLS to function as a crosswalk between synonymous variables included in disparate administrated datasets. This crosswalk's capacity also applies to human curated healthcare concepts (e.g., appendicitis treatments; appendectomy and ileocecectomy) to terms and codes that would otherwise be unrelated in structured data both within languages (CPT to CPT code) and between languages (CPT to ICD codes). Through this use case, we describe the structure, organization, and power of the UMLS and its potential in clinical and translational investigation.

## **Methods and Materials**

## **Clinical Context and Research Problem**

Despite the ubiquity of CPT codes, most health services research examines patient populations defined by families of similar procedures, treated by groups of specialists, focusing on a subset of physiological diagnoses. Lack of a common framework to group procedures across specialties and clinical diagnoses according to the rate of key outcomes has limited analysis of the entire range of surgical procedures. To address this limitation, our prior work applied a modified Delphi consensus methodology to develop the Operative Stress Score (OSS) classifying 565 CPT codes according to the degree of physiologic stress imposed by each procedure, ranging from very low to very high (OSS1–5).<sup>6,7</sup> Although these original 565 CPT codes represented 90% of the procedures performed in Veterans Affairs (VA) hospitals (2010–2014), the original OSS included only 16% of all 3,337 CPT codes present in VA sample. We hypothesized that the relationships inherent within CPT as manifest in the UMLS permits semi-automated expansion of the original OSS ratings to other conceptually related CPT codes (e.g., anatomically and physiologically similar procedures).

### **Data Sources**

We used two data sources to systematically expand the original OSS. First, all 2018 English UMLS sources from the standard UMLS distribution files, freely accessible by any individual requesting a UMLS Terminology Services Account (https://uts.nlm.nih.gov// license.html),<sup>1</sup> including the 2018 American Medical Association curated Category I CPT <sup>8</sup>. Second, the validated original OSS CPT code rating.<sup>6,7</sup>

We validated the added OSS among procedures included in the Veterans Affairs Surgical Quality Improvement Program (VASQIP; 2015–2018) containing CPT code-defined operative procedures and postoperative outcomes.<sup>9,10</sup> Our use of VASQIP data was reviewed by the VA Pittsburgh Healthcare System institutional review board and determined to be exempt (PRO3670).

#### UMLS Semantic graph

The UMLS organizes concepts, codes, and terms from source vocabularies into a unified human-curated network of conceptual relationships. Individual sources may be nonhierarchical or hierarchical but ultimately result in a single unified UMLS graph of the combined Metathesaurus and Semantic Network. This network is rendered in a vast series of

tables with rows of data consisting of, 1) vocabulary source **code** which is the smallest unit within the UMLS, 2) vocabulary source **term** which is a text string that corresponds to the code, 3) **UMLSConcept** defined by Concept Unique Identifiers (CUI), 4) UMLS Concept **preferred term**, and 5) UMLS **semantic type** (Table 1).<sup>5</sup>

The UMLS has limitless, multi-directional connectivity. We took the tables of data and depicted a simplified, hierarchical conceptual model of underlying UMLS relationships between codes, terms, concepts, and semantic types for the diagnosis and treatment of appendicitis (Figure 1). Each source code can be conceptualized, per the UMLS-Graph network, as a terminal branch with terms representing leaves on a tree. As with most codes, the CPT 44960 code has multiple synonymous terms (i.e., leaves), including "Appendectomy", "APPENDEC RPTD APPENDIX ABSC/PRITONITIS", and "Removal of ruptured infected appendix". For simplicity, the UMLS also defines, a single preferred term for each concept. For example, there are related codes both within (CPT 44950) and across vocabularies (SNOMEDCT 80146002, ICD9 47.0), each connected by a common UMLS concept (CUI C0003611 termed "Appendectomy"). Other related concepts (e.g., CUI C0003611 termed "Laparoscopic Appendectomy"), further aggregate into broader concepts. Each concept then consolidates into at least one semantic type. Here, the "Appendectomy" terms, codes, and concepts all emanate from the "Therapeutic or Prevention Procedure" left main branch of the tree. The treatment of appendicitis however requires a diagnosis, often achieved with imaging studies. Therefore, on the right side of the tree the "Diagnostic Procedures" expand into the associated "CT of abdomen and pelvis" leaves.

To expose the relationships between terms within and across vocabularies and facilitate research based on the UMLS, senior author (JS) conceptualized the complexity of this network into a property graph. The UMLS-Graph backend is manifest as a single application programming interface built upon the Neo4j Graph Database Platform (Neo4j, Inc.) that passes Cypher queries. All core relationships (e.g., semantic type-concept, concept-code, code-term, etc.) are included creating a directed property graph allowing for unified query and extraction of all data elements and the pathes between them.

In summary, the UMLS-Graph connects data elements commonly used in clinical research (e.g., CPT 44950, SNOMEDCT 80146002, and ICD9 47.0) which themselves are in separate vocabularies (CPT, SNOMEDCT, ICD9). In the eyes of the computer, each code is completely unrelated (44950, 80146002, 47.0). The UMLS allows these data elements to have a common definition, linking the clinical researcher's concept (e.g., "Appendectomy") with all related data elements for the computer (i.e., CUI C0003611).

## **CPT code Hierarchy**

The CPT code 5-digit numbers ascend from lowest to highest numerically and are organized within an indexed codebook with similar procedures numerically clustered and unrelated procedures numerically distanced. As a result, the CPT codes for an appendectomy include 44950, 44960, and 44970 and an above knee amputation include 27590, 27591, 27592, 27594, and 27596. However, precise relationships between CPT codes are not apparent in the CPT codebook. For example, surgeons understand that a laparoscopic appendectomy

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and open appendectomy are two separate, but related procedures. Within the CPT coding system, these two procedures are defined by codes with similar numbers (i.e., 44950 and 44970). However, transrectal drainage of a pelvic abscess is an unrelated procedure with a numerically similar code (i.e., 45000). As such, the UMLS-Graph organizes digitally functional data elements according to human curated concepts, informed by experience in both clinical care and health services research. Thereby the UMLS-Graph allows extraction of the underlying hierarchy of CPT codes that systematically and explicitly clusters related procedures according to their UMLS Concepts and Semantic Types.

We identified all Category I CPT codes as terminal branches with corresponding terms as leaves in the UMLS-Graph. We then queried UMLS-Graph to define the shortest path (i.e., shortest hierarchical sequence of UMLS Concepts and Semantic Types) from the terminal branch to the single main trunk representing the CPT hierarchical term: "Surgery". CPT codes (children) sharing their most distal branch (parent) were defined as siblings. CPT codes sharing a common branch one proximal to their most distal branch (grandparent) were defined as cousins.

### **OSS Expansion**

The CPT code hierarchy inherent to the UMLS-Graph was exported as a comma separated file for conceptualization and application of the interconnections in a single table of data in wide format. Each row contained a unique 5-digit CPT code (terminal branch) and term (leaves) in its last column with preceding columns including all increasingly broad UMLS Concepts (proximal branches of the CPT code explicit hierarchy). These data were then imported to Excel (Microsoft, Redmond, WA) for creation of the OSS Expansion Tool. In Excel, we used conditional formatting to create visual cues indicating sequential branching and therefore CPT code relatedness within the hierarchy – leaves on a common branch, consolidating branches, and main branches consolidating to a common trunk (eTable 1). We then identified and labeled each of the 565 original OSS rated CPT codes, preserving their location in the hierarchy.

All Category I CPT codes were eligible for expansion. We excluded CPT codes without minimal relatedness to an originally rated OSS including, 1) codes without any originally OSS rated CPT codes consolidating into the main branch, 2) CPT codes without any siblings in the secondary terminal branch.

Three of the separate surgical experts (DH, PS, RS) who led the original Delphi-generated OSS rating, independently reviewed the CPT codes eligible for expansion with the OSS Expansion Tool. Each reviewer added an OSS rating to siblings CPT codes sharing a common terminal branch and/or up the tree hierarchy to first or second cousins. Reviewers' ratings were then assessed for agreement and reconciled in a modified Delphi consensus method, mirroring the original methodology.<sup>6</sup> Unanimously rated codes were assigned (n=1,853). Disagreements within 1 OSS unit and only among one rater (n=461) were reviewed by the minority rater for concession with rate assignment (n=349) or continued disagreement (n=112). If the minority rater did not concede, those ratings in combination with disparate ratings (i.e., all three raters disagreed or >1 units apart, n=29) were discussed via teleconference until unanimity was reached.

To establish OSS ratings in regions of the CPT hierarchy excluded from the original consensus rating, 10 additional CPT codes were rated in accordance with anatomic and physiologic principals using the methods described above.

### **OSS Expansion – UMLS-Graph Use Case Validation**

We included all cases in VASQIP and excluded those with principal procedure CPT code in Category III (emerging technology). We categorized cases by those with an i) original OSS or ii) added OSS rating which together comprise the expanded OSS and excluded cases with no rating. To understand patient characteristics, we compared baseline demographics, preoperative risk factors, and intraoperative data. Postoperative 30-day mortality and complications were stratified by the Risk Analysis Index (RAI), a validated measure of patient frailty independently predicting postoperative morbidity and mortality independent of and synergistically with OSS.<sup>6,11–13</sup>

## Results

The UMLS-Graph, Graphing UMLS Enables Search in Dynamic Trees (https:// guesdt.com/). The semantic property graph includes 5,051,942 codes, 8,752,504 terms, 4,376,259 concepts, and 127 semantic types with a total of 49,172,202 relationships across 122 biomedical vocabularies. An example query within the UMLS-Graph includes searching the code "CPT 44970" which returns five terms: "LAPAROSCOPY APPENDECTOMY", "Laparoscopy, surgical appendectomy", "removal of appendix using an endoscope", "Surgical laparoscopy with appendectomy", and the preferred term "LAPAROSCOPIC APPENDECTOMY". When inputting the "CPT 44970" code into the UMLS-Graph, the graph traverses synonymous concepts for codes across ontologies (SNOMED, ICD9, etc.) and within (CPT) ontologies, for example branching to CPT 44970's parent and concept (CUI C0372525). A subset of the UMLS-Graph schematic is in eFigure 1.

### **CPT code Hierarchy and OSS expansion**

All 5,752 terminal leaf CPT codes in the UMLS-Graph are of the "Treatment and Prevention Procedure" Semantic Type (Figure 2A). The main trunk bifurcated into 19 main branches defining broad concepts (i.e., "Surgical Procedures of the Digestive System") with branches splitting up to five additional times before reaching the terminal leaves. The 19 branches split into 102 secondary (CPT code, n=5,741), 534 tertiary (CPT code, n=5,650), 1,136 quaternary (CPT code, n=3,692), 169 quinary (CPT code, n=670), and 37 senary (CPT code, n=76) additional conceptually more specific branches.

Six of the 19 main branches, containing 356 CPT codes, had no originally rated OSS CPT codes. Four secondary branches included only one CPT code. Therefore, of the 5,187 CPT codes without an original OSS, 4,827 CPT codes (93%) met criteria for potential primary expansion which included 13 main branches, 86 secondary, 486 tertiary, 1,050 quaternary, 119 quinary, and 35 senary subsequent branches (Table 2). Within and beyond the second branch of the tree, 1,049 CPT codes and 2,619 CPT codes had at least one original OSS rated CPT code sibling or cousin.

The conceptualized tree and associated relationships between CPT codes generated nine columns in the OSS Expansion Tool: (1) the main branch, (2–6) potential distal branches, (7) the CPT code and (8) Term leaf, (9) original OSS ratings when applicable. As conceptually depicted in Figure 2b, surgical experts reviewed each terminal leaf procedure term and the original OSS of related CPT codes. As appropriate, the OSS ratings were expanded to siblings (e.g., procedures of the appendix) or to siblings and cousins (e.g., procedures of the pancreas). CPT code relatedness guided the expansion, yet related codes were not automatically or blindly applied. For example, consider sibling CPT codes describing the

primary laparoscopic repair of a reducible (CPT 49654) and incarcerated or strangulated (CPT 49655) incisional hernia. In the original OSS rating, reducible repairs were low (OSS2) and incarcerated repairs moderate (OSS3) stress procedures. Similarly, the rating for laparoscopic repair of a recurrent incarcerated or strangulated incisional hernia (CPT 49657) was given an expansion rating of moderate (OSS3) above that of the originally rated, low stress (OSS2) sibling procedure describing a recurrent reducible repair (CPT 49656).

Guided by the OSS Expansion Tool, surgeon expertise, and formal rating review, the final OSS expansion was applied to 1,768 additional CPT codes, of which 775 had siblings and 1,349 had cousins with an original OSS rated CPT code. Ten additional CPT codes were selected for rating by modified Delphi process by surgical experts without using the OSS Expansion Tool describing unlisted procedures of the stomach (CPT code, 43999) and pancreas (CPT code, 48999) as well as ectopic pregnancy (CPT code, 59120, 59121, 59130, 59135, 59136, 59140, 59150, and 59151). Of the added 1,778 codes, 23% (OSS1, n=409) of procedures were very low, 36% (OSS2, n=641) low, 31% (OSS3, n=548) moderate, 8% (OSS4, n=146) higher, and 2% (OSS5, n=34) very high operative stress.

#### Validation of the Expanded OSS

VASQIP comprised a total of 886,823 cases, of which 786,122 (89%) cases were captured by the original OSS and 71,455 (8%) cases by the added (eFigure 2) which includes 562 (99%) of the original and 848 (48%) of the added CPT codes with OSS scores. The expanded OSS therefore captures 857,577 (97%) cases (mean age, 63 years [standard deviation (SD), 11]; 91% male; 25% non-White; mean RAI, 24 [SD, 7]). Compared to original OSS cases, added OSS cases included more females (14% vs 9%), and frail patients (RAI 30, 25% vs 19%) undergoing more stressful procedures (OSS4 and 5, 11.4% vs 8.2%; all P<.001). Accordingly, added OSS cases include a greater proportion of vascular, neurologic, otolaryngologic, gynecologic, plastic, and thoracic with a subsequent decrease in the proportion of the general and orthopedic surgery cases (Table 3).

Overall, postoperative mortality increased with higher levels of operative stress. Very low stress procedures (OSS1) had <0.5% (original, 0.4% [95%CI, 0.4–0.5%] vs added, 0.9% [95% CI, 0.6–1.2%]) and very high (OSS5) 3.8% (original, 3.5% [95%CI, 3.0–4.0%] vs added, 5.8% [95%CI, 4.6–7.3%]) mortality rate (**eTable 3**). Further, when stratifying by the OSS and patient frailty, the risk of postoperative mortality and complications in the added ratings mirrored the original ratings (Figure 3).

## Discussion

We utilized the network inherent in UMLS's data to create an application program interface to graphically relate over 100 biomedical vocabularies. In this first use case, we used the UMLS-Graph to formalize the relatedness of CPT codes into a single, user-friendly OSS Expansion Tool. We then expanded the OSS rating from the original 565 CPT codes to an additional 1,778 additional CPT codes, together the 2,343 expanded ratings covered 97% of cases included in VASQIP. Cases included by the original, added, and full expanded OSS rating continued to demonstrate that increasing operative stress correlated with increasing postoperative morbidity and mortality.

The relationships defined by the UMLS-Graph were exploited to generate a hierarchical network of CPT codes. Placing the OSS codes assigned by more labor-intensive modified Delphi process into the framework allowed for a systematic and efficient expansion of the OSS requiring smaller numbers of experts.<sup>14</sup> In review of the hierarchy and OSS, a purely automated expansion would be fraught with error. Therefore, the applicability of sibling, parent, and cousin relationships required interpretation by surgical experts. Supported by the conditionally formatted, visual cues within the OSS Expansion Tool the expansion process was accelerated while utilizing the knowledge gained from the original OSS modified Delphi consensus method.

The CPT hierarchy and associated OSS Expansion Tool might assist other health services researchers to improve both simple and complex inefficiencies by facilitating rapid identification, review and evaluation of CPT codes grouped according to clinical diagnoses (i.e., Appendicitis) and specific anatomical systems (i.e., Digestive). Further, the known regional differences in coding preferences, a frequently sited limitation of national administrative data, can be systematically evaluated controlling for defined relatedness within the UMLS-Graph and associated CPT Hierarchy.

The original 565 CPT codes captured approximately 90% of procedures completed in the VA healthcare system (2010–2014). In the more recent (2015–2108), VASQIP dataset, the original OSS captured 97% of surgical cases. The OSS expansion increased the number of rated CPT codes by >400% allowing the physiologic stress of a surgical intervention to be rated for >99% of VASQIP. Although we may have functionally reached an inclusion asymptote in VASQIP, 40% of OSS rated CPT codes were not included as principal procedures in this primarily male cohort. In a private sector cohort including more females, the OSS expansion has been recently validated in the American College of Surgeons National Surgical Quality Improvement Program demonstrating mortality outcome differences between men and women controlling for multiple factors including physiologically-induced operative stress using the UMLS-based OSS expansion.<sup>15</sup>

Medicare claims are determined by multiple code sets including both CPT, ICD, and Healthcare Common Procedure Coding System to quantify reimbursement.<sup>16</sup> Therefore, all patient care activities generate a potentially vast combination of billing codes in addition to an ever-growing list of healthcare related structured and unstructured data included within the UMLS.<sup>17</sup> In this use case, we employed the UMLS-Graph to create the OSS Expansion

Tool table specific to relationships between CPT codes. Investigators commonly face seemingly simple, but at times, insurmountable data management issues. The UMLS-Graph quickly and accurately managed a cross walk between administrative codes across two vocabularies for this work. Other commonly used tools, such as the Elixhauser and Charlson Comorbidity Indices could be easily crosswalked to existing biomedical vocabularies or expanded to include evolving and emerging comorbid conditions (e.g., COVID-19).

The analysis presented here was limited to the CPT vocabulary, and it remains to be seen if the UMLS-Graph can advance surgical research using other datasets and vocabularies. However, the potential applications for the UMLS-Graph are limited only by the imagination of investigators and their ability to conceptualize the complex nature of the UMLS network. For example, a more translational illustration of the utility of the UMLS-Graph is its ability to expose relationships between anatomy, diagnostic syndromes and specific genetic mutations. Anatomic locations can be queried with the Foundation Model of Anatomy vocabulary and linked to associated phenotypic and genetic abnormalities in the Online Mendelian Inheritance in Man vocabulary. For example, searching for genetic and phenotypic abnormalities pertinent to the liver generated 28 phenotypic and 27 genotypic liver related somatic mutations for systematic exploration. The results of this query are available in eTable 5 and include numerous glycogen storage, tumor and metabolic, and familial hypercholesteremia related diseases and genes. Investigators interested in exploring the UMLS-Graph may can access it freely on Github (https://github.com/dbmi-pitt/UMLS-Graph), are encouraged to use it expose the ever expanding and updated relationships within the UMLS.<sup>18</sup>

### Conclusion

In this first use case, we created a UMLS-Graph to systematically and efficiently expand the OSS by >400% and validated the accuracy of our added ratings. We have made available an application program interface for conceptualization of the UMLS-Graph to allow other investigators to harness the power within the UMLS for clinical research and data science applications.

## **Supplementary Material**

Refer to Web version on PubMed Central for supplementary material.

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## Highlights:

• Inherent hierarchical relationships in the UMLS are underutilized

- We graphically assembled into the UMLS-Graph
- The UMLS-Graph allowed for expansion of the Operative Stress Score by >400%
- The powerful UMLS-Graph can be used to similarly accelerate other clinical research



#### Figure 1. UMLS-Graph Schematic

The graphic depicts the hierarchical relationship within the UMLS as a tree, conceptually relating a subset of vocabularies (e.g., ICD9CM, SNOMEDCT, CPT) and the terms within and between vocabularies (i.e., "Appendectomy; for ruptured appendix..." and "Appendectomy") which are conceptually linked (i.e., "Appendectomy") in healthcare and via a CUI (i.e., C0003611) within the UMLS. In this stylized view of the UMLS-Graph graph, the base of the tree (e.g., trunks) include broad semantic types (brown circles) which branch into more specific Unified Medical Language System concepts (brown ovals). Although not depicted here, all concepts relate to each other, but the relationship may be variously close or distant. Branching from within each concept are codes specific to each vocabulary (green ovals) which relate first to the concept and then back to the semantic type. The codes are then described by a variety of synonymous terms (green pentagons) represented here as the most terminal aspect of the tree (e.g., leaves) with some leaves left empty for simplicity. Notably, the UMLS and associated UMLS-Graph contain graphical schema that are both hierarchical (as depicted here) as well as non-hierarchical. For simplicity, the diagram omits both non-hierarchical data and relationships that exist but cannot be easily represented in two dimensions.

Abbreviations: CPT, Current Procedural Terminology; CUI, Concept Unique Identifiers; SNOMEDCT, Systematized Nomenclature of Medicine Clinical Terms; ICD9CM,

*International Classification of Diseases*, 9<sup>th</sup> edition, Clinical Management; UMLS, Unified Medical Language System.

### A Sample Hierarchy Development - Fewest Nodes from Leaf to Trunk



# Figure 2. Sample Development (Panel A) and Sample Use (Panel B) of the *Current Procedural Terminology* Code Hierarchy

Panel A illustrates the development of the *Current Procedural Terminology* code hierarchy. The proximity of the leaf and terminal branch and main trunk were determined by the shortest distance, fewest number of common branches, along the tree. Panel B is an example of the *Current Procedural Terminology* code hierarchy structure and expansion of the OSS to siblings of the same color (e.g., red to red ovals) or cousins of different colors (e.g., red to orange ovals).

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# Figure 3. Postoperative mortality (Panel A) and complications (Panel B), stratified by patient frailty

Categories of Operative Stress Score rated *Current Procedural Terminology* codes are defined as, 1, very low stress; 2, low stress; 3, moderate stress; 4, high stress; 5, very high stress. Frailty is categorized in accordance with the validated Risk Analysis Index (RAI) as, RAI 20, robust (dark blue); RAI 21–29, normal (light blue); RAI 30–39, frail (orange); RAI 40, very frail (brown). 30-day postoperative mortality (A) and complications (B) for all cases included in the Expanded Operative Stress Score by frailty categorization (left) and for the individual frailty categorization (right) comparing cases rated by the Original (closed circle, solid line) and Added (open square, dashed line) Operative Stress Score. All error bars correspond with 95% Confidence Intervals.

### Table 1.

## Pertinent biomedical informatics and UMLS specific definitions

Term	Definition	Examples	
Ontologies	Formalized subject area with a defined set of properties and relationships with other categories	SNOMEDCT	
Vocabulary	Vocabulary Information organized by indexing or categorizing defined terms		
Vocabulary code	Smallest unit within a vocabulary within the UMLS	CPT Code 5-digit number: 48150	
Vocabulary term Text string that corresponds to the vocabulary code		Whipple; with pancreatojejunostomy	
UMLS concept Concepts describing and capturing codes/terms which may be specific to single or spanning across many vocabularies		Resection of pancreas	
UMLS Concept Unique Identifier (CUI) Alphanumeric code for a UMLS concept		C0372232	
UMLS preferred term	Text string corresponding to the UMLS concept, preferentially used by the UMLS	Resection of pancreas	
Semantic types	Description of concepts with varying levels of granularity and specificity, organized in a hierarchy	Therapeutic or Prevention Procedure	
Graph schema Network connecting codes, terms, concepts, and semantic types both with and without a hierarchical structure and relatedness		UMLS-Graph	
Property graph Graphing model where the connections between codes, terms, concepts, and sematic types themselves have a meaning (i.e., a name) and properties.		"branch_of", "component_of	

Abbreviations: SNOMEDCT, Systematized Nomenclature of Medicine Clinical Terms; CPT, Current Procedural Terminology, MeSH, medical subject headings, UMLS, Unified Medical Language System.

#### Table 2.

#### Current Procedural Terminology Codes Eligible for Primary Operative Stress Score Expansion

		Branches (no. branches)						
		Main <sup>*</sup> (13)	Second <sup>*</sup> (86)	Third (486)	Fourth (1,050)	Fifth (119)	Sixth (35)	
CI	CPT code, no. (%)							
	Terminal branch	0	43 (1%)	1,659 (34%)	2,642 (55%)	418 (9%)	65 (1%)	
	OSS-Rated siblings?	0	10 (1%)	622 (59%)	329 (31%)	81 (8%)	7 (1%)	
	OSS-Rated cousins?	0	10 (1%)	1,416 (54%)	1,028 (39%)	126 (5%)	39 (1%)	
M	Main branch system description $^{/\!/}$ , no. CPT code per branch (%) $^{\#}$							
	Auditory	92 (1.9%)	92 (1.9%)	88 (1.8%)	39 (1.2%)	0	0	
	Cardiovascular	660 (13.7%)	660 (13.7%)	660 (13.8%)	500 (16.0%)	173 (35.8%)	17 (26%)	
	Digestive	757 (15.7%)	757 (15.7%)	757 (15.8%)	464 (14.8%)	80 (16.6%)	1 (2%)	
	Endocrine	19 (0.4%)	19 (0.4%)	18 (0.4%)	12 (0.4%)	0	0	
	Female Genital	207 (4.3%)	207 (4.3%)	202 (4.2%)	89 (2.8%)	9 (1.9%)		
	Hemic and Lymphatic	57 (1.2%)	57 (1.2%)	56 (1.2%)	32 (1.0%)	0	0	
	Integumentary	367 (7.6%)	367 (7.6%)	348 (7.3%)	279 (8.9%)	37 (7.7%)	31 (48%)	
	Male Genital	126 (2.6%)	126 (2.6%)	124 (2.6%)	45 (1.4%)	0	0	
	Mediastinum and Diaphragm	14 (0.3%)	14 (0.3%)	14 (0.3%)	5 (0.2%)	0	0	
	Musculoskeletal	1,460 (30.2%)	1,460 (30.2%)	1,450 (30.3%)	903 (28.9%)	42 (8.7%)	0	
	Nervous	494 (10.2%)	494 (10.2%)	494 (10.3%)	403 (12.9%)	108 (22.4%)	16 (25%)	
	Respiratory	285 (5.9%)	285 (5.9%)	285 (6.0%)	196 (6.3%)	0	0	
	Urinary	289 (6.0%)	289 (6.0%)	288 (6.0%)	158 (5.1%)	34 (7.0%)	0	

All CPT codes consolidating to a main branches which did not contain an originally rated CPT code were excluded from the primary expansion (n=356). All CPT codes terminating in the second branch without any siblings, independent of original OSS rating, were excluded (n=4).

<sup>?</sup>CPT code relatedness is defined only in the most distal, and therefore contextually specific, available branch.

<sup>//</sup>Main branches with CPT codes excluded from primary expansion include: Fine needle aspiration (CPT code, n=2), Intersex surgery

(CPT code, n=2), Operating microscope procedures (CPT code, n=1), Reproductive system procedures (CPT code, n=1), Maternity care (CPT code, n=64), Eye and ocular (CPT code, n=286).

<sup>#</sup>Includes any CPT code consolidating onto each branch, not just solely the CPT codes at their most terminal branches. For example, there are 92 CPT codes which consolidate onto the main Auditory branch. Of which, all traverse a second branch, 88 traverse and 49 terminate on a third branch, the remaining 39 terminate at a fourth branch.

Abbreviations: OSS, Operative Stress Score; CPT, Current Procedural Terminology.

### Table 3.

## Patient and case characteristics

Variable	Expanded OSS	Original OSS	Added OSS
	n=857,577	n=745,142	n=14,819
Age, years, mean (SD)	64 (11)	64 (11)	62 (12)
Male sex, no. (%)	777,912 (91%)	716,507 (91%)	61,405 (86%)
Race, no. (%)			
White	612,332 (75%)	563,330 (76%)	49,002 (73%)
Black	152,797 (19%)	138,573 (19%)	14,224 (21%)
Asian or Pacific Islander	3,420 (<1%)	3,015 (<1%)	405 (1%)
Other	43,272 (5%)	39,743 (5%)	3,529 (5%)
Ethnicity, no. (%)			
Not Hispanic	784,371 (93.2%)	720,100 (93.3%)	64,271 (92.1%)
Hispanic	45,307 (5.4%)	40,976 (5.3%)	4,331 (6.2%)
Risk Analysis Index, no. (%)		-	
Robust (20)	213,859 (27.3%)	194,235 (26.9%)	19,624 (30.6%)
Normal (21–29)	416,963 (53.1%)	388,675 (53.9%)	28,288 (44.2%)
Frail (30–39)	123,416 (15.7%)	110,942 (15.4%)	12,474 (19.5%)
Very frail ( 40)	30,556 (3.9%)	26,876 (3.7%)	3,680 (5.7%)
American Society of Anesthe	thesiologists Classification, no. (%)		
1	4,595 (0.5%)	4,020 (0.5%)	575 (0.8%)
2	134,518 (15.7%)	124,584 (15.9%)	9,934 (14.0%)
3	599,296 (69.9%)	552,123 (70.3%)	47,173 (66.3%)
4	116,546 (13.6%)	103,296 (13.1%)	13,250 (18.6%)
5	2,090 (0.2%)	1,862 (0.2%)	228 (0.3%)
Operative Stress Score, no. (9	%)	-	
1	44,778 (5.2%)	39,842 (5.1%)	4,936 (6.9%)
2	377,023 (44.0%)	343,954 (43.8%)	33,069 (46.3%)
3	363,320 (42.4%)	337,986 (43.0%)	25,334 (35.5%)
4	64,562 (7.5%)	57,644 (7.3%)	6,918 (9.7%)
5	7,894 (0.9%)	6,696 (0.9%)	1,198 (1.7%)
Procedure type, no. (%)			
General	225,105 (26.5%)	213,605 (27.4%)	11,500 (16.3%)
Orthopedic	249,714 (29.4%)	237,460 (30.4%)	12,254 (17.4%)
Vascular	127,549 (15.0%)	109,198 (14.0%)	18,351 (26.0%)
Gynecologic	12,528 (1.5%)	8,795 (1.1%)	3,733 (5.3%)
Urologic	99,899 (11.7%)	94,525 (12.1%)	5,374 (7.6%)
Neurologic	66,068 (7.8%)	58,809 (7.5%)	7,259 (10.3%)
Otolaryngologic	23,967 (2.8%)	20,447 (2.6%)	3,520 (5.0%)
Plastics	15,948 (1.9%)	11,357 (1.5%)	4,591 (6.5%)

	Variable	Expanded OSS	Original OSS	Added OSS	
		n=857,577	n=745,142	n=14,819	
	Thoracic	29,900 (3.5%)	25,865 (3.3%)	4,035 (5.7%)	
30	30-day postoperative outcome, no. (%)				
	Any complication	65,313 (7.6%)	57,913 (7.4%)	7,400 (10.4%)	
	Mortality	9,658 (1.1%)	8,432 (1.1%)	1,226 (1.7%)	

Abbreviations: no.: Number; SD: standard deviation; OSS, Operative Stress Score.