

Accepted Manuscript

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PII: S0012-3692(18)30725-6

DOI: [10.1016/j.chest.2018.04.037](https://doi.org/10.1016/j.chest.2018.04.037)

Reference: CHEST 1717

To appear in: *CHEST*

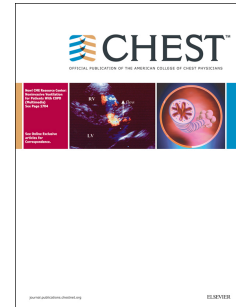
Received Date: 7 February 2018

Revised Date: 6 April 2018

Accepted Date: 27 April 2018

Please cite this article as: Sanchez-Pinto LN, Luo Y, Churpek MM, Big Data and Data Science in Critical Care, *CHEST* (2018), doi: 10.1016/j.chest.2018.04.037.

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Big Data and Data Science in Critical Care

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Word count: 3,495

Conflicts of Interest: MMC has a patent pending for risk stratification algorithms for hospitalized patients. He is also supported by a career development award from the National Heart, Lung and Blood Institute and a research project grant program award from the National Institute of General Medical Sciences. All others declare no conflicts of interest.

ABSTRACT

The digitalization of the healthcare system has resulted in a deluge of clinical Big Data and has prompted the rapid growth of data science in medicine. Data science, which is the field of study dedicated to the principled extraction of knowledge from complex data, is particularly relevant in the critical care setting. The availability of large amounts of data in the intensive care unit, the need for better evidence-based care, and the complexity of critical illness makes the use of data science techniques and data-driven research particularly appealing to intensivists. Despite the increasing number of studies and publications in the field, so far there have been few examples of data science projects that have resulted in successful implementations of data-driven systems in the intensive care unit. However, given the expected growth in the field, intensivists should be familiar with the opportunities and challenges of Big Data and data science. In this paper, we review the definitions, types of algorithms, applications, challenges, and future of Big Data and data science in critical care.

INTRODUCTION

The digitalization of the healthcare system is changing the way we practice medicine and conduct clinical research.^{1,2} The widespread implementation of electronic health records (EHRs) is paving the way for Big Data research and is bringing the world of data science to the patient's bedside.²⁻⁴ Within the healthcare system, the intensive care unit (ICU) presents a particularly convincing case for using data science to improve patient care.⁵ For one, the evidence supporting many of the interventions performed in the ICU is rather scarce and practice variability is abundant.^{5,6} In addition, the complexity of critical illness makes the traditional reductionist approach to medical research insufficient, that is, single drug intervention trials or single pathway biomarker studies are unlikely to satisfy the clinical realities of the ICU.⁵ Critical care research requires an integrative approach that embraces the complexity of critical illness and the computational technology and algorithms that can make it possible.^{7,8} Moreover, the data required to do this are being generated and digitized in troves. EHRs, bedside monitors, medication pumps, and ventilators are continuously generating new minable data, and soon the advancement of modern molecular diagnostics will result in a deluge of "omics" data derived from the genome, transcriptome, microbiome, and a long list of other "-omes" (Figure 1).

As Big Data and data science gradually infiltrate most aspects of clinical research and – ultimately – clinical care in the ICU, it is increasingly evident that intensivists should be familiar with the promise and perils of these approaches. In this paper, we will review the definitions, types of algorithms, applications, challenges, and future of Big Data and data science in critical care.

DEFINITIONS IN DATA SCIENCE

Big Data can be defined as digital data that is generated in high volume and high variety and that accumulates at high velocity, resulting in datasets too large for traditional data-processing systems.^{2,9} In practice, Big Data in healthcare depends on both the breadth and the depth of the data being captured. For example, administrative healthcare datasets with few data elements per patient record (low depth) are usually considered Big Data problems when they contain millions of records (wide breadth). On the other hand, when applying next-generation sequencing and other "omics" approaches (high depth), just a few dozen patients can become a Big Data problem (narrow breadth).¹⁰

Data science can be defined as "the set of fundamental principles that support and guide the principled extraction of information and knowledge from data."⁹ A closely related term is *data mining*, which is the actual extraction of knowledge from data via machine learning algorithms that incorporate data science principles.⁹ *Machine learning* is the field of study that focuses on how computers learn from data and the development of algorithms that make this learning possible.¹¹ Finally, another important concept in data science is *domain expertise*, which in healthcare can be defined as the understanding of real-world clinical problems and the realities of patient care that help frame and contextualize the application of data science to healthcare problems.^{8,9,11}

These and other pertinent definitions in data science are presented in Table 1.

THE ECOSYSTEM OF DATA SCIENCE IN HEALTHCARE

The data revolution in healthcare would not be possible if it were not for several key developments, including: (1) the data science movement that has transformed other industries; (2) the extraordinary growth in computational power; (3) the availability of open source tools and low-cost equipment to perform advanced analyses; and (4) the increasing availability of educational resources and advanced degrees in data science and related fields.¹² Open source programming and scripting languages, such as R and Python, have extensive libraries of statistical packages and machine learning algorithms that are relatively easy to use by researchers with some training and have greatly democratize the access to data science techniques. Educational resources, such as graduate programs in data science, are also increasingly available. Notable amongst these are massive open online courses (MOOCs), which have led to increased popularity of data science and the applications of machine learning techniques to real-world problems.¹³ There are now a multitude of courses from prestigious institutions that cover data science and machine learning, and many of them are free. This provides the interested researcher with educational opportunities from world-class experts at the touch of a button. Other websites contain user-created code to run machine learning algorithms from scratch (e.g., <https://github.com/>), or host data science competitions for participants around the world (e.g., <https://www.kaggle.com/>). This rich online environment is ideal for data scientists to learn and grow, with the field of medicine benefitting from advances shared around the world freely through the internet.

TYPES OF ALGORITHMS IN DATA SCIENCE

Machine learning algorithms are generally divided into two categories: supervised and unsupervised learning algorithms.¹¹ Semi-supervised algorithms represent a hybrid of the two but have been used less often in healthcare problems. Finally, deep learning algorithms defy this classification, even though they derive from artificial neural network algorithms which are generally classified as supervised algorithms. The most defining characteristic of deep learning is their focus on learning data representations (or features) that can then be used in supervised, unsupervised, or semi-supervised problems. We review these types of algorithms in more detail below (and also in Figure 2 and Table 2).

Supervised learning algorithms

Supervised learning algorithms are used to uncover the relationship between variables of interest and one or more target outcomes.^{11,14} For supervised problems, the target outcome(s) must be known. For example, if researchers want to know whether a set of clinical features (e.g., vital signs, laboratory tests, etc.) can predict ICU mortality, they could apply a supervised learning algorithm to a dataset in which each patient record contains the set of clinical features of interest and a label specifying their outcome (“survived” or “not survived” in this case) (Table 1). Examples of supervised learning algorithms include regression-based methods (e.g. linear and logistic regression, lasso, elastic net), tree-based methods (e.g.,

classification and regression trees, random forest, gradient boosted trees), k-nearest neighbor, artificial neural networks, and support vector machines (Table 2).¹⁵

Unsupervised learning algorithms

Unsupervised learning algorithms are used to uncover naturally occurring patterns or groupings in the data, without targeting a specific outcome.¹¹ The most compelling use case of unsupervised learning in healthcare is in precision medicine, in which the goal is to uncover subsets of patients who share similar clinical or molecular characteristics and are—in theory—more likely to respond to targeted therapies directed at their shared underlying pathobiology.^{16,17} For example, an unsupervised learning algorithm may be used to uncover subgroups of patients with sepsis that have distinct molecular and clinical characteristics and will respond differently to specific drugs, such as corticosteroids.¹⁸ Some examples of unsupervised learning algorithms include clustering algorithms (e.g., hierarchical clustering, k-means clustering), latent class analysis, and principal component analysis (Table 2).^{11,14,19}

Deep learning algorithms

Deep learning algorithms are designed to extract meaningful features from the data in order to represent information in an increasingly higher order of hierarchical complexity in the form of stacked layers of nodes (or “neurons”).²⁰ For example, if the input is a photo of several people, the first layer of nodes might simply extract straight lines, curves, and color hues. Deeper layers may combine some of those lines, curves, and hues to represent eyes, noses, ears, and other more complex features. After deeper layers of nodes have perceived increasingly more complex features in an unsupervised way, they can then be used to perform specific tasks, like match the faces in the photo to certain specific people with known features. In medical applications, deep learning has been used, for example, to detect diabetic retinopathy on funduscopy images,²¹ detect cancer in skin photographs,²² or predict clinical outcomes using EHR data.^{25,26}

DATA SCIENCE APPLICATIONS IN CRITICAL CARE

Predictive and prognostic models

The most common applications of data science to critical care problems are predictive and prognostic models using supervised learning algorithms. Although identical from a modeling perspective, predictive and prognostic models can be distinguished semantically by the fact that *predictive models* are generally trained to predict the likelihood of a condition, event, or response, while *prognostic models* are specifically trained to predict the likelihood of a condition-related endpoint or outcome, such as mortality (Table 1).^{16,27} This distinction, however, is not always clear in the literature and, depending on the use case, might be irrelevant.

One of the oldest and best-known prognostic models to estimate risk of mortality in ICU patients is the Acute Physiology and Chronic Health Evaluation (APACHE) score, which was first developed in the 1980's by Knaus and colleagues using logistic regression.^{28,29} Since then, many other groups have developed predictive and prognostic models using larger, more granular datasets and applying modern machine learning methods. For example, Churpek and colleagues developed a logistic regression model in a dataset of over 250,000 hospital

admissions that accurately estimated the risk for ICU transfer, cardiac arrest, or death in ward patients.³¹ In a follow-up study, the same group showed that more modern machine learning methods, such as random forests and gradient boosted machines, could more accurately predict clinical deterioration compared to classical logistic regression.¹⁵ In another example, Joshi and colleagues used 54 clinical variable time series to predict 30-day mortality in ICU patients in the publicly-available Multiparameter Intelligent Monitoring in Intensive Care (MIMIC) dataset.³² They clustered the physiologic measurements into organ specific patient states and achieved a state-of-the-art 30-day mortality prediction area under the receiver operating characteristic curve (AUC) of 0.91.

Predictive models aimed at identifying patients with specific condition or those more likely to respond to a specific therapy are more commonly used in the field of oncology, with multiple examples of biomarker-based models used to diagnose particular subtypes of cancer that respond to targeted therapy.^{4,16} However, there are some examples in the critical care literature, particularly in sepsis and septic shock.^{33,34} For example, Wong and colleagues used a combination of a classification and regression tree-based biomarker risk model and gene expression profiles in pediatric sepsis patients to identify a subgroup of patients who were more likely to benefit from corticosteroids.¹⁸

Clustering and phenotyping

Unsupervised learning algorithms in critical care have mainly been used to uncover naturally occurring subgroups or clusters of patients who share similar clinical and/or molecular characteristics. These clusters are oftentimes called phenotypes, subphenotypes, or subtypes, although there is still little consensus on the terminology.¹⁶ For example, Calfee and colleagues applied Latent Class Analysis and identified two subphenotypes of acute respiratory distress syndrome (ARDS) using clinical and cytokine data from two ARDS randomized controlled trials.³⁷ The subphenotypes identified had distinct differences in inflammatory profiles, response to ventilator strategies, and clinical outcomes. Knox and colleagues used self-organizing maps and k-means clustering to identify four distinct clusters of patients with sepsis-associated multiple organ dysfunction syndrome that were independently associated with outcomes after adjusting for severity of illness.³⁸ Luo and colleagues analyzed multiple physiological variable trends of patients in the MIMIC dataset and applied non-negative matrix factorization to group related trends, which were shown to effectively predict 30-day mortality while maintaining model interpretability.³⁹ Finally, Vranas and colleagues applied clustering analysis to discover and validate six clinically recognizable subgroups of ICU patients that differed significantly in all baseline characteristics and clinical trajectories despite sharing common diagnoses.⁴⁰

Applications with non-traditional data types

Natural language processing

Much of the data used in critical care studies, such as vital signs or laboratory test results, are structured data that can be easily entered into a relational database or spreadsheet and be sorted, summarized, etc. However, there is a significant amount of clinical information contained in the form of unstructured clinical narratives (e.g., progress notes, discharge summaries, nursing notes, diagnostic reports, etc.).⁴¹ Methods for analyzing narrative data,

generally known as natural language processing (NLP), are designed to extract features from texts that can be then used in task-specific algorithms for different purposes, for example, prognostic modeling. Lehman and colleagues used clinical data and unstructured progress notes from the first 24 hours of ICU admissions to estimate the risk of in-hospital mortality.⁴² They inferred topic models from progress notes and achieved an AUC of 0.82, which was superior to severity of illness scores based only on structured clinical variables. Ghassemi and colleagues further investigated the prognostic power of topics as features from the first 24 hours and achieved an AUC of 0.85 for in-hospital mortality when combining text and structured data.⁴³ Weissman and colleagues applied NLP to analyze discharge documents of ARDS survivors and found that ARDS itself is rarely mentioned in those documents, as opposed to more frequent mentions of “mechanical ventilation” and “ICU stay”.⁴⁵ On the other hand, their NLP based document classifier reported 100% accuracy for ARDS identification, suggesting that NLP can be used to effectively identify patients with certain types of conditions.

Physiologic waveform analysis

Physiologic waveform data from bedside monitors and wearable devices is increasingly being used in data science studies in critical care. Many institutions collect and store physiologic monitor data, such as electrocardiography, photoplethysmography, impedance pneumography, invasive arterial manometry, end-tidal capnography, and electroencephalography. The publicly available MIMIC databases contains physiologic waveform data for ICU patients at Beth Israel Deaconess Medical Center, which has facilitated the development of the state-of-the-art waveform analysis in the field.⁴⁶ For example, researchers have used waveform data to estimate cardiac output data using pulse contour analysis techniques,⁴⁷ detect hypovolemia using photoplethysmography data,⁵¹ and predict hyperlactatemia using combined physiologic data.⁵⁴

Image analysis

The advancement in the field of deep learning, which is particularly useful for image analysis, has resulted in a rapid increase in the number of studies in this area in the last few years.⁵⁵ However, none of the current published studies have tested the usefulness of automated image analysis in an ICU setting. The rapid growth of this field, however, will undoubtedly result in many use cases applicable to critical care situations. Perhaps most pertinent to critical care clinicians is the advancement in techniques to detect pulmonary pathology in chest radiographs,^{23,24} as well as normal and abnormal findings in brain and abdominal imaging.⁵⁵ These techniques could be particularly helpful in ICUs with limited availability of specialists who can accurately interpret radiographic images in a timely fashion, but their effectiveness and safety should first be thoroughly tested before any clinical implementation may be considered.

CHALLENGES AND PITFALLS OF DATA SCIENCE IN CRITICAL CARE

Like most emerging technologies, the products of data science research in critical care will undoubtedly go through a series of hype and disillusionment cycles before becoming accepted and proven assets in the study and care of critically ill patients. One of the first challenges that data science faces in critical care is that, despite the increasing number of

studies and publications in the field, so far there have been few examples of data science projects that have resulted in successful implementations of data-driven systems in the ICU.¹¹ This lack of exposure in the clinical setting inevitably results in a degree of mistrust by clinicians in these data-driven systems.^{58,59} While clinicians are happy to use similar systems to browse their smart TVs, shop online, or interact with social media apps, they are wary of the idea of sharing clinical decision-making responsibilities with machine learning algorithms, particularly if they view them as “black boxes”.⁵⁹ It is likely that only the implementation of well-designed, interpretable, and effective data-driven systems in the ICU will make clinicians start to gain trust in them. Furthermore, the implementation of these data-driven systems must be performed under the rigorous auspices of well-controlled experimental studies, including (but not limited to) simulation testing, pre- and post-intervention studies, and randomized controlled trials. The medical informatics literature has good examples of using scientifically rigorous approaches to the implementation and testing of digital solutions like clinical decision support tools and can serve as a model to follow.⁶⁰⁻⁶²

Clinicians and researchers appraising a data-driven system and the literature that supports it must be aware of common pitfalls that can raise concerns about its value. The effectiveness of a data-driven system goes beyond a measure of performance, like an AUC or a p-value. In order to be effective, a data-driven system must produce actionable outputs for the right patients, at the right time. For example, the output can be predictive information that can help a clinician decide the most effective treatment for a particular patient as soon as a diagnosis is made. Furthermore, when evaluating the clinical implementation of such a system, it is important to know whether it has been tested in an experimental setting and whether it has shown a meaningful impact in a population similar to the one for which it is being considered.

Unfortunately, bad data science abounds. We must make a collective effort to ensure that only good data science evolves into data-driven systems that can be safely tested and used in critically ill patients. The ease of access to large amounts of data and computing power can lead to data mining “fishing expeditions” that can result in low-quality research.³ Poorly framed clinical problems, bad data, or debatable methodologies will result in flawed data science, and it can create more problems than it solves.^{3,6,59} Using epidemiological best practices to analyze retrospective data, including thoughtfully adjusting for confounding variables, is just as important in large datasets as it is in smaller ones. In addition, a model may fit well only on the training data but generalize poorly to other data, a phenomenon known as overfitting (see Table 1). Overfitting may happen when algorithms learn from idiosyncrasies –or noise– in the training data and techniques such as cross-validation and regularization can be used to mitigate this problem.^{63,64}

Poorly implemented digital technologies can harm patients,⁶⁵ and only a rigorous approach to their evaluation and implementation can mitigate this risk. Partnerships between data scientists, clinical domain experts, medical informaticians, and implementation science specialists will result in more effective and safer data-driven systems. Clinicians with data science skills, clinical research expertise and an intimate knowledge of the clinical realities in the ICU can help data science teams capture the right data, address the right clinical problems, and produce the right actionable knowledge.¹⁰ Furthermore, clinician input can help minimize the number of unnecessary alerts or prompts these systems might produce, thereby reducing

the risk of alert fatigue, which is another common problem amongst front-line providers working with novel digital technologies.⁶⁶

Another common concern amongst clinicians is the perceived loss of autonomy in the face of increasingly more sophisticated computational systems. This is despite the fact that clinicians will readily acknowledge that the complexity of medicine nowadays far exceeds the capacity of the unaided human mind and that perhaps these novel computational systems can help manage some of this complexity.¹⁰ To put it in perspective, humans typically make decisions using less than six data points, since anything more than that becomes cognitively too expensive.⁶⁷ Yet, an ICU patient can generate thousands of data points in a single day, and when you add fatigue, interruptions, and the clinicians' own cognitive biases, it is not surprising that many clinical decisions end up being suboptimal.⁶⁸ On the other hand, computers can sift seamlessly through tens of thousands of data points, they can easily analyze complex non-linear interactions between variables, they never sleep, and can multitask effortlessly. However, clinical thinking and medical decision-making are not reproducible by current technologies.¹⁰ The qualitative aspect of clinical decision-making –the so-called “art of medicine”– is impossible to model quantitatively. Furthermore, many factors influencing clinical decision-making, including clinical, societal, and personal factors, are not necessarily reflected in the digital records, so any output from a data-driven system will need to be first evaluated, interpreted, and enriched by clinicians before any action is taken. However, in order to achieve a successful partnership between clinicians and computers, we must first improve the skills of bedside clinicians at interpreting and using the output from these data-driven systems.⁵⁹

Finally, another challenge faced by data science teams in critical care is balancing the need for data openness and reproducibility with the demand for data privacy and security. The open data science movement calls for transparent and reproducible research with seamless data sharing across institutions. Indeed, a recent study showed an alarming lack of reproducibility in data science studies using the same ICU data, which suggests that algorithms, study procedures, computer code and even datasets should be openly available in order to ensure reproducibility.⁶⁹ However, this data openness must not result in poor data governance, lack of data security, or loss of confidentiality, all of which are necessary in order to perform ethical research and maintain public trust.⁷⁰

THE FUTURE OF BIG DATA AND DATA SCIENCE IN CRITICAL CARE

We imagine a future in critical care where data-driven systems and clinicians work hand-in-hand. Large quantities of clinical, physiologic and “omics” data are analyzed by computational systems and served to the bedside clinicians in the form of manageable, interpretable, and actionable knowledge that augment the clinician's decision-making capacity. Predictive models perform diagnostic and therapeutic recommendations, while clinicians contextualize these recommendations and coordinate their implementation. False alerts are kept to a minimum and systems are continuously improved through a collaborative and scientifically rigorous approach.

Data science can be transformative. There is a real opportunity that this scenario will become a reality in the near future, but there is still a lot of work ahead of us. Our patients entrust us with their precious data and we –clinicians, researchers, data scientists, and leaders in critical care– have an obligation to use it in the best possible way.

TABLES

Term	Definition
Big Data	Digital data that is generated in high volume and high variety and that accumulates at high velocity, resulting in datasets too large for traditional data-processing systems.
Data science	The set of fundamental principles that support and guide the principled extraction of information and knowledge from data.
Data mining	The extraction of knowledge from data via machine learning algorithms that incorporate data science principles.
Domain expertise	The understanding of real-world problems in a given domain (e.g., critical care medicine) that helps frame and contextualize the application of data science to solve these problems.
Machine learning	The field of study that focuses on how computers learn from data and the development of algorithms that make this learning possible.
Features	The data elements, also known as <i>independent variables</i> , used to train a model. Features can be simple transformations of the raw data (e.g., average heart rate in the last 24 hours) or complex transformation like the ones performed by neural networks (see Table 2).
Outcomes	The data elements, also known as <i>dependent variables</i> , that represent the target for training in a supervised learning model. Outcomes can be categorical (e.g., yes/no) or continuous (e.g., length of hospital stay). Categorical binary outcomes are the most common in medicine (e.g., died or alive by 28 days). Binary outcomes are typically represented as a Boolean logic (i.e., true/false or 1/0) but can also be represented using fuzzy logic (i.e., a range of probabilities -or degrees of truth- between 0 and 1).
Supervised learning	Algorithms that are used to uncover the relationship between a set of features and one or more known outcomes.
Unsupervised learning	Algorithms that are used to uncover naturally occurring patterns or groupings in the data, without targeting a specific outcome.
Model training	The process through which machine learning algorithms develop a model of the data by learning the relationships between features and, in supervised learning, between features and outcomes. This is also referred to as <i>model derivation</i> or <i>data fitting</i> .
Model validation	The process of measuring how well a model fits new, independent data. For example, evaluating the performance of a supervised model at predicting an outcome in new data. This is also referred to as <i>model testing</i> .
Predictive model	A model generally trained to predict the likelihood of a condition, event, or response. The Food and Drug Administration specifically considers <i>predictive strategies</i> as those geared towards identifying groups of patients more likely to respond to an intervention.
Prognostic model	A model specifically trained to predict the likelihood of a condition-related endpoint or outcome, such as mortality. In general, the goal is to estimate a prognosis given a set of baseline features, regardless of what ultimately leads to the outcome.
Overfitting	The phenomenon that occurs when an algorithm learns from idiosyncrasies in the training data, usually referred to as <i>noise</i> . Noisy data is data that is randomly present in the training dataset but does not represent the generalizable truth (usually referred to as <i>signal</i>) that explains the relationships between the features and the outcomes. Overfitting will generally lead to poor performance of the model in an independent

	validation dataset.
Digitization	The conversion of something analog or physical (e.g., paper documents, printed images, etc.) into a digital format (i.e., bits or 1s and 0s)
Digitalization	The wide adoption of digital technologies by an organization in order to leverage their digitized data with the goal of improving operations and performance. The adoption of electronic health records and other digital technologies (e.g., picture archiving and communication systems for medical images, pharmacy management systems, billing systems, etc.) are examples of digitalization in healthcare.
Data curation	The process of integrating data from different sources, structuring it, authenticating it, and annotating it in order to ensure its quality, add value, and facilitate its use and reuse.
Structured data	Data –usually discrete or numeric– that is easy to search, summarize, sort, and quantify. Examples include vital signs (e.g., heart rate) or laboratory test results (e.g., complete blood count).
Unstructured data	Data that does not conform to a pre-specified structure, like a written narrative, images, video, or audio. Unstructured data is generally harder to search, sort, and quantify. Examples include clinician notes, pathology slides, and radiology images.

Table 1. Definitions of common terms in data science

Algorithm class	Examples	Description
Classic regression	Linear regression, logistic regression	Linear regression is a supervised learning algorithm that models the relationship between one or more features and a continuous outcome by fitting a regression line that minimizes the sum of all the residuals, which are the distances between each feature in the training data and the line being fitted to model them. Logistic regression is a generalization of the linear model that uses the logistic function to estimate the probability of a binary outcome. To do this, the fitted sigmoid-shaped curve of the logistic function maps the feature values into a probability between 0 and 1.
Regularized regression	Lasso, ridge regression, elastic net	An extension of the classic regression algorithms in which a penalty is imposed to the fitted model to reduce its complexity and decrease the risk of overfitting (see Table 1).
Tree-based	Classification and regression trees, random forest, gradient boosted trees	A class of supervised learning algorithm based on decision trees. Decision trees are a sequence of “if-then-else” splits that are derived by iteratively separating the data into groups based on the relationship of the features with the outcome. Random forest and gradient boosted trees are example of ensemble tree models. <i>Ensemble models</i> combine the output of many trained models to estimate an outcome.
Support vector machines	Linear, polynomial, radial basis kernel	A class of supervised learning algorithms that represents the data in a multidimensional feature space and then fits a “hyperplane” that best separates the data based on the outcomes of interest.
K-nearest neighbor	K-nearest neighbor	A type of supervised learning algorithm that represents data in multidimensional feature space and uses local information about observations closest to a new example to predict the outcome for that example.
Bayesian	Naïve Bayes, Bayesian network	A class of supervised learning algorithms that use Bayes’ theorem of conditional probability, which is the probability that something will happen given that something else has already occurred. In general, Bayesian algorithms work by iteratively updating the probability of an outcome (or posterior belief) given new data.
Neural network	Artificial neural network, deep neural network	A class of non-linear algorithms built using layers of nodes that extract features from the data and perform combinations that best represent the underlying structure, usually to predict an outcome. Neural networks can be shallow (like a perceptron with two layers) or deep (multiple layers), which form the basis for the field of <i>deep learning</i> .
Dimensionality reduction algorithms	Principal component analysis, linear discriminant analysis.	A class of unsupervised learning algorithms that exploit the inherent structure in the data to describe data using less information. Principal components, for example, summarize a large set of correlated features into a smaller number of representative features.
Latent class analysis	Latent class analysis	A type of unsupervised learning algorithm that identifies unseen subgroups, or latent classes, in the data. Class membership is unknown for each example so the probability of class membership is indirectly estimated by measuring the patterns in the data.
Cluster analysis	K-means, hierarchical cluster analysis	A class of unsupervised learning algorithm that use the inherent structures in the data to best organize the data into subgroups of maximum commonality based on some distance measure between features.

Table 2. Examples of algorithms used in data science**FIGURE LEGENDS**

Figure 1. Some of the major sources of Big Data in the intensive care unit (ICU). The term “*Omics*” refers to the data derived from modern molecular techniques, e.g. genomics, transcriptomics, proteomics, metabolomics, microbiomics, etc.

Figure 2. Types of machine learning algorithms applicable to critical care. Supervised learning algorithms (**A**) can be used, for example, to uncover the relationship between patient clinical features (such as laboratory tests and vital signs) and mortality in order to predict the outcome in future cases. Unsupervised learning algorithms (**B**) can be used to uncover naturally occurring groupings or clusters of patients based on their clinical characteristics, without targeting a specific outcome. Deep learning algorithms (**C**) can be used, for example, to extract meaningful features from imaging data (such as a chest X-ray) in order to represent information in an increasingly higher order of hierarchical complexity and be able to make predictions, such as the presence of pathologic findings.

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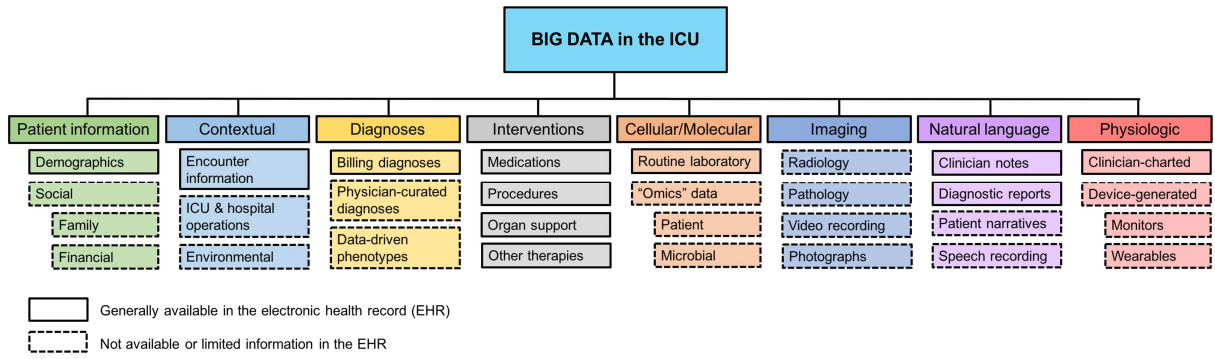
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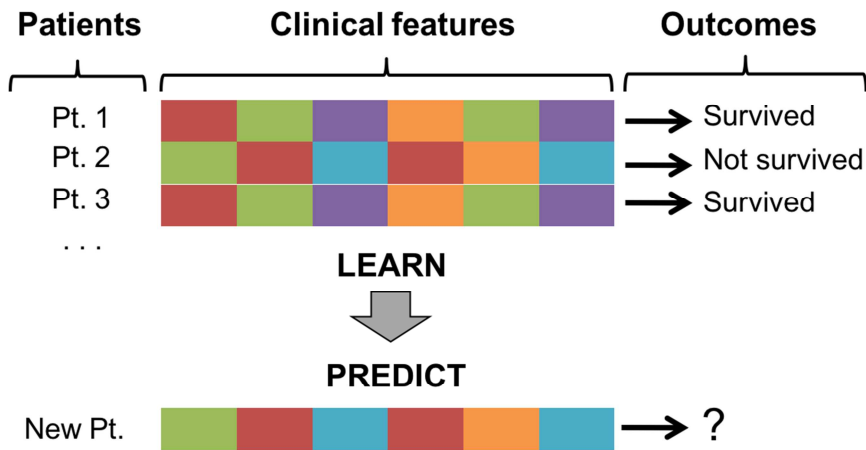
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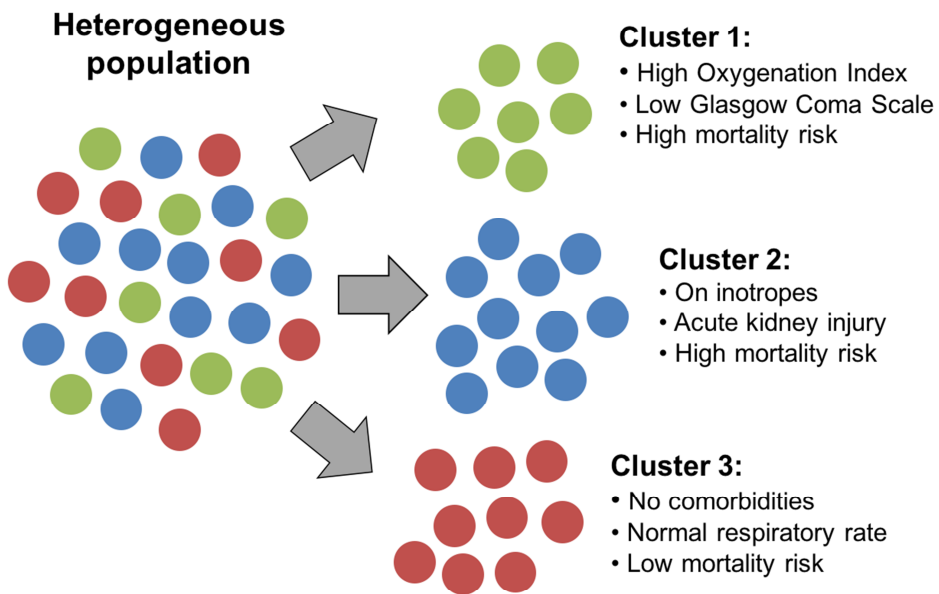


ACCEPTED MANUSCRIPT

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