ARTIFICIAL INTELLIGENCE IN HEALTH CARE

RICHARD VAN ECK associate dean for teaching and learning monson endowed chair for medical education

PRESENTATION FOR THE DAKOTA CONFERENCE ON RURAL AND PUBLIC HEALTH

1

DISCLOSURES

- No financial disclosures
- Some of this information comes from an MIT online course, including examples and some figures and images. These are identified and credited.

• OBJECTIVES

- Define supervised and unsupervised learning
- Define natural language processing, deep learning, neural networks
- Define the process by which supervised learning models are created, trained, and deployed
- List key considerations for designing deep learning models, including cost, bias, risk, explainability, and interpretability
- List examples of how Al is being used in health care environments.

3

OVERVIEW

- Defining Al
 - Supervised Machine Learning
 Natural Language Processing
 Neural Networks/Deep Learning
 - Unsupervised Machine Learning
 Generative Al
- Customized Al Solutions
 - Training Data Sets
 - Annotation
 - Curation

- Risk, Bias, Best Practices
 - Interpretability/Explainability
 - Adversarial Training
- Case Studies of Al in Health Care
 - Hospital Optimization
 - Reducing Burden
 - Augmenting Clinical Workflows
 - Risk Stratification
 - Mental Health
 - Invisibles













SUPERVISED MACHINE LEARNING

- Machines learn from curated, annotated datasets
- Humans do the curation and annotation
- Humans are involved in monitoring results and refining the models























REDUCING BURDEN, AUGMENTING CLINICAL WORKFLOWS, AND IMPROVING OUTCOMES



BENEFITS OF AI

- Al often seen as a barrier between doctor and patient
 - But can remove barriers and create time for more patient interaction
- Potential benefits
 - Remove burden of repetitive tasks
 - Augment patient workflows
 - Improve outcomes through customized care





MEDICAL DOCUMENTATION AT POINT OF CARE

- Chief complaints
 - Can vary, which makes it harder to standardize
 - Many "downstream" tasks rely on chief complaint
 Enrolling patients in clinical trials
 - Conducting retrospective QI studies
 - Take time to generate
- Increase standardization and save time?
- Redesigned workflow
 - Nurses normally assign chief complaint first
 - Instead, had nurse asks questions, write triage note
- Al processes note in real-time
 Suggests chief complaint











ER: CHIEF COMPLAINT TIME SAVINGS

- Initially took 11.6 keystrokes to enter chief complaint
- After Al use, took 0.6 keystrokes
 - Sometimes chief complaint was in the top five listed, so no typing in the text field at all

27







BREAST CANCER AND AI

- Dr. Connie Lehman
 - Director of Breast Imaging and Co-Director of the Avon Comprehensive Breast Evaluation Center, Massachusetts General
- Breast density
 - Federal mandate to inform women with high density
 - Higher risk of tumors, tumors are harder to detect
- Radiologists are inconsistent
 - Classifications of high density ranged from 6% to 85% for same set of images





Images contain billions of data points

- Pixels per image
- Multiple images from different angles
- Most of that data is not used
 - Condensed into 1-2 page summary
 - Further condensed into a few categories • Cancer grade, hormonal status
- 4 women with same categorization
- 3 were fine, 1 metastatic recurrence Image from lecture by Regina Barzilay, MIT Delta Electronics



Image from lecture by Regina Barzilay, MIT Delta Electronics Professor of EE and Computer Science, course on AI in Health Care.















	Λ			
		Li M, Ladner D, Miller S, Classen D. Identifying hospital patient safety problems in real-time with electronic medical record data using an ensemble machine learning medel. Int J Clin Med Inform 2018;1(1):43-58.	The adverse event risk score at the 0.1 level could identify 57.2% of adverse events with 26.3% accuracy fram 9.2% of the validation sample. The adverse event risk score of 0.04 could identify 85.3% of adverse events	
		Murth HJ, Frizhenry F, Matheny MB, Gentry N, Kotter KJ, Crimin K, et al. Automated identification of postoperative complications within an electronic medical record using natural language processing. JAMA 2011 Aug 24;306(8):848-855. doi: 10.1001/jama.2011.1204	NLP identitied 82% of acute renal tailure cases compared with 38% for patient satety indicators. Similar results were obtained for venous thromboembolism (59% vs 46%), pneumonia (64% vs 5%), sepsis (89% vs 34%), and postoperative myocardial infarction (91% vs 89%)	
]/{		Wong Y, Coiera E, Runciman W, Magrabi F. Using multiclass classification to automate the identification of patient safety incident reports by type and severity. BMC Med Inform Decis Mak 2017 Jun 12;17(1):84 doi: 10.1186/s12911-012-0483-8	For severity level, the F score for severity assessment code (SAC) 1 (extreme risk) was 87.3 and 64% for SAC4 (low risk) on balanced data. With stratified data, a high recall was achieved for SAC1 (82.8%-84%), but precision was poor (6.8%-11.2%). High-risk incidents (SAC2) and medium-risk incidents (SAC3) were often misclassified. Reports about fails, medications, pressure injury, aggression, and blood tests were identified with high recall and precision	
		Rosenbaum M, Baron J. Using Machine Learning-Based Multianalyte Delta Checks to Detect Wrong Blood in Tube Errors. Am J Clin Pathol 2018 Oct 24;150(6):555-566. doi: <u>10.1093/cicp/cay085</u>	In contrast to the univariate analysis, the best performing multivariate delta check model (SVM) identified errors with a high degree of accuracy (0.97)	
		McKnight SD. Semi-supervised classification of patient safety event reports. J Patient Saf 2012 Jun;8(2):60-64. doi: 10.1097/PTS.0b013e31824ab987	The semisupervised model categorized patient safety reports into their appropriate patient safety topic and avoided over- laps; 85% of unlabeled reports were assigned correct labels. It helped NCPS analysts to develop policy and mitigation decisions	
		Marella WM, Sparnon E, Finley E. Screening Electronic Health Record-Related Patient Safety Reports Using Machine Learning. J Patient Saf 2017 Mar;13(1):31-36. doi: 10.1097/PTS.000000000000104	The NB kernel performed best, with an AUC of 0.927, accu-racy of 0.855, and F score of 0.877. The overall proportion of cases found relevant was compara-ble between manually and automatically screened cases; 334 reports identified by the model as relevant were identified as not relevant, implying a false-positive rate of 13%. Manual screening identified 4 incorrect predictions, implying a false-negative rate of 29%	
		Ye C, Wang O, Liu M, Zhang L, Xia M, Hao S, et al. A Real-Time Early Warning System for Monitoring Inpatient Mortality Risk: Prospective Study Using Electronic Medical Record Data. J Med Internet Res 2019 Jul 6521(7):e1371 9 doi: 10.2196/18719	The modified early warning system accurately predicted the possibility of death for the top $13.3\% (34/255)$ of patients at least 40.8 hours before death	
1	2 2	Fong A, Adams KT, Gount MJ, Howe JL, Kallegg KM, Ratwani RM. Identifying health information technology related safety event reports from patient safety event report databases. J Biomed Inform 2018 Oct;86:135-142 doi: <u>10.1016/j.bi.2018.09.007</u>	Unigram models performed better than Bigram and combined models. It identified HIT-related events trained on PSE free-text descriptions from multiple states and health care systems. The unigram IR model gave an AUC of 0.931 and nF1 score of 0.765. It also showed potential to maintain a faster runtime when more reports are analyzed. The final HIT model had less complexity and was more easily sharable	
	$\frac{1}{2}$	Simon ACR, Holleman F, Gude WT, Hoekstra JBL, Peute LW, Jaspers MWM, et al. Safety and usability evaluation of a web-based insulin self-titration system for patients with type 2 diabetes mellitus. Artif Intell Med 2013 Sep;59(1):23-31. doi: 10.1016/j.ortmed.2013.04.009	27 out of 74 (36.5%) PANDIT advice differed from those provided by diabetes nurses. However, only one of these (1.4%) was considered unsafe by the panel	
	$\langle \rangle$	² Song D, Chen Y, Min Q, Sun Q, Ye K, Zhou C, et al. Similarity-based machine learning support vector machine predictor of drug-drug interactions with improved accuracies. J Clin Pharm Ther 2019 Apr 18;44(2):268-275. doi:10.1111/jour.12766	The 10-fold crossvalidation improved the identification of drug-drug interaction with $AUC>0.97$, which is significantly greater than the analogously developed ML model (0.67)	9

Ò



	Li Q, Spooner SA, Kaiser M, Lingren N, Robbins J, Lingren T, et al. An end-to-end hybrid algorithm for automated medication discrepancy detection. BMC Med Inform Decis Mak 2015 May 06;15:37 doi: 10.1186/s12911-015-0160-8	The hybrid algorithm yielded precision (P) of 95.0%, recall (R) of 91.6%, and F value of 93.3% on medication entity identification, and P=98.7%, R=99.4%, and F=99.1% on attribute linkage. The combination of the hybrid system and medication matching system gave P=92.4%, R=90.7%, and F=91.5%, and P=71.5%, R= 65.2%, and F=68.2% on classifying the matched and the discrepant medications, respectively
	Carrell DS, Cronkite D, Palmer RE, Saunders K, Gross DE, Masters ET, et al. Using natural language processing to identify problem usage of prescription opioids. Int J Med Inform 2015 Dec;84(12):1057-1064. doi: <u>10.1016/j.ijmedinf.2015.09.002</u>	The NLP-assisted manual review identified an additional 728 (3.1%) patients with evidence of clinically diagnosed problem opioid use in clinical notes.
	Tinoco A, Evans RS, Staes CJ, Lloyd JF, Rathschild JM, Haug PJ. Comparison of computerized surveillance and manual chart review for adverse events. J Am Med Inform Assoc 2011;18(4):491-497 dei: 10.136/amiaini-2011-20018Z	CSS detected more hospital-associated infections than man- ual chart review (92% vs 34%); CSS missed events that were not stored in a coded format
	Oney A, Oney M, Abul O. Classification of nervous system withdrawn and approved drugs with TaxPrint features via machine learning strategies. Comput Methods Programs Biomed 2017 Apr;142:-91-96. doi:10.1016/j.cmpb.2017.02.004	The Gaussian SVM model yielded 78% prediction accuracy for the drug dataset, including all diseases. The ensemble of bagged tree and linear SVM models in-volved 89% of the accuracies for psycholeptics and psycho- analytic drugs
	Cai R, Liu M, Hu Y, Melton BL, Matheny ME, Xu H, et al. Identification of adverse drug-drug interactions through causal association rule discovery from spontaneous adverse event reports. Artif Intell Med 2017 Feb;76:7-15 doi: 10.1016/j.artmed.2017.01.004	CARD demonstrated higher accuracy in identifying known drug interactions compared to the traditional method (20% vs 10%); CARD yielded a lower number of drug combinations that are unknown to interact (50% for CARD vs 79% for association rule mining).
	Dandala B, Joopudi V, Devarakonda M. Adverse Drug Events Detection in Clinical Notes by Jointly Modeling Entities and Relations Using Neural Networks. Drug Saf 2019 Jan;42(1):135-146. doi: 10.1007;420:4-018-076-4-x	Joint modeling improved the identification of adverse drug events from 0.62 to 0.65
	Dey S, Luo H, Fokoue A, Hu J, Zhang P. Predicting adverse drug reactions through interpretable deep learning framework. BMC Bioinformatics 2018 Dec 28;19(Suppl 21):476 doi: <u>10.1186/s12859-</u> 018-2544-0	Neural fingerprints from the deep learning model (AUC=0.72) outperformed all other methods in predicting adverse drug reactions. The model identified important molecular substructures that are associated with specific adverse drug reactions
	Yang X, Bian J, Gong Y, Hogan WR, Wu Y, MADEx: A System for Detecting Medications, Adverse Drug Events, and Their Relations from Clinical Notes. Drug Saf 2019 Jan;42(1):123-133 doi: 10.1007/s4024-018-02-61-0	MADEx achieved the top-three best performances (F1 score of 0.8233) for clinical name entity recognition, adverse drug effect, and relations from clinical texts, which outperformed traditional methods
	Chapman AB, Peterson KS, Alba PR, DuVall SL, Patterson OV. Detecting Adverse Drug Events with Rapidly Trained Classification Models. Drug Saf 2019 Jan 16;42(1):147-156 doi: <u>10.1007/s40264- 018-0763-y</u>	The micro-averaged F1 score was 80.9% for named entity recognition, 88.1% for relation extraction, and 61.2% for the integrated systems
Ç	Lian D, Khoshneshin M, Street WN, Liu M. Adverse drug effect detection. IEEE J Biomed Health Inform 2013 Mar;17(2):305-311. doi: 10.1109/TITB.2012.2227272	Experimental results showed the usefulness of the proposed pattern discovery method by improving the standard baseline adverse drug reaction by 23.83%
مر	Huang L, Wu X, Chen JY. Predicting adverse side effects of drugs. BMC Genomics 2011 Dec 23;12 Suppl 5:511 doi: 10.1186/1471-2164-12-55-511	The proposed computational framework showed that an in silico model built on this framework can achieve satisfactory cardiotoxicity adverse drug reaction prediction performance (median AUC=0.771, accuracy=0.675, sensitivity=0.632, and specificity=0.789).























[°]EMERALD DEVICE

- Like Wi-Fi router
- Measures disturbances in electromagnetic waves throughout the home
- Al models interpret the patterns
- Breathing and heart rate
- As accurate as in-office measurement
- With more validity (in-situ)



Image and descriptions by Dina Katabi, Andrew and Erna Viterbi Professor of EE and Computer Science, MIT







WHY THE FUTURE OF AI IS NOT QUITE **BRIGHT ENOUGH** FOR SHADES

AND THOSE TOO YOUNG TO KNOW WHO THEY WERE)



EXPENSIVE AND TIME-CONSUMING

- Annotation is expensive
- Curation takes time and is easy to get wrong

 - Who makes up your population?
 Different imaging machines, preprocessing data (typos, IDC-9 vs IDC-
- Al needs to be connected to the EHR to be most effective
 - Requires significant expertise
 - If you don't have expertise, have to hire outside help
 AND must deidentify EHR data in ADDITION
- Complex problems require complex Al
 - Will need outside help in most cases
 - Interpretability will be challenging



UNDERSTANDING AND TRUST

- Al as a "Black Box"
- Power of Al: do things humans cannot do
 Or do them faster
 - Often makes it less understandable
- Working to build interpretable Al
 List the features to which it is "paying attention"
- Challenges of interpretability
 - Interpretable for doctor \neq interpretable for patient
 - Interpretable at what level of detail? • Biological prediction of drug target?
 - Full biological mechanism of processing by body?



HOW AI CAN GO WRONG EVEN WHEN DOING OUR BEST

- Risk stratification and hospital optimization
 Predict sickest patients and align resources and care
- Optum AI developed for use by hospitals
 Data vectors included IDC codes
 - More codes = more treatment = sicker patient
- Results
 - White people tended to be classified as higher risk
 - Black people tended to be classified as lower risk
- Why?
 - IDC codes confounded with insurance levels and cost
 - Better insurance = more treatment ≠ sicker people!
- Explainability
 - If Al said what it was "paying attention" to...

Obermeyer Z, Powers B, Vogeli C, Mullainathan S. Dissecting racial bias in an algorithm used to manage the health of populations. Science. 2019;366(6464):447-453. doi:10.1126/science.aax2342











THE (NEAR) FUTURE OF AI

• Al factories

- Automates data collection, preprocessing, labeling, augmentation
- Continually integrates and adjusts to incoming (new) data
- Preliminary model development
- Selects and "tunes" algorithms to refine and select best models
- Deploys testing frameworks to test models against validation datasets
- Monitors model performance, alerts to model drift and anomalies

• Will exponentially accelerate deep learning development and remove expertise and resource barriers

• Must be used by humans as productivity tool for SML, not as automomous stand-alone



WORKS CITED AND ADDITIONAL READINGS

- Alvarez-Melis, David and Tommi S. Jaakkola. "A Causal Framework for Explaining the Predictions of Black-Box Sequence-to-Sequence Models." ArXiv:1707.01943, November 14, 2017. http://arxiv.org/abs/1207.01943
- Athalye, Anish, Logan Engstrom, Andrew Ilyas, and Kevin Kwok. "Synthesizing Robust Adversarial Examples." Last modified June 7, 2018. http://arxiv.org/abs/1707.07397.
- Bahl, Manisha, Regina Barzilay, Adam B. Yedidia, Nicholas J. Locascio, Lili Yu, and Constance D. Lehman. 2018. "High-Risk Breast Lesions: A Machine Learning Model to Predict Pathologic Upgrade and Reduce Unnecessary Surgical Excision." Radiology 286, no. 3: 810–818. https://doi.org/10.1148/radiol.2017170549.
- Bennett, Charles L., Tammy J. Stinson, Victor Vogel, Lyn Robertson, Donald Leedy, Patrick O'Brien, Jane Hobbs, et al. 2000. "Evaluating the Financial Impact of Clinical Trials in Oncology: Results From a Pilot * Study From the Association of American Cancer Institutes/Northwestern University Clinical Trials Costs and Charges Project." Journal of Clinical Oncology 18, no. 15: 2805–2810. https://doi.org/10.1200/JCO.2000.18.15.2805.
- Bertsimas, Dimitris, and Jean Pauphilet. Forthcoming. "Holistic Hospital '
 Optimization." Management Science.

Ganin, Yaroslav, Evgeniya Ustinova, Hana Ajakan, Pascal Germain, Hugo Larochelle, François Laviolette, Mario Marchand, and Victor Lempitsky. "Domain-Adversarial Training of Neural Networks." Last modified May 26, 2016. https://arxiv.org/abs/1505.07818.

- Hartmann, Lynn C., Thomas A. Sellers, Marlene H. Frost, Wilma L. Lingle, Amy C. Degnim, Karthik Ghosh, and Robert A. Vierkant, et al. 2005. "Benign Breast Disease and the Risk of Breast Cancer." The New England Journal of Medicine 353: 229–237. https://www.neim.org/dai/full/10.1056/NEJMoa0444383.
- Kabelac, Zachary, Christopher G. Tarolli, Christopher Snyder, Blake Feldman, Alistair Glidden, Chen-Yu Hsu, Rumen Hristov, E. Ray Dorsey, and Dina Katabi. "Passive Monitoring at Home: A Pilot Study in Parkinson Disease." Digital Biomarkers 3, no. 1 (April 30, 2019): 22– 30. https://doi.org/10.1159/000498922.
- Khullar, Dhruv. 2019. "A.I. Could Worsen Health Disparities." Nev York Times, January 31, 2019. https://www.nytimes.com/2019/01/31/opinion/ai-biashealthcare.html.
- Shaaban, A.M., J.P. Sloane, C.R. West, F.R. Moore, C. Jarvis, E.M. Williams, and C.S. Foster. 2002. "Histopathologic Types of Benign Breast Lesions and the Risk of Breast Cancer. Case-Control Study." The American Journal of Surgical Pathology 26, no. 4: 421–430. https://www.ncbi.nlm.nih.gov/pubmed/11914619.
- Yala, Adam, Regina Barzilay, Laura Salama, Molly Griffin, Grace Sollender, Aditya Bardia, Constance Lehman, et al. 2017. "Using Machine Learning to Parse Breast Pathology Reports." Breast Cancer Research and Treatment 161: 203–211. https://doi.org/10.1007/s10549-016-4035-1.



