

If almost 60 percent of our members say that they spend three hours or more of every working day on administrative tasks, they cannot be there for their patients during this time. I simply think it's a scandal how much manpower and working time is wasted on data collection and documentation.

- *Chairman of the „Marburger Bund“, Dr. Susanne Johna (2022)*

[Ma22] Marburger-Bund: Zu wenig Personal, zu viel Bürokratie, unzulängliche Digitalisierung (quotation translated from German)

Evaluating Adapter-based Knowledge-enhanced Language Models in the Biomedical Domain

Alexander Fichtl

16.10.2023, Master Thesis Final Presentation

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Department of Computer Science
School of Computation, Information and Technology (CIT)
Technical University of Munich (TUM)
www.matthes.in.tum.de

1. Motivation

2. Background

- Knowledge-enhanced Language Models (KELMs)
- Adapters
- Evaluation Tasks

3. Research Questions

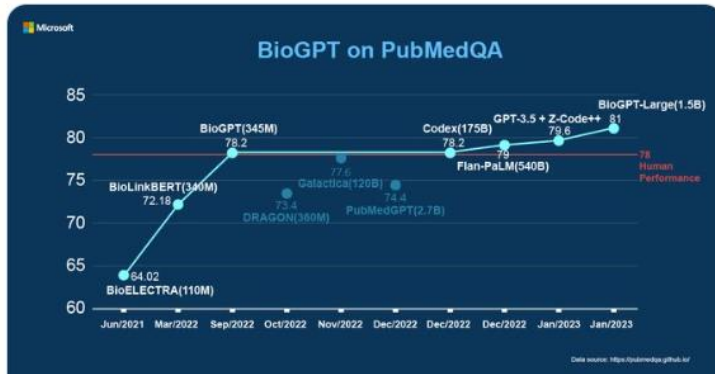
4. Methodology and Results

- Literature Review
- Model Experiments
- Research Survey

5. Shortcomings and Future Research



BioGPT, a domain-specific generative model pre-trained on large-scale biomedical literature, has achieved human parity, outperformed other general and scientific LLMs, and could empower biologists in various scenarios of scientific discovery. Learn more: msft.it/6014eAnLq



oup.com
BioGPT: pre-trained transformer for biomed text generation & mining

Original Investigation

April 28, 2023

Comparing Physician and Artificial Intelligence Chatbot Responses to Patient Questions Posted to a Public Social Media Forum

John W. Ayers, PhD, MA^{1,2}; Adam Poliak, PhD³; Mark Dredze, PhD⁴; et al

Key Points

Question Can an artificial intelligence chatbot assistant, provide responses to patient questions that are of comparable quality and empathy to those written by physicians?

Findings In this cross-sectional study of 195 randomly drawn patient questions from a social media forum, a team of licensed health care professionals compared physician's and chatbot's responses to patient's questions asked publicly on a public social media forum. The chatbot responses were preferred over physician responses and rated significantly higher for both quality and empathy.

Meaning These results suggest that artificial intelligence assistants may be able to aid in drafting responses to patient questions.

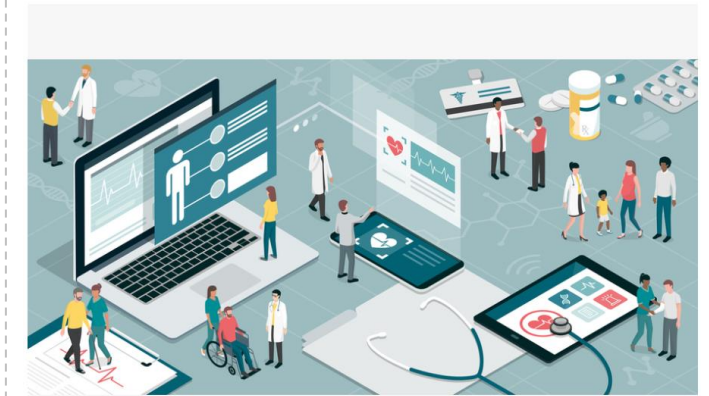
MIT News

ON CAMPUS AND AROUND THE WORLD

Large language models help decipher clinical notes

Researchers used a powerful deep-learning model to extract important data from electronic health records that could assist with personalized medicine.

Rachel Gordon | MIT CSAIL
December 1, 2022

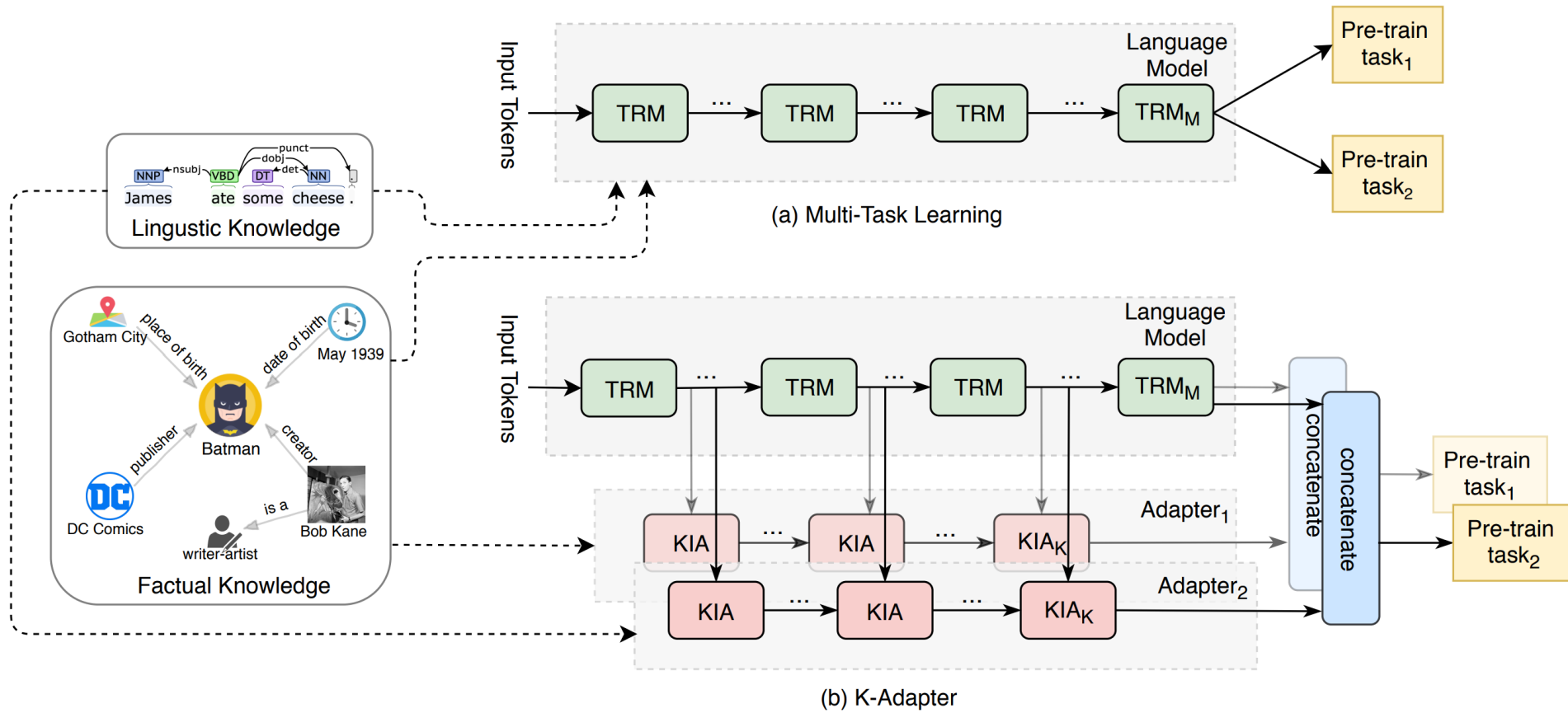


[Mi23] Microsoft: *BioGPT: generative pre-trained transformer for biomedical text generation and mining*

[Go22] Gordon, R.: *Large language models help decipher clinical notes*

[Ay23] Ayers, J., Poliak, A., Dredze, M., et al.: *Comparing Physician and Artificial Intelligence Chatbot Responses to Patient Questions Posted to a Public Social Media Forum*

Background: What are KELMs?

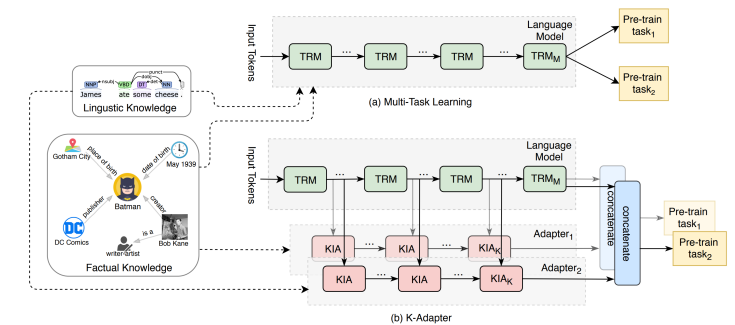


[Wu21a] Wang, R., Tang, D., Duan, N., Wei, Z., Huang, X., Ji, J., Cao, G., Jiang, D., Zhou, M.: K-ADAPTER: Infusing Knowledge into Pre-Trained Models with Adapters

Background: What are KELMs?

Medical natural language inference task (NLI):

- **Patient Premise:** No history of blood clots or DVTs, has never had chest pain prior to one week ago
- **Hypothesis:** Patient has angina

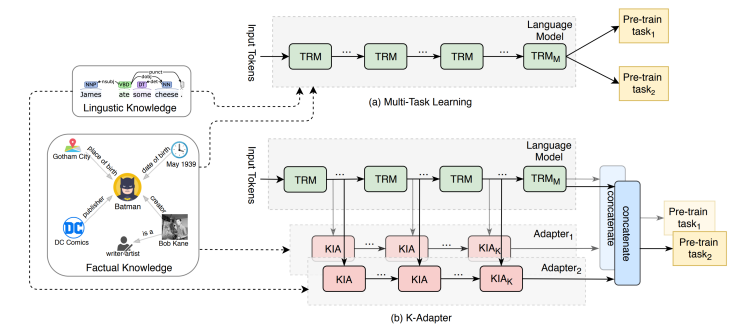


[Lu21] Lu, Q., Dou, D., Nguyen, T.H.: Parameter-Efficient Domain Knowledge Integration from Multiple Sources for Biomedical Pre-trained Language Models

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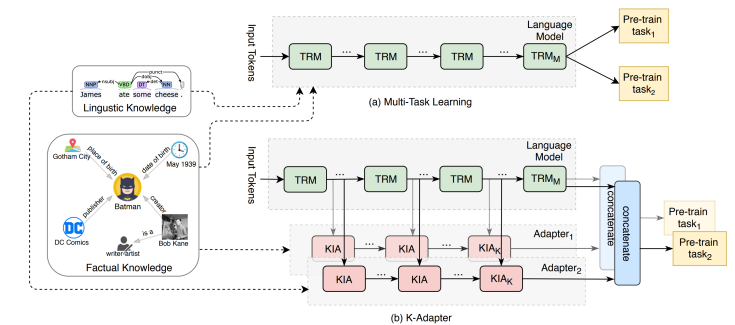
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➔ Correct inference more likely if model specifically learned synonyms and relations



Background: What are KELMs?

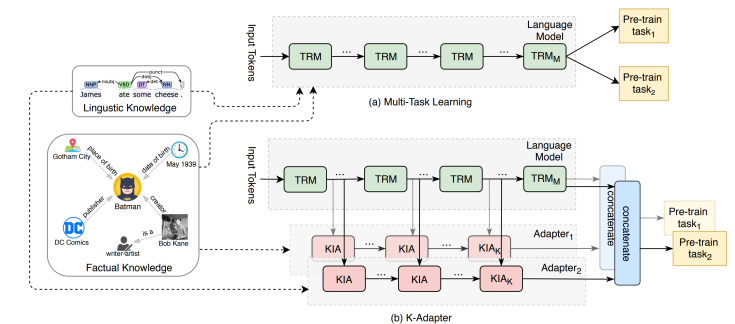
Relevance

- Active research area
- Can address hallucinations
- Superior performance over vanilla LMs

Shortcomings

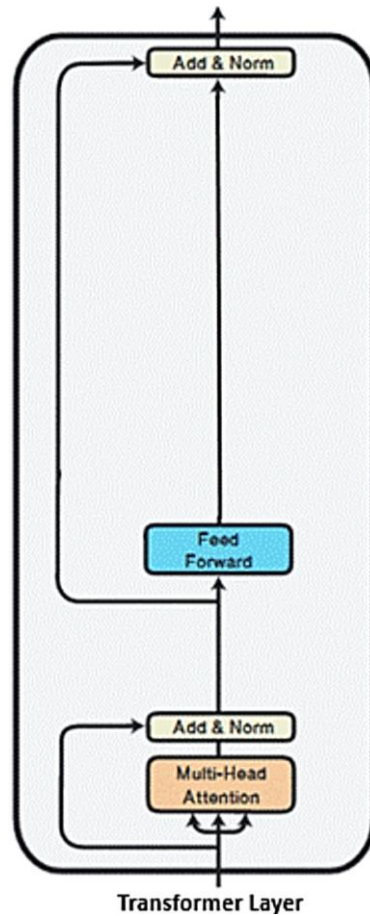
- Long Training Periods
- Catastrophic forgetting

➔ Lightweight “Adapters”



[We21a] Wei, X., Wang, S., Zhang, D., Bhatia, P., Arnold A.: Knowledge Enhanced Pretrained Language Models: A Comprehensive Survey
[Wa21a] Wang, R., Tang, D., Duan, N., Wei, Z., Huang, X., Ji, J., Cao, G., Jiang, D., Zhou, M.: K-ADAPTER: Infusing Knowledge into Pre-Trained Models with Adapters

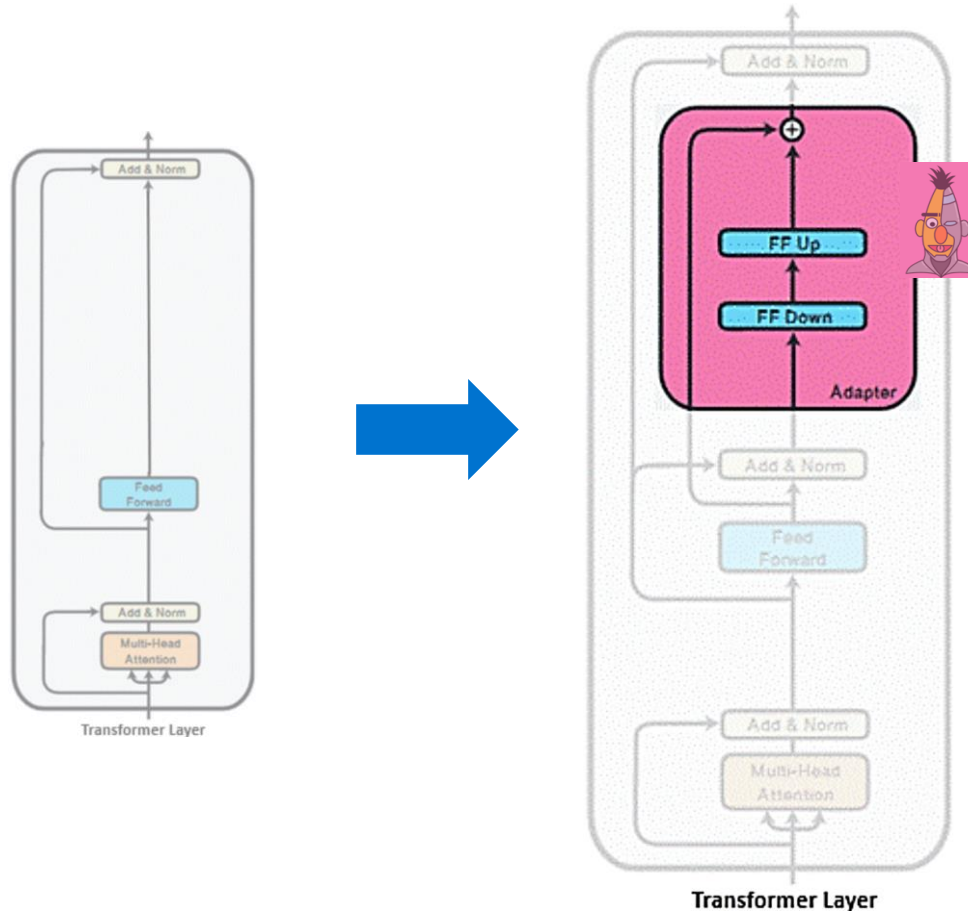
Background: What is an Adapter?



[Va17] Vaswani, A., Shazeer, N., Parmar, N., Uszkoreit, J., et al.: Attention Is All You Need

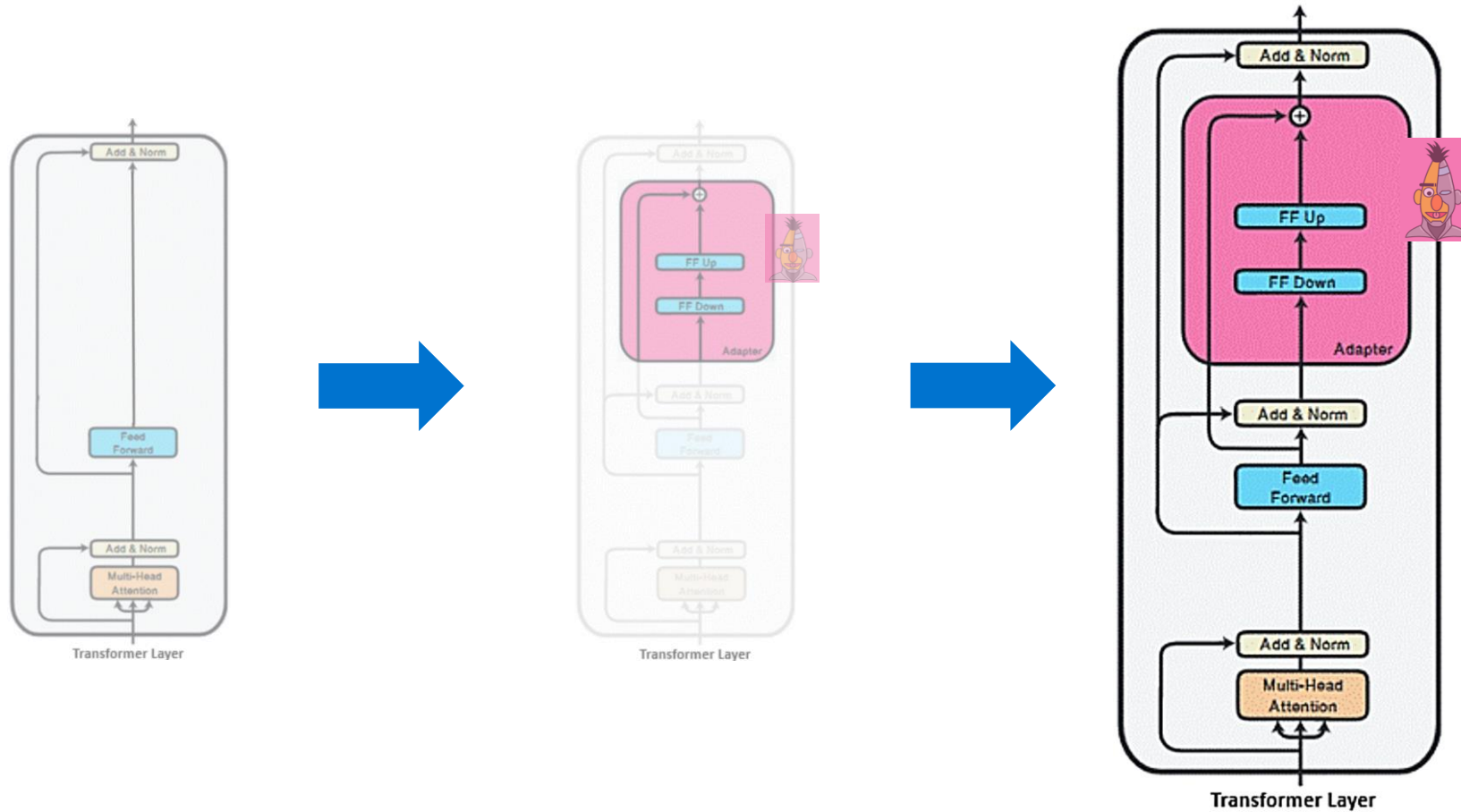
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Dataset	Task	Train	Dev	Test	EvaluationMetrics
HoC	DocumentClassification	1295	186	371	MicroF1
PubMedQA	QuestionAnswering	450	50	500	Accuracy
BioASQ	QuestionAnswering	670	75	140	Accuracy
MedNLI	Natural Language Inference	11232	1395	1,422	Accuracy



BLURB

Biomedical Language Understanding
and Reasoning Benchmark



RQ1: What adapter-based approaches to knowledge-enhancement exist, and how do they compare to each other?

- Systematic Literature Review (SLR)

RQ2: Can we improve existing approaches with new methods and data from a private ontology?

- SLR
- Thesis experiments with OntoChem KGs

RQ3: Is the research on biomedical KELMs relevant to medical professionals, and what factors hinder or support the deployment of the technology in practice?

- Research survey



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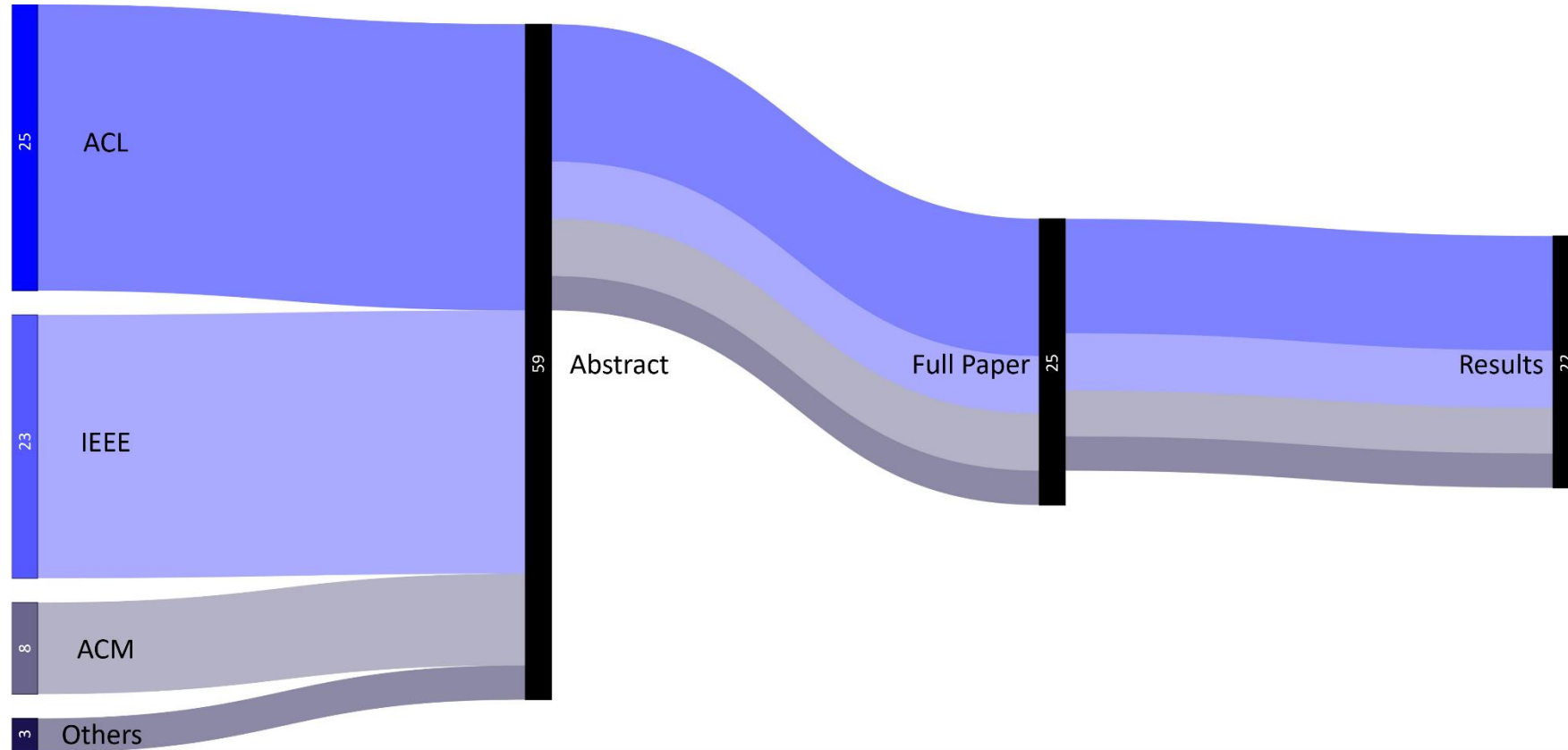
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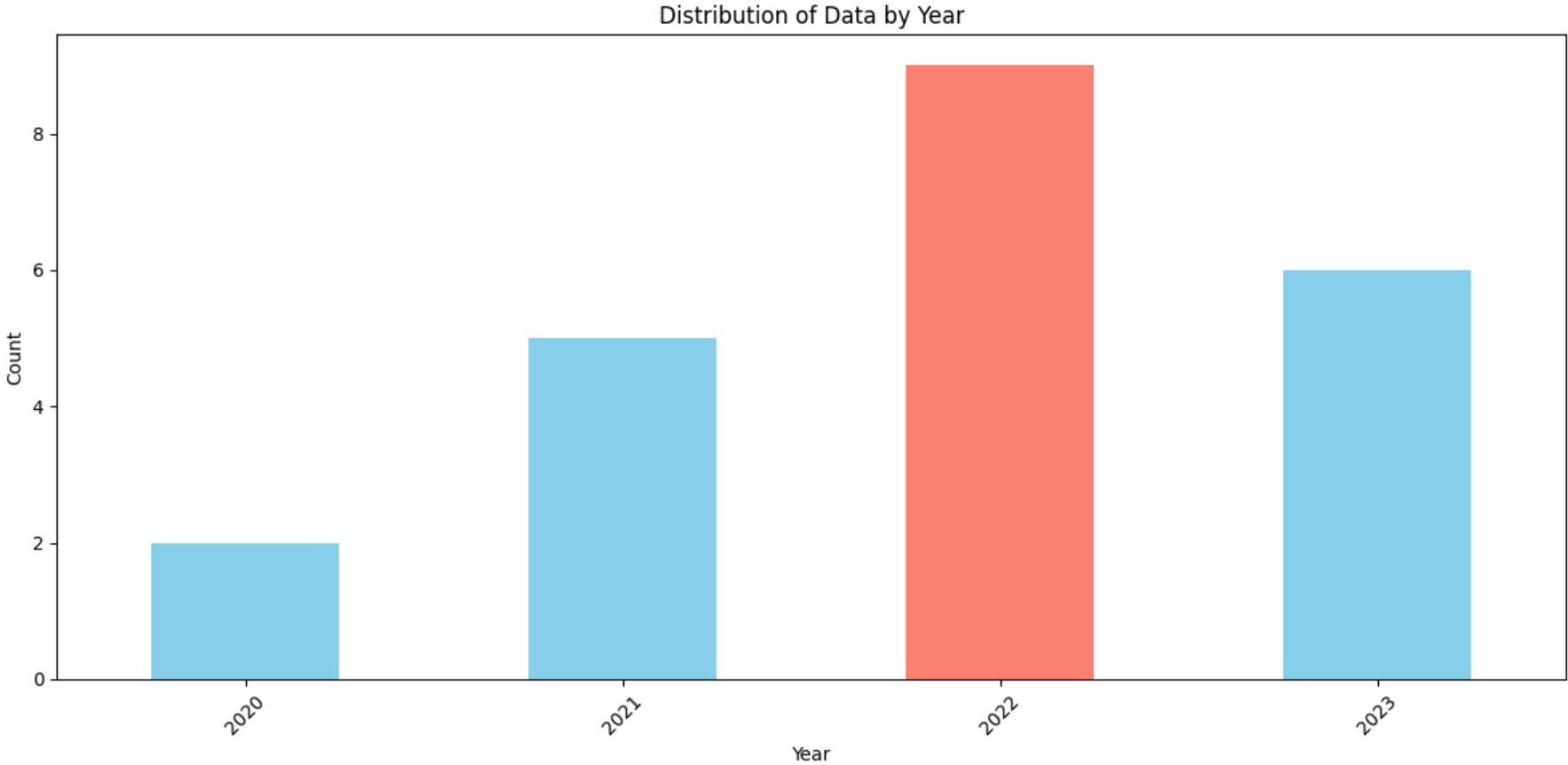
- SLR
- Thesis experiments

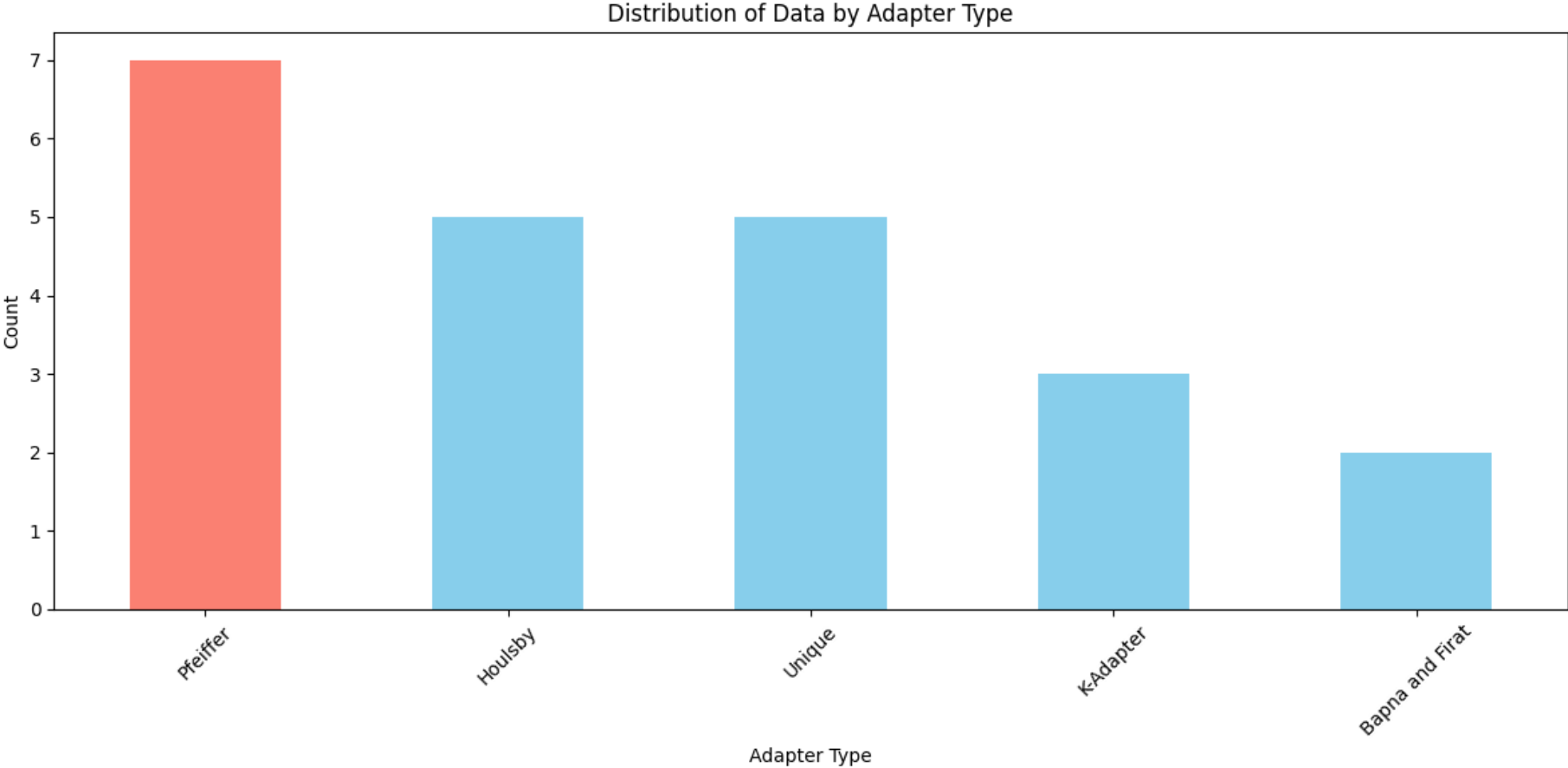
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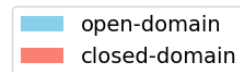
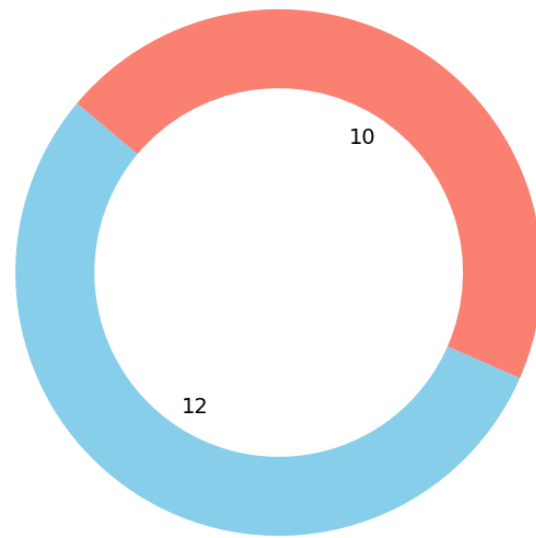
Criteria	Inclusion Criteria	Exclusion Criteria
Platforms	Papers published on ACM, ACL , or IEEE Xplore	Papers from other platforms (if not essential)
Search String	Papers matching the search string ("adapter" OR "adapter-based") AND ("language model" OR "nlp" OR "natural language processing") AND ("injection" OR "knowledge")	Papers that do not match the search string
Thematic Relation	Papers which address the topic of adapter-based knowledge-enhanced language models	Papers where Adapters were used for NLP, but for use-cases other than knowledge-enhancement
Language	Papers written in English or in German are included.	Papers not written English or German
Publication Date	Papers published after February 2, 2019 (publication of the Houlby Adapter, the first LLM adapter)	Papers published before February 2, 2019
Duplicates	Publications not yet part of the selection process	Duplicate versions of the same Papers



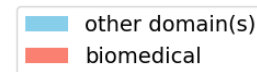
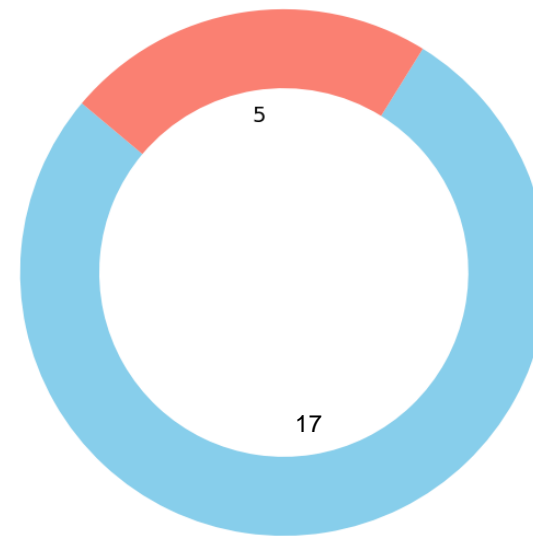




Domain Scope Distribution



Biomedical Domain Distribution



paper	adapter type	scope	coverage	biomed	task	nickname
Zou et al. (2022)	K-Adapter	open	/	no	RC	K-MBAN
Moon et al. (2021)	Houlsby	open	/	no	MT	/
Yu et al. (2023)	Unique	open	/	no	SL	CSBERT
Quian et al. (2022)	Unique	open	/	no	SR	/
Li et al. (2023)	Houlsby	closed	multi	no	SF	/
Li et al. (2023)	Unique	open	/	no	SC	CKGA
Nguyen et al.(2023)	Pfeiffer	open	/	no	SA	/
Lai et al. (2023)	Pfeiffer	closed	single	yes	QA, NLI, EL	KEBLM
Guo et al. (2022)	Unique	open	/	no	NER	/
Chronopoulou et al. (2023)	Bapna and Firat	closed	both	no	LM	Adapter-Soup
Wold et al. (2022)	Houlsby	open	/	no	LAMA	/
Chronopoulou et al. (2022)	Unique	closed	multi	no	LM	/
Hung et al. (2022)	Pfeiffer	closed	multi	no	TOD	DS-TOD
Emelin et al. (2022)	Houlsby	closed	multi	no	TOD	/
Xu et al. (2022)	Bapna and Firat	open	/	no	KGD	KnowBERT
Kær et al. (2021)	Pfeiffer	closed	multi	yes	NER, STC	mDAPT
Lu et al. (2021)	K-Adapter	closed	single	yes	NLI	DAKI
Majewska et al. (2021)	Pfeiffer	open	/	no	EE	/
Lauscher et al. (2020)	Houlsby	open	/	no	GLUE	/
Meng et al. (2021)	Pfeiffer	closed	single	yes	BLURB	MoP
Wang et al. (2020)	K-Adapter	open	/	no	RCL, ET, QA	K-Adapter
Xie et al. (2022)	Pfeiffer	closed	single	yes	ES	KeBioSum

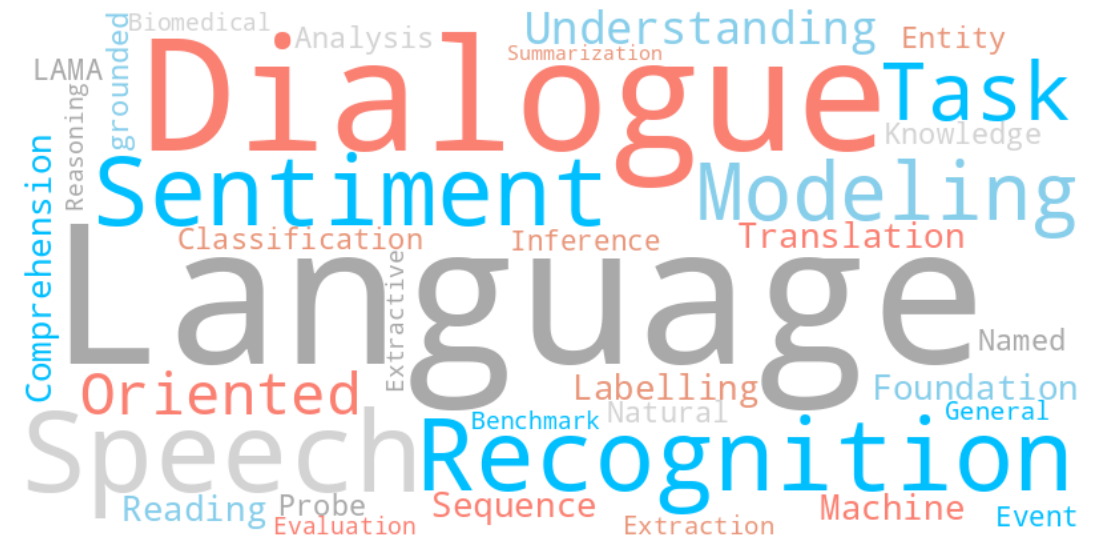
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RQ1: What adapter-based approaches to knowledge-enhancement exist, and how do they compare to each other?

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- The SLR portrayed, analysed, and compared existing approaches
- We found the best setting for our model experiments
- We provide a novel and extensive resource for other researchers in the field





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Fact Finder

Semantic knowledge extraction results in information organized into triples (e.g. "sumatriptan" "treats" "migraine headache"). Those triples come from millions of documents of various sources and can be queried individually or they can even be chained together in order to perform a shortest path analysis. [More](#)

Entity 1 [Edit](#)
Substances; Natural Pro...
aspirin
Ontological

183 triples
induces
Ontological

Entity 2 [Edit](#)
Diseases
All concepts from available domains
[Add entity](#)

Relations: Narrow [i](#) Direct matches: Show [Blocklists: Select blocklists](#) [Manage](#) [Clear all](#) [Save search](#) [Search](#)

[Shortest path](#) **[Full path](#)** [Triple](#) [Sub path](#)

183 matching paths

[Export](#)

<input type="checkbox"/>	#	Entity 1	Relation 1	Entity 2	Occurrences
<input type="checkbox"/>	1	acetylsalicylic acid	induces	Ulcer	129
<input type="checkbox"/>	2	acetylsalicylic acid	induces	Urticaria	72
<input type="checkbox"/>	3	acetylsalicylic acid	induces	Hemorrhage	68

Model Experiments: OntoChem Knowledge Graphs

Onto20Rel	#Triplets
relates to	708,076
induces	502,512
modulates	326,534
treats	225,279
inhibits	219,720
is analyzed by	195,291
produces	173,979
increases activity of	148,673
contains	133,241
increases	110,803
detects	93,373
decreases activity of	85,425
prevents	82,574
increases expression of	80,771
expresses	62,142
attenuates	54,865
decreases expression of	51,152
binds to	49,206
is a	47,435
affects expression of	37,399
Total	3,388,450

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Model Experiments: OntoChem Knowledge Graphs

Onto20Rel	#Triplets	OntoType20Rel	#Triplets
relates to	708,076	[protein] relates to [disease]	295,841
induces	502,512	[substance] induces [physiology]	282,721
modulates	326,534	[food] contains [compound]	269,211
treats	225,279	[substance] treats [disease]	247,348
inhibits	219,720	[biomarker] of [disease]	205,604
is analyzed by	195,291	[substance] is analyzed by [method]	130,275
produces	173,979	[plant] produces [compound]	102,270
increases activity of	148,673	[protein] induces [physiology]	85,411
contains	133,241	[compound] increases activity of [protein]	85,196
increases	110,803	[compound] decreases activity of [protein]	72,311
detects	93,373	[substance] inhibits [physiology]	68,728
decreases activity of	85,425	[protein] is a [biomarker]	65,558
prevents	82,574	[anatomy] produces [protein]	64,206
increases expression of	80,771	[substance] prevents [disease]	60,260
expresses	62,142	[protein] induces [disease]	59,577
attenuates	54,865	[substance] modulates [protein]	54,533
decreases expression of	51,152	[protein] is analyzed by [method]	54,250
binds to	49,206	[method] treats [disease]	35,768
is a	47,435	[method] detects [physiology]	33,504
affects expression of	37,399	[protein] modulates [physiology]	24,332
Total	3,388,450		2,296,904

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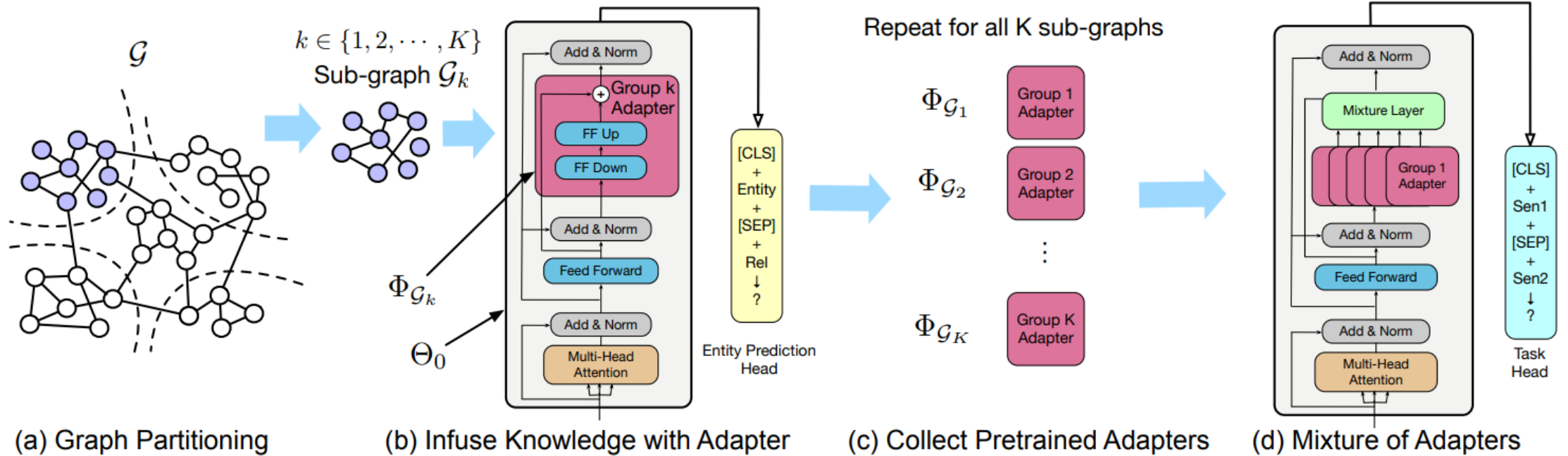
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[Me21] Meng, Z., Liu, F., Clark, T., et al.: Mixture-of-Partitions: Infusing Large Biomedical Knowledge Graphs into BERT
 [Pf21] Pfeiffer, J., Kamath, A., Rücklé, A., et al.: AdapterFusion: Non-Destructive Task Composition for Transfer Learning

Model Experiments: Results

↓ model dataset →	HoC	PubMedQA	BioASQ7b	MedNLI
SciBERT-base	80.52 \pm 0.60	57.38 \pm 4.22	75.93 \pm 4.20	81.19 \pm 0.54
+ MoP	81.79 [†] \pm 0.66 ↑	54.66 \pm 3.10	78.50 [†] \pm 4.06 ↑	81.20 \pm 0.37
+ KEBLM	/	59.0	/	82.14
BioBERT-base	81.41 \pm 0.59	60.24 \pm 2.32	77.50 \pm 2.92	82.42 \pm 0.59
+ MoP	82.53 [†] \pm 1.08 ↑	61.04 \pm 4.81 ↑	80.79 [†] \pm 4.40 ↑	82.93 \pm 0.55 ↑
+ KEBLM	/	68.00	/	84.24
+ DAKI	/	/	/	83.41
PubMedBERT-base	82.25 \pm 0.46	55.84 \pm 1.78	87.71 \pm 4.25	84.18 \pm 0.19
+ MoP	83.26 [†] \pm 0.32 ↑	62.84 [†] \pm 2.71 ↑	90.64 [†] \pm 2.43 ↑	84.70 \pm 0.19 ↑
+ OntoType20Rel (ours)	82.17 \pm 0.62	55.40 \pm 5.57	86.36 \pm 3.07	83.94 \pm 0.63
+ Onto20Rel (ours)	82.39 \pm 0.65 ↑	56.12 \pm 2.91 ↑	84.36 \pm 4.73	83.97 \pm 0.59
BioLinkBERT-base	82.21 \pm 0.87	56.76 \pm 3.00	91.29 \pm 3.18	84.1 \pm 0.03
+MoP (ours)	82.36 \pm 0.57 ↑	63.62 [†] \pm 5.31 ↑	91.50 \pm 2.25 ↑	83.78 \pm 0.09
+OntoType20Rel (ours)	82.37 \pm 0.42 ↑	60.46 \pm 5.81 ↑	92.14 \pm 2.30 ↑	82.84 \pm 0.34
+Onto20Rel (ours)	82.24 \pm 1.25 ↑	63.28 [†] \pm 4.46 ↑	90.57 \pm 3.14	83.69 \pm 0.55

[Me21] Meng, Z., Liu, F., Clark, T., et al.: Mixture-of-Partitions: Infusing Large Biomedical Knowledge Graphs into BERT

[Gu20] Gu, Y., Tinn, R., Cheng, H., et al.: Domain-Specific Language Model Pretraining for Biomedical Natural Language Processing

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+ Onto20Rel (ours)	82.39 \pm 0.65 \uparrow	56.12 \pm 2.91 \uparrow	84.36 \pm 4.73	83.97 \pm 0.59
BioLinkBERT-base	82.21 \pm 0.87	56.76 \pm 3.00	91.29 \pm 3.18	84.1 \pm 0.03
+MoP (ours)	82.36 \pm 0.57 \uparrow	63.62 \pm 5.31 \uparrow	91.50 \pm 2.25 \uparrow	83.78 \pm 0.09
+OntoType20Rel (ours)	82.37 \pm 0.42 \uparrow	60.46 \pm 5.81 \uparrow	92.14\pm2.30 \uparrow	82.84 \pm 0.34
+Onto20Rel (ours)	82.24 \pm 1.25 \uparrow	63.28 \pm 4.46 \uparrow	90.57 \pm 3.14	83.69 \pm 0.55

[Me21] Meng, Z., Liu, F., Clark, T., et al.: Mixture-of-Partitions: Infusing Large Biomedical Knowledge Graphs into BERT

[Gu20] Gu, Y., Tinn, R., Cheng, H., et al.: Domain-Specific Language Model Pretraining for Biomedical Natural Language Processing

Model Experiments: Results (Qualitative Probing)

Question: Can Diazepam be beneficial in the treatment of traumatic brain injury?

Context: The present experiment examined the effects of diazepam, a positive modulator at the GABA(A) receptor, on survival and cognitive performance in traumatically brain-injured animals.

Predictions:

BioLinkBERT:	no
BioLinkBERT	
+OntoType20Rel:	yes
True Label:	yes

Entity 1: Substances; Natural Pro... diazepam

Entity 2: Diseases; Proteins, Gen... brain injury

1 triple

All relations or choose

Add entity

1 matching path

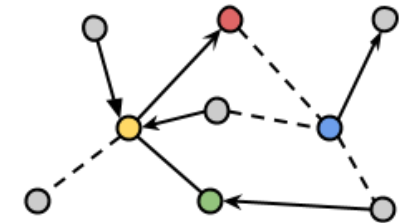
Choose number of hops: 1

Export

<input checked="" type="checkbox"/>	#	Entity 1	Relation 1	Entity 2	Occurrences
<input checked="" type="checkbox"/>	1	diazepam	treats	brain injury	1

RQ2: Can we improve existing approaches with new methods and data from a private ontology?

- We were able to improve KELMs with OntoChem KGs
- We proposed Onto20Rel and OntoType20Rel
- Diversification of KG Sources





RQ1: What adapter-based approaches to knowledge-enhancement exist, and how do they compare to each other?

- Systematic Literature Review (SLR)

RQ2: Can we improve existing approaches with new methods and data from a private ontology?

- SLR
- Thesis experiments with OntoChem KGs

RQ3: Is the research on biomedical KELMs relevant to medical professionals, and what factors hinder or support the deployment of the technology in practice?

- Research survey

- **Extent:** Twenty questions, including a brief introduction to biomedical NLP
- **Targeting:** Medical centres and hospitals in Munich and the surrounding countryside, student groups at TUM and LMU, personal contacts of the authors and advisor
- **Related work:** Survey focus on NLP in biomedicine not yet represented in literature. Comparison with survey on AI in general

Survey on the Use of Language Models in Medicine and Biomedical Research

In recent years, there has been significant progress in the development of large language models (LLMs). In addition to well-known LLMs such as ChatGPT, there also exists an increasing number of specialized language models for use and research in medicine and biomedicine. Our research at the Technical University of Munich deals with the applications and relevance of these specialized models. Your insights help us assess the impact and importance of ongoing research.

This survey consists of 20 short questions and takes a maximum of 10 minutes to complete.

If you are unfamiliar with LLMs and their applications in the (bio)medical field, please read the brief [introduction](#) provided for you and then return to the survey.

If you have any questions or comments regarding the survey, please email us at: alexander.fichtl@tum.de

We appreciate your help and thank you for your time!

*Indicates required question

Informed Consent

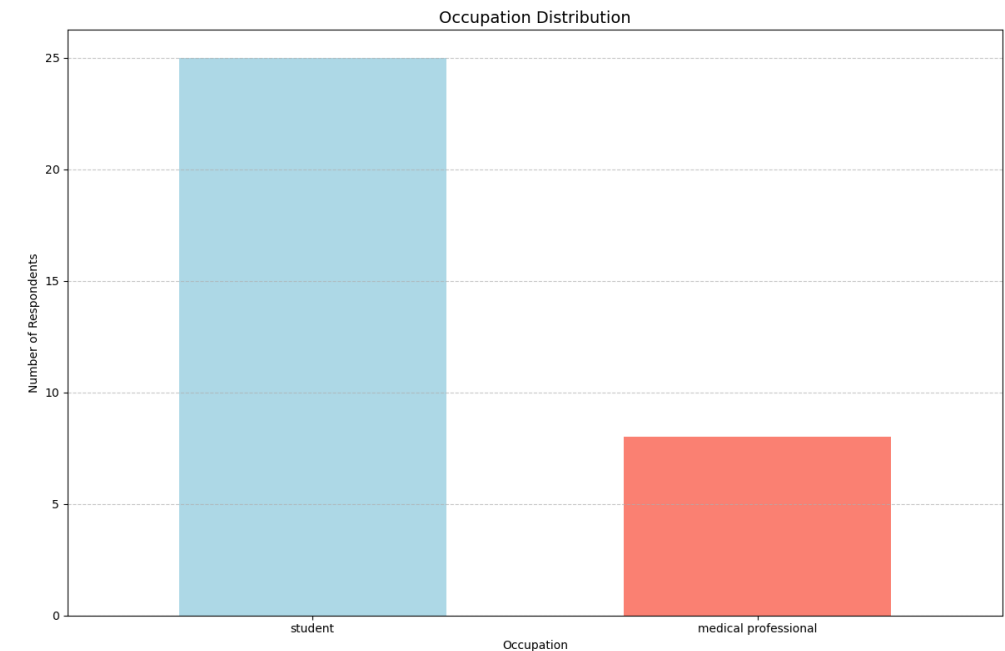
All information given on this survey will be completely anonymous. All data is used exclusively for research purposes at the Technical University of Munich. If you wish to have your data deleted after you have already submitted the survey, please contact: alexander.fichtl@tum.de.

1. Do you wish to participate? *

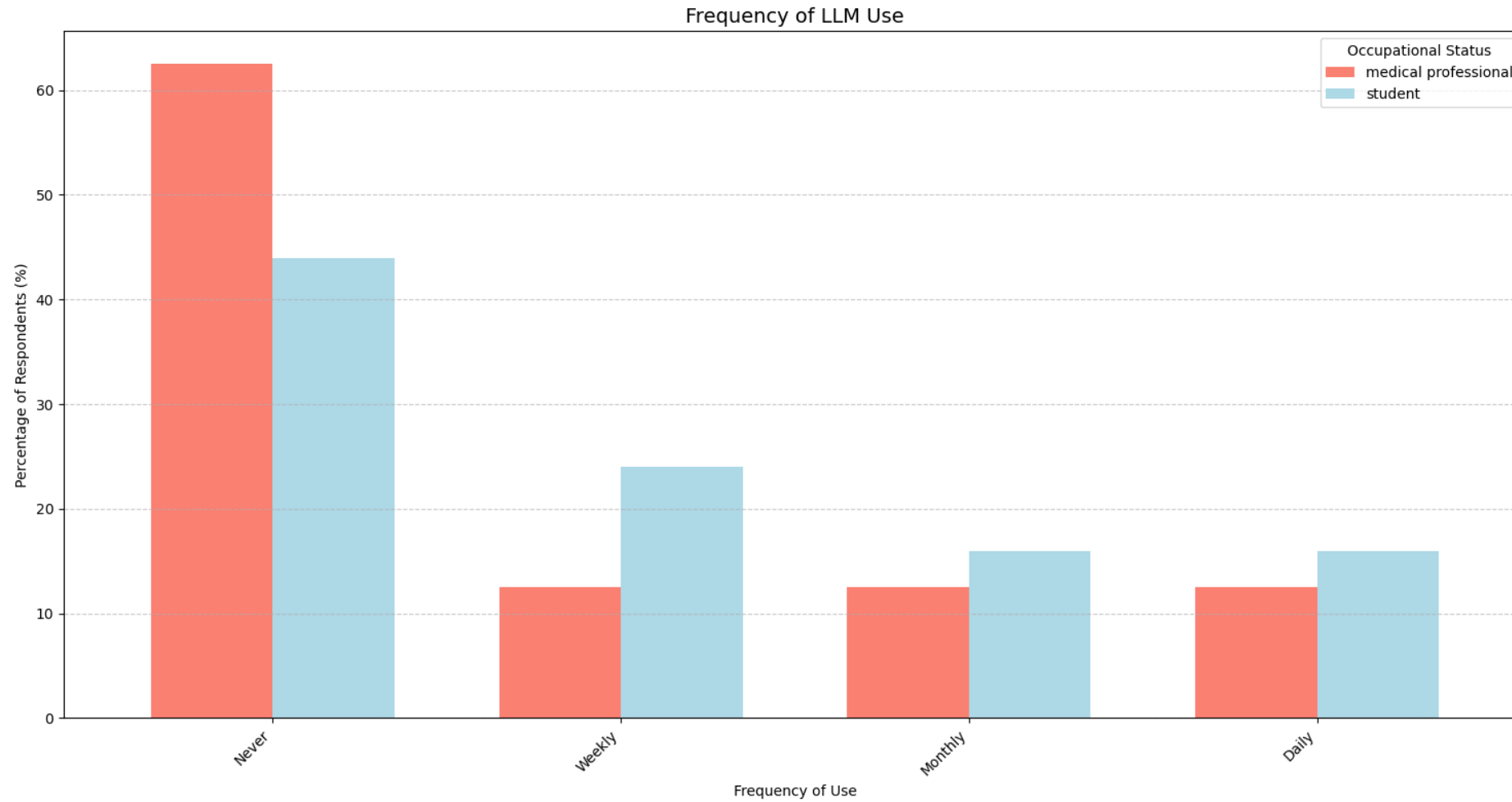
Mark only one oval.

- Yes *Skip to question 2*
- No *Skip to section 2 (Declined Participation)*

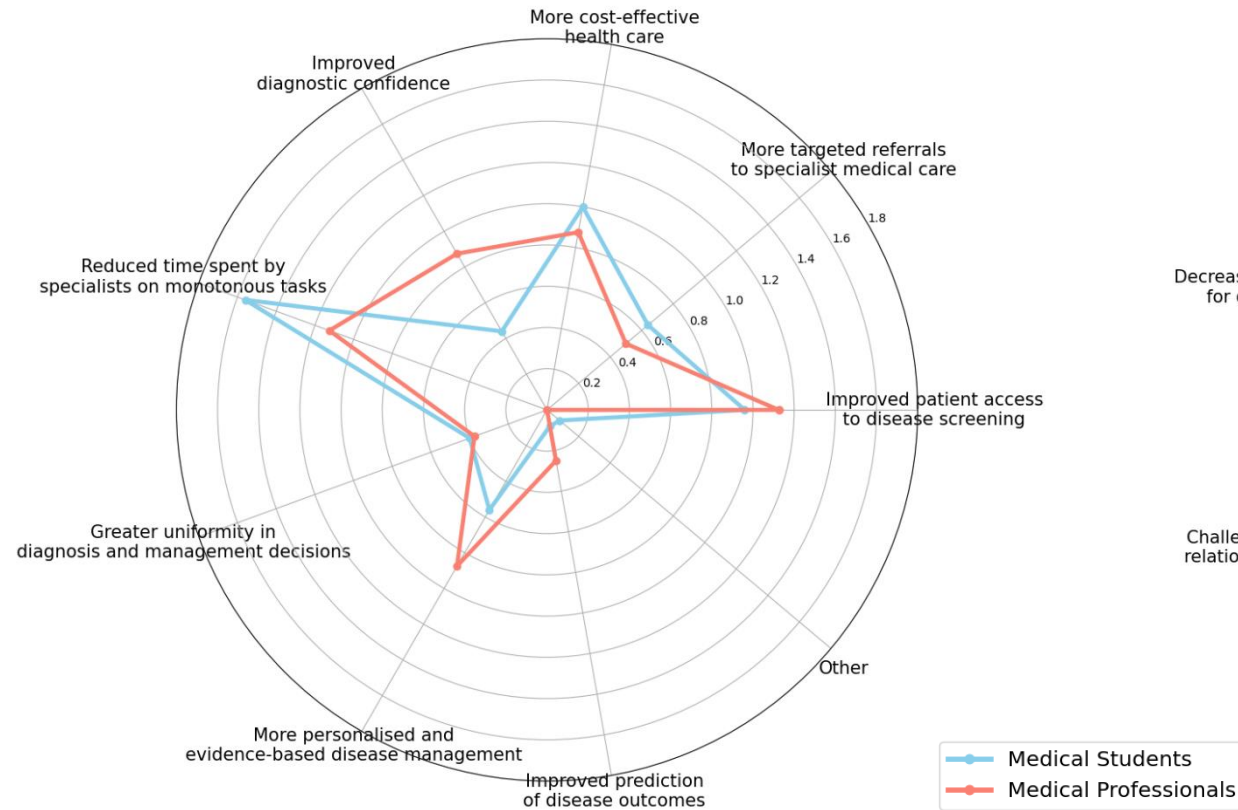
- **Participants:** 34 participants, with 33 usable questionnaires
- **Distribution:** 25 students, 8 medical professionals
 - Experience levels from <5 to >30 years
 - Specialties vary strongly across medical disciplines
- Preliminary survey in preparation for survey at TUM Klinikum rechts der Isar (MRI)



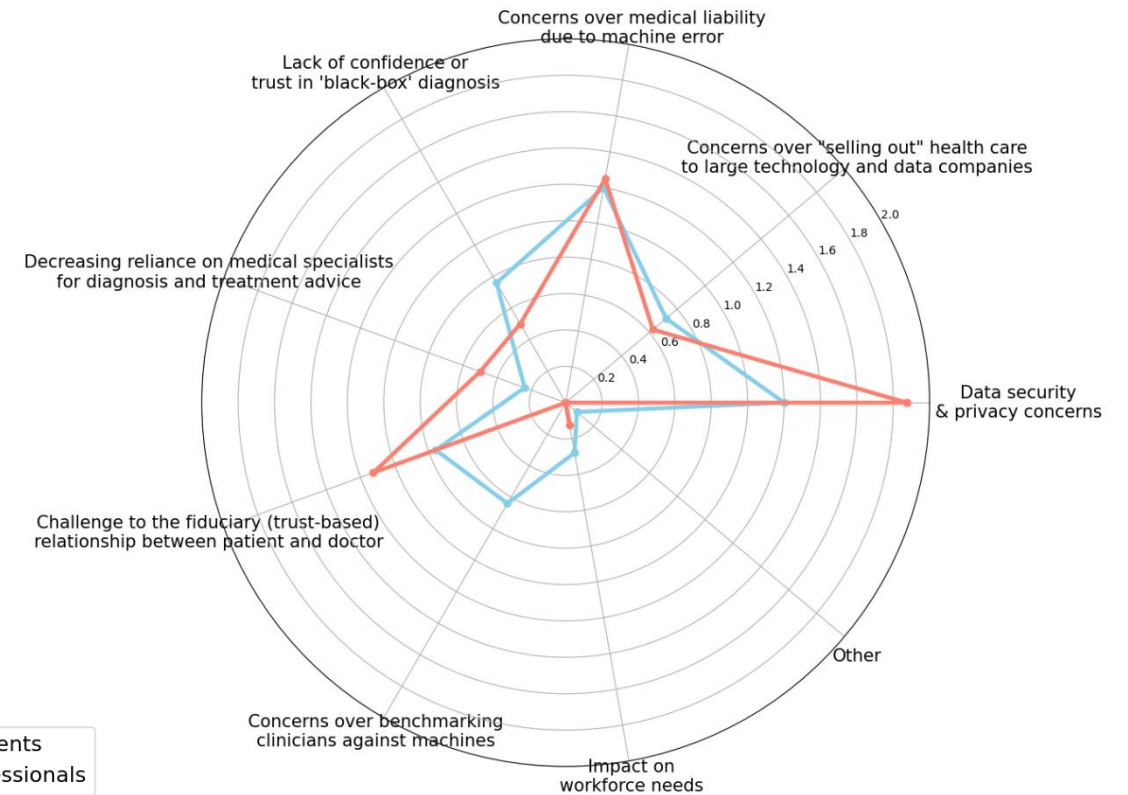
Research Survey: Results



Chances



Concerns



“For me, teaching and training needs to be more rigorous so that those using the technology can detect potential issues. I do think that a free chatbot replacing an expensive doctor’s visit to tell you that you need bed rest and plenty of water is viable, but I think that it will take a decade to become mainstream.”

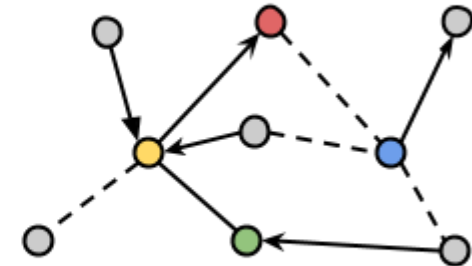
- Medical student

“More information about possible use cases should be provided in training. Also, experts should be hired who, in collaboration with doctors, can devise customized strategies or concepts tailored to individual needs.”

- Pathologist, 10-20 years of experience,
translated from German

RQ3: Is the research on biomedical KELMs relevant to medical professionals, and what factors hinder or support the deployment of the technology in practice?

- The medical community is gaining familiarity with NLP and LLMs
- Participants recognize and anticipate chances and advantages
- There are prevalent concerns, like data security and the impact on the doctor-patient trust relationship, but notably not regarding model performance

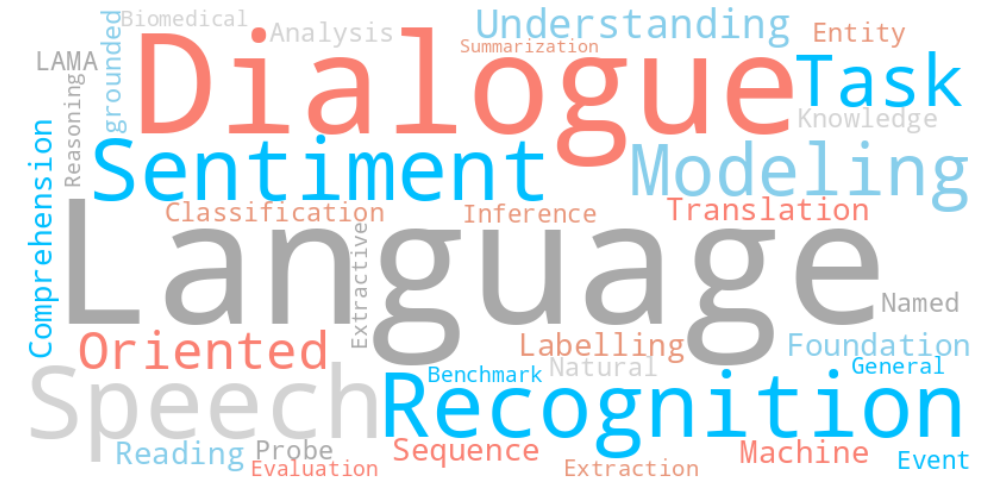


Main Shortcomings:

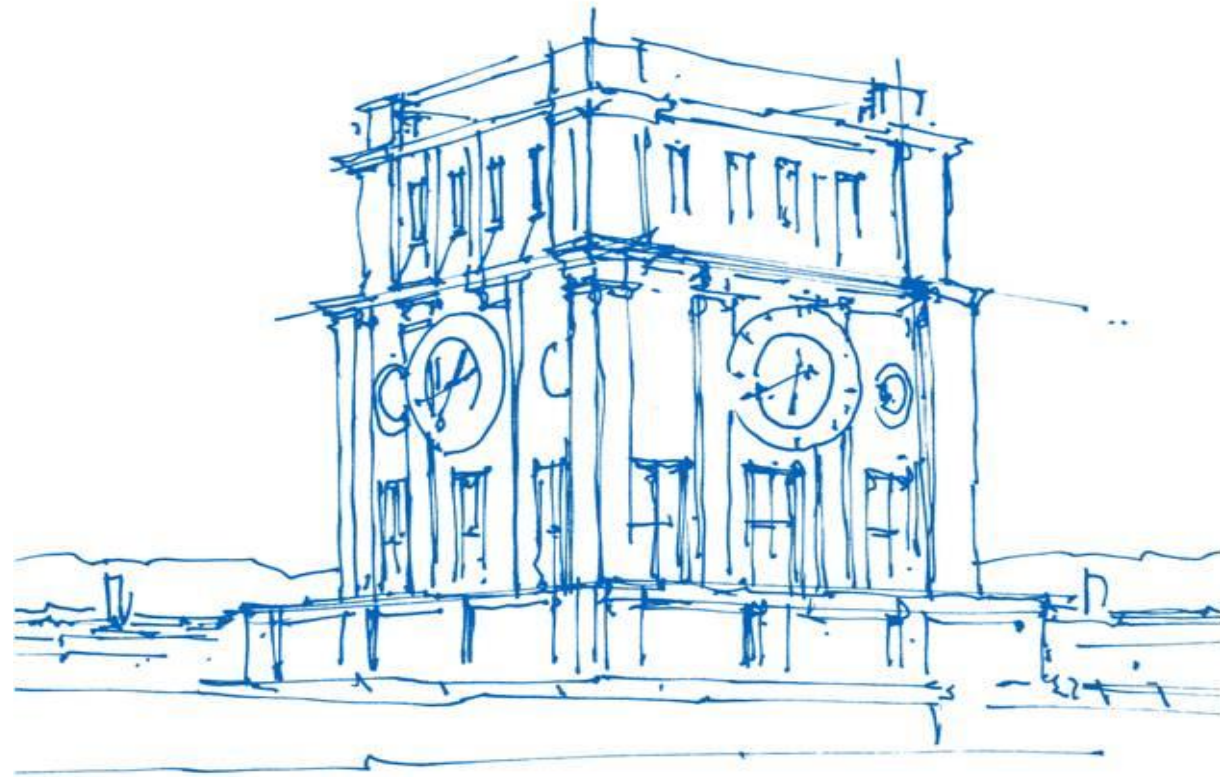
- Incomplete KGs
- Survey Response Rates

Future Research Opportunities:

- OntoChem Potential
- KG Merging
- ...



Thanks!



Uhrenturm der TUM



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Question:

Do preoperative statins reduce atrial fibrillation after coronary artery bypass grafting?

Context:

(Objective) Recent studies have demonstrated that statins have pleiotropic effects, including anti-inflammatory effects and atrial fibrillation (AF) preventive effects [...]

(Methods) 221 patients underwent CABG in our hospital from 2004 to 2007. 14 patients with preoperative AF and 4 patients with concomitant valve surgery [...]

(Results) The overall incidence of postoperative AF was 26%. *Postoperative AF was significantly lower in the Statin group compared with the Non-statin group (16% versus 33%, $p=0.005$).* Multivariate analysis demonstrated that independent predictors of AF [...]

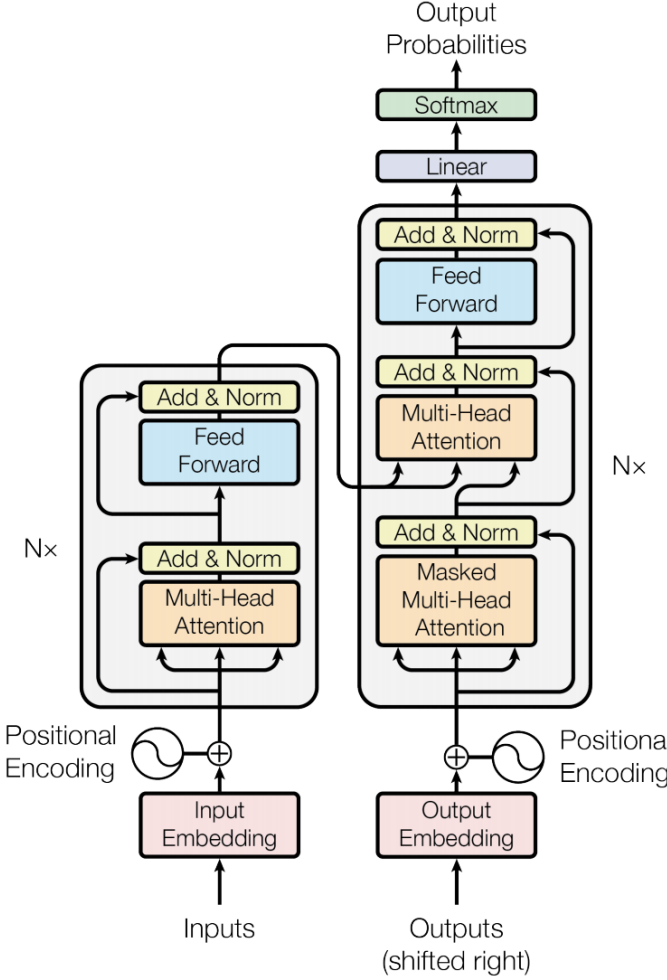
Long Answer:

(Conclusion) Our study indicated that preoperative statin therapy seems to reduce AF development after CABG.

Answer: yes

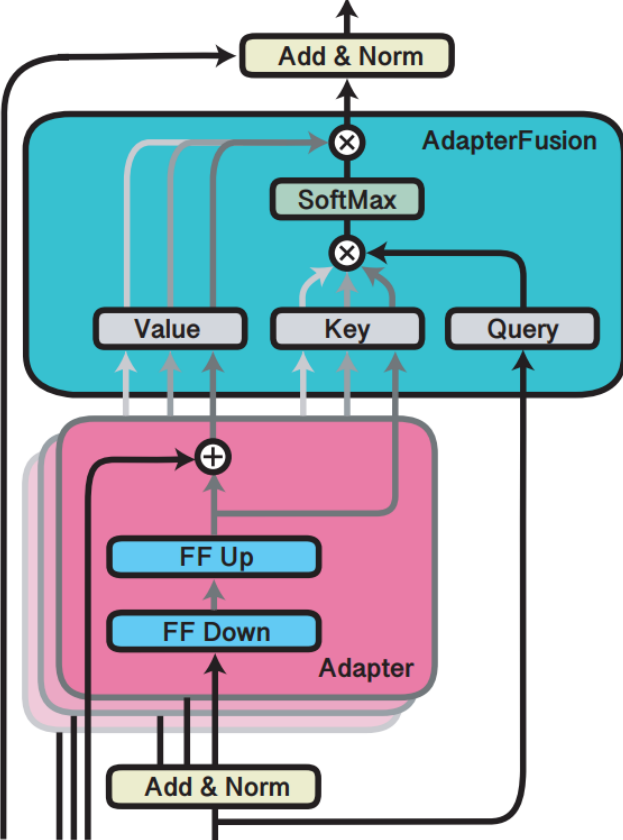
Figure 1: An instance (Sakamoto et al., 2011) of PubMedQA dataset: Question is the original question title; Context includes the structured abstract except its conclusive part, which serves as the Long Answer; Human experts annotated the Answer yes. Supporting fact for the answer is *highlighted*.

Appendix B: Transformers



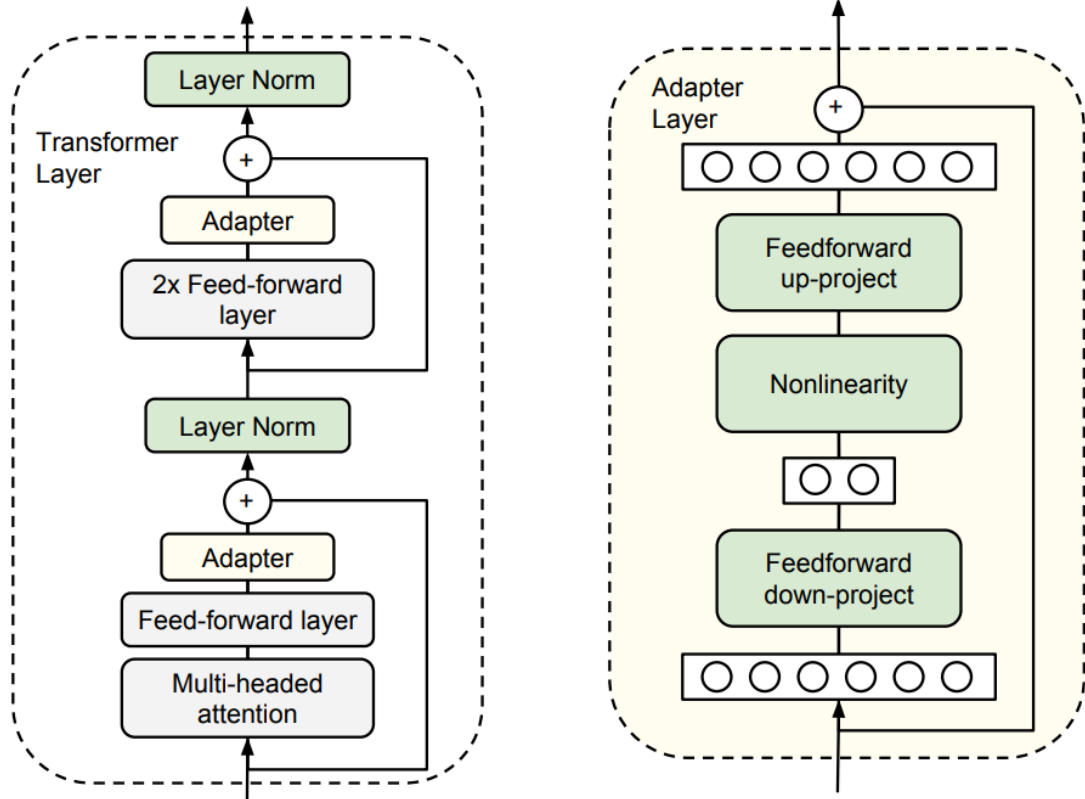
[Va17] Vaswani, A., Shazeer, N., Parmar, N., Uszkoreit, J., et al.: Attention Is All You Need

Appendix C: Adapter Fusion



[Pf21] Pfeiffer, J., Kamath, A., Rücklé, A., et al.: AdapterFusion: Non-Destructive Task Composition for Transfer Learning

Appendix D: Houlsby Adapter



[Pf21] Houlsby, N., Giurgiu, A., Jastrzebski, S., et al.: Parameter-Efficient Transfer Learning for NLP:



Can adapter-based approaches outperform other knowledge injection methods in downstream tasks (Blurb, claim verification)?

- Literature review
- Thesis experiments

How does the performance of KELMs in closed domains compare to open domain performance?

- Literature review
- Thesis experiments

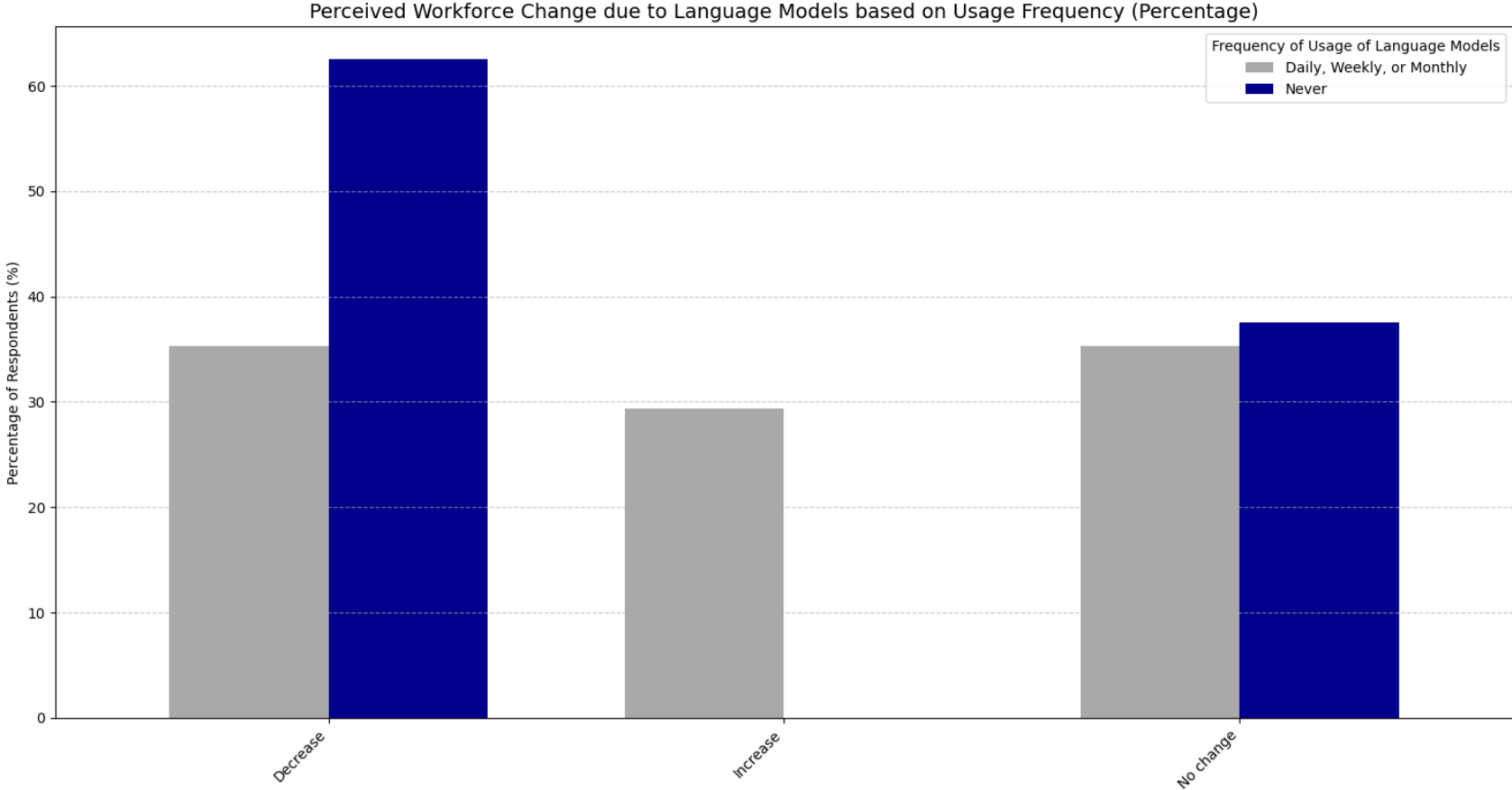
How do models trained on a private ontology (e.g., OntoChem) compare to models trained on public ontologies (e.g., UMLS)?

- Thesis experiments
- SciWalker

Is there interest in the results of this thesis amongst medical professionals and can they make use of biomedical KELMs?

- Survey and Interviews
- Mini-workshops on knowledge enhancement with adapters

Appendix F: Perceived Implication



Appendix G: Survey Participant Background

1. What is your (aspired) specialty?	2. Years of practice in your field of expertise?
Psychiatry	Currently in training/Student
Genetics	Currently in training/Student
Microbiology	Currently in training/Student
Data Scientist (Medical)	<5 years
August Staimann	Currently in training/Student
biostatistics	<5 years
Optometry	Currently in training/Student
training/Student	Currently in training/Student
medical technology	Currently in training/Student
Undecided	Currently in training/Student
Plastic surgery	Currently in training/Student
designing	<5 years
Data science	5-10 years
Consultant medical doctor (obstetrics and gynaecology)	Currently in training/Student
Brain computer interfaces	Currently in training/Student
Innere	Currently in training/Student
Psychiatrie	Currently in training/Student
Chirurgie	Currently in training/Student
Experimentelle unfallchirurgische Forschung	<5 years
Innere Medizin (hausärztlich)	20-30 years
Psychiatrie und Psychotherapie	Currently in training/Student
Innere Medizin	Currently in training/Student
Allgemeinmedizin	Currently in training/Student
Innere	Currently in training/Student
Pädiatrie	Currently in training/Student
Physiologie Forschung	>30 years
Öffentliche Apotheke	<5 years
Notfallpflege	>30 years
Orthopädie	Currently in training/Student
Chirurgin	Currently in training/Student
TBD	Currently in training/Student
Pathologie	10-20 years