SMART precision cancer medicine: a FHIR-based app to provide genomic information at the point of care

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ABSTRACT

Background Precision cancer medicine (PCM) will require ready access to genomic data within the clinical workflow and tools to assist clinical interpretation and enable decisions. Since most electronic health record (EHR) systems do not yet provide such functionality, we developed an EHR-agnostic, clinico-genomic mobile app to demonstrate several features that will be needed for point-of-care conversations.

Methods Our prototype, called Substitutable Medical Applications and Reusable Technology (SMART)© PCM, visualizes genomic information in real time, comparing a patient’s diagnosis-specific somatic gene mutations detected by PCR-based hotspot testing to a population-level set of comparable data. The initial prototype works for patient specimens with 0 or 1 detected mutation. Genomics extensions were created for the Health Level Seven© Fast Healthcare Interoperability Resources (FHIR©) standard; otherwise, the prototype is a normal SMART on FHIR app.

Results The PCM prototype can rapidly present a visualization that compares a patient’s somatic genomic alterations against a distribution built from more than 3000 patients, along with context-specific links to external knowledge bases. Initial evaluation by oncologists provided important feedback about the prototype’s strengths and weaknesses. We added several requested enhancements and successfully demonstrated the app at the inaugural American Society of Clinical Oncology Interoperability Demonstration; we have also begun to expand visualization capabilities to include cancer specimens with multiple mutations.

Discussion PCM is open-source software for clinicians to present the individual patient within the population-level spectrum of cancer somatic mutations. The app can be implemented on any SMART on FHIR-enabled EHRs, and future versions of PCM should be able to evolve in parallel with external knowledge bases.

Keywords: health information management, electronic health records, genomics, neoplasms, information science, mobile health

BACKGROUND AND SIGNIFICANCE

The definition of cancer and the care of cancer patients are increasingly being driven by tumor genomics, aka molecular profiling.1, 2 As the number of clinically relevant findings with prognostic implications rapidly expands, human cognitive capacity, as predicted in 1989,3 will no longer be able to keep up. Vanderbilt University Medical Center (VUMC) was an early adopter of near-universal genotyping of cancer specimens for a number of disease-specific “actionable” mutations, through the use of SNaPshot multiplexed PCR mutation panels.4–7 Even with the limited number of genes tested in SNaPshot, it is already very difficult for an individual clinician to be intimately familiar with the population distributions of genomic alterations and their implications.8, 9 These difficulties will only grow worse as next-generation sequencing (NGS) will utilize cancer gene panels approaching 500 genes and potentially tens of thousands of variants per specimen.9–11 Unfortunately, commercially available electronic health records do not provide ready means to display clinical genomic data, nor additional functionalities such as links to external knowledge bases including My Cancer Genome, a curated oncology gene variant knowledge base, among others. This gap must be addressed for clinicians to achieve the goals set forth in President Barack Obama’s Precision Medicine Initiative.12

OBJECTIVE

We sought to develop an open-source application based on the Substitutable Medical Applications and Reusable Technology (SMART) Health IT platform (www.smarthealthit.org), an open-access application programming interface (API) that enables apps to run broadly across the health care ecosystem.13 We used Health Level Seven International (HL7©)’s Fast Healthcare Interoperability Resources (FHIR©) standard and its extension features for native representation of molecular profile data. The purpose of the resulting SMART Precision Cancer Medicine (PCM) app is to present population-level genomic health information to oncologists and their patients in real time as a component of clinical practice. We also wanted to demonstrate the ease of including seamless links to external knowledge bases within the app.

MATERIALS AND METHODS

The SMART API, SMART on FHIR, and genomics extensions

The SMART platform has been previously described.14, 15 HL7 is a standards development organization that has developed several widely used standards in the health care space. The newest HL7 standard is FHIR, currently a draft standard for trial use. Similarly to HL7 V3, FHIR is a constraint on the HL7 Reference Information Model,16 with some minor modifications. In addition, FHIR is based around the latest Web technologies, such as representational state transfer APIs, and can be represented in extensible markup language, JSON, and other formats. SMART on FHIR provides an app platform for health applications that integrates with EHRs, patient portals, personal health records,
and data warehouses. There are 3 key aspects of SMART on FHIR: (1) a data access layer based on FHIR, combined with a set of constraining profiles that lock down optionality and align vocabularies with Meaningful Use requirements,17,18 (2) a security layer that provides narrowly scoped authorization to specific portions of a patient’s record via OAuth 2.0,19 and (3) a single-sign-on layer using OpenID Connect.20 SMART on FHIR apps can be integrated into the context of an existing EHR or patient portal session, conveying the current patient, encounter, and other details of the host environment, or they can launch independently, such as on a mobile phone or device.

The SMART on FHIR Genomics API provides additional functionality to SMART on FHIR by extending the FHIR Observation resource to support clinical genomic data.21 These extensions are the basis of the FHIR Standard Profile for Genetics, which was published on September 23, 2015 (http://www.hl7.org/FHIR/observation-genetics.cg-prf-1a.html).

Translating local data into FHIR-compliant data
We were interested in displaying the following information for clinical consumption: (1) demographics including name, medical record number, gender, and age, (2) primary cancer diagnosis, and (3) molecular profile results. Comorbidities were not displayed at this pilot stage. Solid tumor oncology patients with certain histologies (eg, lung cancer, melanoma) seen at VUMC have, since July 2010, routinely undergone molecular profiling with SNaPshot, a fast, high-throughput, multiplex mutational profiling method based on the Applied Biosystems SNaPshot platform.5,7 SNaPshot assays for common somatic mutations across multiple cancer-associated genes, as defined in the Catalogue of Somatic Mutations in Cancer (COSMIC),22 SNaPshot panels have been performed on more than 4500 cancer specimens at VUMC (as of August 2015).

SNaPshot data are stored in the VUMC EHR and related data warehouses using an internally developed local code set. We transformed these data into unambiguous codable concepts after reviewing various terminology options for diseases, genes, gene alterations, and protein alterations (see Supplementary Table 1). This candidate list was generated by searching for codes for a representative disease (melanoma) seen at VUMC have, since July 2010, routinely undergone molecular profiling with SNaPshot, a fast, high-throughput, multiplex mutational profiling method based on the Applied Biosystems SNaPshot platform.5,7 SNaPshot assays for common somatic mutations across multiple cancer-associated genes, as defined in the Catalogue of Somatic Mutations in Cancer (COSMIC).22 SNaPshot panels have been performed on more than 4500 cancer specimens at VUMC (as of August 2015).

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### Table 1: Examples of local code mapping to standardized codable concepts.

<table>
<thead>
<tr>
<th>Local Code</th>
<th>Human Gene Naming Consortium ID</th>
<th>National Cancer Institute Thesaurus</th>
<th>Consensus Coding Sequence</th>
<th>Human Genome Variant Society</th>
<th>Display Name</th>
<th>Gene Wiki URL</th>
</tr>
</thead>
<tbody>
<tr>
<td>E790M</td>
<td>HGNC:3236</td>
<td>C17757</td>
<td>c.2369C&gt;T &lt;br&gt;p.T790M</td>
<td>EGFR T790M</td>
<td>~Epidermal_growth_factor_receptor</td>
<td></td>
</tr>
<tr>
<td>G12CM</td>
<td>HGNC:6407</td>
<td>C25785</td>
<td>c.34G&gt;T &lt;br&gt;p.G12C</td>
<td>KRAS G12C</td>
<td>~KRAS</td>
<td></td>
</tr>
<tr>
<td>G13CM</td>
<td>HGNC:6407</td>
<td>C25785</td>
<td>c.37G&gt;T &lt;br&gt;p.G13C</td>
<td>KRAS G13C</td>
<td>~KRAS</td>
<td></td>
</tr>
</tbody>
</table>

**SMART PCM app overview**
A team of software developers within the Vanderbilt-Ingram Cancer Center (VICC)’s Research Informatics Core developed a native iOS app optimized for the iPad and iPad Mini tablet devices (Apple Inc., Cupertino, CA, USA). Development proceeded in a continuous delivery build cycle with frequent input from clinical subject matter experts, especially with respect to how to properly model the clinical genomic information.
data; the actual implementation of FHIR was straightforward. An integral component of the development process was implementation and refinement of the app at the FHIR Connectathon 7 (September 13–14, 2014) and a Connectathon hosted by the American Society of Clinical Oncology (May 4–5, 2015); details are in Supplementary Materials.

The PCM app is accessed from an icon on the home screen of an iPad. The user authenticates by using their standard VUMC username and password. As required by the OAuth 2.0 process, the user must authorize the app for initial data access. A splash-screen is presented and followed by a simple query interface, which allows a practitioner to look up a patient by name or medical record number.

A successful patient query brings up the patient-centric view, as shown in Figure 2. In the example, the patient has lung cancer that harbors a KRAS p.G12C mutation. To prevent visual clutter in the initial series of pie charts, only common occurrences are labeled. The clinician can see the fully labeled information by interacting with the charts, as shown in Figure 3. The “dive-in” detail includes full labels and exact patient counts. From that view, the user can link to external Gene Wiki content by touching the desired gene (Figure 4).

User feedback survey
After completion of the initial development cycle, which included FHIR Connectathon 7, we performed user testing with oncology clinicians. The SMART PCM app was evaluated in the VICC medical oncology clinic, with the data source being 3800 patients who had undergone SNaPshot testing with 0 or 1 mutation detected. All queries for names, medical record numbers, and genomic data were performed in real time from a local SMART on FHIR server, and data was fed to the app via FHIR bundle and resource objects. This initial evaluation focused on collecting clinician feedback and was not carried out in the

Figure 1: A snippet of the JSON FHIR code for a patient with lung cancer and a p.T790M mutation detected in the epidermal growth factor receptor gene. Three extensions to the FHIR Observation Resource are shown: (1) assessed.gene, which uses the NCI Thesaurus to represent the gene name in HGNC-compliant format; (2) assessed.referenceSeq, which uses the CCDS database to represent the gene reference sequence; (3) assessed.variant, which represents the observed gene mutation (c.2369C>T) and predicted protein alteration (p.T790M) directly in Human Genome Variant Society syntax. The full code for this patient is available in Supplementary Table 2.

```json
{
  "url": "https://fhir/Mutation#assessed.gene",
  "valueCodeableConcept": {
    "coding": [
      {
        "system": "http://ncit.nci.nih.gov",
        "code": "C17757",
        "display": "EGFR"
      }
    ]
  },
  "url": "https://fhir/Mutation#assessed.referenceSeq",
  "valueCodeableConcept": {
    "coding": [
      {
        "code": "5514.1",
        "display": "CCDS 5514.1"
      }
    ]
  },
  "url": "https://fhir/Mutation#assessed.variant",
  "valueCodeableConcept": {
    "coding": [
      {
        "system": "http://www.hgvs.org/mutnomen",
        "code": "c.2369C>T",
        "display": "c.2369C>T"
      },
      {
        "system": "http://www.hgvs.org/mutnomen",
        "code": "p.T790M",
        "display": "T790M"
      }
    ]
  }
}
```
presence of patients. Similar to other “apps” in the VUMC ecosystem (eg, WebPACS), the app is launched independently from the EHR and can be used synchronously or asynchronously per user needs. Two groups of users were targeted for evaluation of the application: fellows within the oncology training program and practicing attending oncologists. Fourteen users (9 fellows and 5 attendings) were approached for user evaluation, with a goal to ascertain the majority of problems and experiences for each group. After they had freely explored the app’s functionality without a time limit, they were directed to complete a short online survey. Survey questions were created based on informal needs assessments and conversations between 2 clinical oncologists. Questions were created around user concepts and specific potential improvements. Additionally, users were able to enter open-ended free-text comments. All users completed the survey, which is described in the Supplementary Materials.

Statistical Analysis
Survey responses were exported from Research Electronic Data Capture into the R statistical package for analysis. Free-text comments were coded by 2 independent raters (J.L.W., M.J.R.) for content, with discrepancies adjudicated by joint consultation; interrater reliability was calculated using Cohen’s kappa (κ). Hypothesis testing was performed using Fisher’s exact test and Mann-Whitney U; all statistical tests were 2-sided.

Ethics and software availability
The described tool is a quality improvement initiative with the intent to implement a practice to improve the quality of patient care, and was thus determined by the Vanderbilt Institutional Review Board to be non-research per section 45 CFR 46.102(d) of the Health Insurance Portability and Accountability Act.


RESULTS
Implementing SMART on FHIR Genomics at VUMC
VUMC’s extensive research data warehouse is a relational database containing detailed information on over 3 million patients, dating back to 1992. This includes most data from the EHR, such as clinical
progress notes and provider-patient communications, as well as data feeds directly from laboratory and billing systems. Our app was built to interface with a local SMART on FHIR server, using data directly from the research data warehouse, analogous to previous work with i2b2. The local server was similar to the SMART on FHIR prototype described above, except that OAuth 2.0 was linked to a custom internal authentication and role-based authorization service, which utilizes lightweight directory access protocol for access to client applications. Access to the server was only enabled within the VUMC firewall and was audited.

User feedback

User evaluations were highly variable across all nominal domains tested, with no correlation among variables (eg, users who found the system quick and easy to use would not necessarily use it clinically; see Supplementary Figure 1). There was no statistical difference between fellows and attendings in their responses to any of the questions ($P > 0.05$).

Table 2 shows how clinicians responded to the survey’s list of 8 categories of potential functional additions. The top request was for assistance with selecting the right targeted drug for a given mutation, followed by a request for more links to external knowledge bases, population-level outcome information, and treatment cost information.

The survey results included 35 free-text comments about the app. These comments were coded to 5 categories (navigation, annotation, clinical utility, speed, general) with good interrater agreement ($\kappa = 0.84$). The most common categories were technical aspects of navigation and general comments (Table 3). Several representative responses to the question of “What features could be improved?” are as follows:

“I think this could be a useful clinical tool if additional features were built in. I would not show this to the average patient that I see, because I generally feel that too much information is confusing to the average pt.”

“Can include more information such as access to my cancer genome. Link to available targeted agents, cost, survival etc.”

“More clear deliniation [sic] of the patient’s findings. All to be legible. Speed may deteriorate when dataset increases. There needs to be a demo to show you how it works. All in all great though!”

Implementing SMART PCM at the ASCO interoperability demonstration

Based upon several of the themes that emerged from the survey results, we implemented additional functionalities. The variant pie visualization was activated so that a user interacting with this data would be brought to the disease- and variant-specific page of My Cancer Genome. Additional icons were added to bring the user to
The enhanced app was competitively reviewed for inclusion in the American Society of Clinical Oncology (ASCO)’s inaugural interoperability demonstration, and was accepted along with 11 other vendor products. SMART PCM received a FHIR-compliant message containing genomic information about a synthetic patient with BRAF-mutated colon cancer in real time from a third-party clinical genomics laboratory, GenoSpace LLC (Cambridge, MA, USA). For the demonstration, the synthetic patient was compared to a locally cached synthetic colon cancer patient population (N = 415); visualization construction was performed in real time.

Feasibility of expanding to NGS
Further modifications to the app realized the function of showing data from NGS test results, using synthetic patient data (N = 150,

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**Table 2: Features clinical users would like to see in future PCM designs.**

<table>
<thead>
<tr>
<th>Additional feature</th>
<th>Percent of fellows requesting feature (n = 9) (%)</th>
<th>Percent of attendings requesting feature (n = 5) (%)</th>
</tr>
</thead>
<tbody>
<tr>
<td>Decision support (eg, what drugs will work for my patient?)</td>
<td>67</td>
<td>60</td>
</tr>
<tr>
<td>More external knowledge content (eg, links to My Cancer Genome, COSMIC)</td>
<td>44</td>
<td>40</td>
</tr>
<tr>
<td>Outcome information (eg, survival)</td>
<td>44</td>
<td>40</td>
</tr>
<tr>
<td>Cost information (eg, how much will the drugs cost that could work based on observed mutation?)</td>
<td>33</td>
<td>40</td>
</tr>
<tr>
<td>Larger populations (eg, state- or country-level statistics)</td>
<td>0</td>
<td>20</td>
</tr>
<tr>
<td>Stratify by criteria such as age, gender, treatment exposure, stage</td>
<td>11</td>
<td>0</td>
</tr>
<tr>
<td>Additional visualizations</td>
<td>0</td>
<td>0</td>
</tr>
<tr>
<td>All of the above</td>
<td>11</td>
<td>20</td>
</tr>
</tbody>
</table>

**Table 3: User comments by category.**

<table>
<thead>
<tr>
<th>Coded user comment category</th>
<th>Comments count</th>
</tr>
</thead>
<tbody>
<tr>
<td>Navigation</td>
<td>8</td>
</tr>
<tr>
<td>Annotation</td>
<td>6</td>
</tr>
<tr>
<td>Clinical utility</td>
<td>2</td>
</tr>
<tr>
<td>Speed</td>
<td>1</td>
</tr>
<tr>
<td>General</td>
<td>8</td>
</tr>
</tbody>
</table>

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**Figure 4:** The SMART PCM app allows for access to external knowledge sources that would otherwise be unavailable to the user through their native EHR system. Shown here is the Gene Wiki page for the gene KRAS, embedded within the app.
see Supplemental Methods). A pie chart was used to stand out gene mutations detected in 1 patient from all gene mutations observed in similar patients. Further analysis of the distribution of gene mutations was also performed by local outlier factor (LOF) analysis. LOF is an efficient method to identify density-based outliers (see Supplemental Methods). An example is shown in Figure 5 and Figure 6, and Supplementary Figure 2.

DISCUSSION
Cancer is a set of complex diseases whose treatment calls for highly individualized diagnosis and rapidly evolving treatments; indeed, the knowledge base in the genetics of cancer is expanding exponentially. This complexity has been recognized by the broad research and patient care community, and cancer is the first area to be tackled through the Precision Medicine Initiative. As knowledge of the genomic drivers of cancer grows along with the increasing number of druggable targets, matching patients to treatments has become increasingly important. Many oncologists treat a wide variety of cancers, and the potential to move from the existing paradigm of more than 100 distinct types of cancer to thousands or more distinct subtypes is daunting. Given the pace of knowledge generation and the transition to large cancer gene panels and routine whole-exome or whole-genome sequencing, it will be a challenge to incorporate genomic cancer data into existing EHR platforms on a near–real time basis.

We decided that an appropriate initial scope for our clinico-genomic prototype would be to provide information to the clinician for situating a patient relative to other patients in the clinical genomics space. Providing contextually useful patient population comparisons may be a “nice to have” for traditional clinical tests. However, for the...
complex and highly differentiating results associated with genomic testing, such information is essential for any type of diagnostic/ prognostic support, since more active clinical decision rules may depend entirely upon the particular evidence base for the clinical condition of the patient. The informational support provided by the PCM prototype immediately informs the clinician about the patient in a cancer population context and it allows the physician to share—show and tell—this information with the patient.

The PCM app prototype demonstrates how the “app ecosystem” path can keep pace with the underlying medical science. Our prototype demonstrates how to achieve end-to-end integration with a data warehouse operating in near–real time with the accompanying EHR system. The SMART on FHIR components enable role-based authentication and authorization for obtaining patient context and population-level data within a firewalled and audited environment. Given the security concerns surrounding PHI, this is a critical aspect. The information required by SMART PCM demonstrates the convenience of the FHIR data model: to satisfy the clinical data requirement, we only had to stand up 4 FHIR Resources: Patient, DiagnosticReport, Specimen, and Observation. Although FHIR had no off-the-shelf way to package the gene and variant data requirements, existing FHIR Resources were readily extended to capture the required information.

By writing the prototype app as a SMART app, we have made it deployable on different EHR systems that have exposed the same FHIR Resources and Extensions. The SMART design also allows accessing patient-specific gene and variant data from a secondary source, such as a commercial DNA sequencing data service. Indeed, a key way in which SMART on FHIR Genomics could stimulate innovation is to offer the right combination of predictable data payloads and secure architectures to simplify “mashing up” of data originating in differing locations, so that capabilities such as those displayed in SMART PCM are no longer exotic. This technical solution does not obviate the need for appropriate data governance policies to ensure that the appropriate level of trust, privacy, and security persists throughout the health care ecosystem. With the growing impetus for post–Meaningful Use interoperability such as The Office of the National Coordinator for Health Information Technology (ONC)’s Federal Health IT Strategic Plan 2015–2020 and embracing of the app concept by some EHR vendors, the time for interoperable standards-based apps is nigh. Through the Argonaut Project (https://hl7-fhir.github.io/argonauts.html), an industry-academic consortium, the SMART on FHIR API is being incorporated into 5 major EHR vendor products. The Health Services Platform Consortium, a nonprofit organization with multiple vendor participants, is also promoting the use of apps with a FHIR service layer and EHR integration using SMART. The uptake of SMART by vendors leads to a “win-win” situation where innovation can freely occur and vendors are also free to reuse and adapt open-source app contents into their own products.

Feedback from testers revealed several themes. Foremost is the heterogeneous expectations of clinical oncology users for technology in delivering information. Ranges for the continuous variables in Supplementary Figure 1 averaged 82.75 points out of 100. Additionally, users who rated the app highly with regard to speed or ease of use were not more or less likely to say they would use it clinically. This illustrates a second theme among users: that speed and ease of use must be present, but it is content that will drive the ultimate utility of a clinical app. As demonstrated in Table 2, features such as enhanced decision support and more links to external knowledge bases were the most desired features, whereas additional
visualization and the ability to display data by demographics were less desired. Somewhat surprisingly, only 1 user (out of 14) requested inclusion of larger databases such as COSMIC into the population-level displays. This may reflect the large size of the institutional VUMC database and it is likely that implementers with a smaller local database might want external data included to some degree.

As we found in our assessment of coding terminologies (Supplementary Table 1), there are at least 26 structured vocabularies, syntaxes, or Web resources for diseases, 24 for genes, 9 for gene alterations, and 7 for protein alterations. There are also additional vocabularies that have terms explicitly linking 1 or more of these categories (eg, Orphanet links genes to proteins). Although this list may not be comprehensive, choosing from among these was still by no means straightforward. Issues such as version control, provenance, and compatibility must all be considered when selecting terminologies, and for this project we found that the consensus approach was best to reconcile these challenges. However, this process can be time-consuming and does not always guarantee optimal results.

As we demonstrated after the initial user feedback sessions, the addition of more external links is straightforward to implement. External linking capabilities could easily be extended in future work, including patient-centric resources such as Cancer.net and the National Center for Biototechnology Information library of cancer information, and shared decision-making resources such as ClinicalTrials.gov, a clinical trial aggregator. In the absence of an indisputable authoritative knowledge source, multiple links could be provided.

The function of drawing a pie chart showing multiple gene mutations observed in a single specimen substantiates the feasibility of expanding to NGS. However, Figure 5 also demonstrates some significant visualization challenges. First, for the large amount of gene mutations observed, it is hard to integrate all information in one figure. If we display all gene names and their occurrence level, the figure would become too crowded for clinicians to catch things that really matter, while much information would be omitted if we stand out only the most important information. Although alternative visualizations such as bubble plots might overcome some of the weakness of the pie/donut representation, they still suffer from clutter at this scale. Some have advocated circular plots that show connections between objects or between positions, which is an attractive way to display multidimensional cancer genomics data for scientific publication, but also tends to be over-cluttered for clinical use. Even more challenging, there is no consensus on how-identifying the most valuable information; sometimes high levels of occurrence do not tie in with high importance. For example, the most commonly mutated gene in cancer, tumor protein p53, does not have a therapeutic target. Second, inner connections (eg, physical/genetic interactions) may exist among gene mutations detected; thus, network analysis such as LOF holds the potential to decode potential relationships of mutations including variants of undetermined significance. Visualization of clinico-genomic data is a complex issue that will not be solved for some time. The National Cancer Institute has recognized this complexity and has recently issued a request for applications for Visualization Genomic Data Centers.

CONCLUSION

We built and tested a standards-based clinico-genomic app, with immediate applicability to many cancer patients. The ultimate goals of the SMART PCM app are to (1) provide practitioners with context-dependent population-level cancer mutation information, (2) act as a within-workflow intermediary to select external knowledge bases, and (3) enable a patient-centered and gene-driven shared decision-making model. Patients, caregivers, and clinicians do not wish “precision medicine” to be a mere buzzword, but rather want to know the context of the disease (eg, untreated BRAF-mutated metastatic colon cancer), the prognosis of the disease (eg, the median survival for a patient with untreated BRAF-mutated metastatic colon cancer), and what can be done about the disease (eg, published efficacious treatment regimens for untreated BRAF-mutated metastatic colon cancer). At the same time, personalization does not mean consignment to isolation. If anything, a well-defined niche diagnosis can bring about a sense of solidarity with others so afflicted, as evidenced by the numerous and strong rare disease coalitions, and sites such as PatientsLikeMe (Cambridge, MA, USA). This can especially be the case when membership in such a group brings the option of targeted therapy, which may also include a unique set of side effects. Apps such as SMART PCM can evolve quickly and nimbly, and offer a new and innovative resource for precision medicine.

Physicians have been described as professional information consumers, and with the increasing power of mobile computing they will increasingly turn to mobile applications for information. These applications will have to be built upon robust data standards to ensure accuracy, speed, and interoperability; however, it will ultimately be content that drives clinical utility and adoption.

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COMPETING INTERESTS

The authors have declared that no competing interests exist.

CONTRIBUTORS

J.L.W., D.K., K.D.M., I.S.K., and G.A. conceived the SMART PCM app. D.C., R.O., and L.W. built the SMART PCM app with clinical subject matter expertise provided by J.L.W. J.L.W. and D.C. participated in FHIR Connectathon 7. J.M. hosted the SMART on FHIR server at FHIR Connectathon 7. J.L.W. and M.J.R. conducted the user testing, and M.J.R. carried out the data analysis. S.Z. and H.Y. developed the NGS analysis expansion to SMART PCM. All authors contributed to the initial draft manuscript and approved the final manuscript.

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