

When Chain-of-Thought Backfires: Evaluating Prompt Sensitivity in Medical Language Models

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Abstract

Large language models are increasingly deployed in medical settings, yet their sensitivity to prompt formatting remains poorly characterized. We evaluate MedGemma (4B and 27B variants), Google’s medical-specialist language model, on two benchmark datasets: MedMCQA (4,183 questions) and Pub-MedQA (1,000 questions). Our experiments reveal several concerning findings. Chain-of-thought prompting decreases accuracy by 5.7% compared to direct answering, contradicting the assumption that reasoning traces improve performance. Few-shot examples degrade performance by 11.9%, with position bias increasing from 0.14 to 0.47. Shuffling answer options causes the model to change its prediction 59.1% of the time, with accuracy dropping by up to 27.4 percentage points. Truncating context to 50% causes accuracy to plummet from 45.0% to 14.1% (4B) and from 38.2% to 23.4% (27B)—worse than providing no context at all. Surprisingly, MedGemma-27B performs best when given only study results (40.0%), exceeding its full-context accuracy (38.2%), suggesting larger models may benefit from selective rather than comprehensive context. These results demonstrate that prompt engineering techniques validated on general-purpose models do not transfer to domain-specific medical LLMs, and that deployment requires rigorous empirical validation rather than assumed best practices.

1 Introduction

Medical question answering is hard. Unlike general knowledge tasks where multiple reasonable answers might exist, clinical decisions often hinge on precise distinctions between similar-sounding options. A model that performs well on average may still fail catastrophically on the specific cases where accuracy matters most.

Large language models have shown impressive performance on medical licensing exams [Singhal et al., 2023a, Nori et al., 2023]. This has fueled enthusiasm for deploying LLMs in clinical decision support. But benchmark accuracy tells only part of the story. How these models respond to variations in prompt format, question ordering, and context presentation remains unclear.

We focus on MedGemma [DeepMind, 2024], Google’s medical-specialist LLM built on the Gemma architecture. The model was specifically trained on medical literature and clinical data, making it a natural candidate for healthcare applications. Our goal is not to benchmark raw accuracy, but to stress-test the model’s robustness to common prompt engineering variations.

1.1 The Prompt Engineering Assumption

A widely held belief in the LLM community is that certain prompting strategies reliably improve performance. Chain-of-thought prompting, where models are instructed to reason step-by-step before answering, has shown gains across mathematical and reasoning tasks [Wei et al., 2022]. Few-shot learning, where examples are provided in-context, helps models understand desired output formats [Brown et al., 2020]. These techniques are often treated as “best practices” that should transfer across domains.

But should they? Domain-specific models may have learned different response patterns during training. A model trained extensively on medical text might already internalize structured reasoning, making explicit chain-of-thought prompts redundant or even harmful. Similarly, few-shot examples from one medical specialty might mislead the model when applied to another.

1.2 Contributions

We present a systematic evaluation of MedGemma’s sensitivity to prompt variations across three primary experimental conditions. First, we conduct a prompt ablation study comparing zero-shot, chain-of-thought, and few-shot prompting strategies on the full MedMCQA validation set of 4,183 questions, measuring not only accuracy but also position bias under each condition. Second, we test option order sensitivity by shuffling answer choices and measuring how often the model changes its prediction when the same question is presented with reordered options—a direct test of whether the model is responding to semantic content or superficial position cues. Third, we evaluate evidence conditioning on 1,000 PubMedQA questions, systematically varying the amount and type of context provided to understand how incomplete or section-specific information affects model accuracy.

Our findings challenge several conventional assumptions about prompt engineering in medical AI, and we believe they have important implications for anyone considering deployment of language models in clinical or biomedical settings. We release our evaluation framework and results to support further research on LLM robustness in high-stakes domains.

2 Related Work

2.1 Medical Language Models

The application of large language models to medicine has accelerated rapidly. GPT-4 achieved accuracy rates of 93.2%, 95.0%, and 92.0% on USMLE Steps 1, 2CK, and 3 respectively, exceeding the passing threshold by over 20 points [Nori et al., 2023]. Med-PaLM was the first AI system to surpass the 60% passing mark on USMLE-style questions, and Med-PaLM 2 subsequently achieved 86.5% on MedQA [Singhal et al., 2023a,b]. These results have fueled enthusiasm for deploying LLMs in clinical decision support.

MedGemma, introduced at Google I/O 2025, represents the latest generation of medical-specialist models [DeepMind, 2024]. Built on the Gemma architecture and trained on medical literature and clinical data, MedGemma-27B achieves 87.7% on MedQA, within 3 points of larger models like DeepSeek R1 but at approximately one-tenth the inference cost. However, Google emphasizes that MedGemma is not intended for direct clinical use without further validation—a caveat our results strongly support.

Despite impressive benchmark numbers, concerns persist about real-world reliability. Benchmark performance may not capture the model’s behavior under realistic deployment conditions where prompts vary, context is incomplete, and question formatting differs from training distributions.

2.2 Prompt Sensitivity and Robustness

The fragility of LLM predictions to prompt variations is well-documented in general domains. Lu et al. [2022] demonstrated that few-shot example ordering significantly affects performance. The ProSA framework introduced PromptSensiScore to quantify this sensitivity, finding that performance can swing by up to 45% depending on prompt formulation [Jia et al., 2024]. Larger models generally exhibit enhanced robustness, but even state-of-the-art systems remain vulnerable.

2.3 Position Bias in Multiple-Choice Questions

Zheng et al. [2024] showed that modern LLMs are vulnerable to option position changes due to inherent “selection bias”—they prefer specific option IDs (like “Option A”) regardless of content. In their analysis of 20 LLMs across three benchmarks, llama-30B selected options A/B/C/D with frequencies of 34.6%/27.3%/22.3%/15.8% respectively, despite balanced ground truth distributions. This bias stems from token-level preferences where models assign more probabilistic mass to certain option ID tokens. Their proposed PriDe debiasing method separates prior bias from predictions, but requires additional inference overhead.

2.4 Chain-of-Thought: When Reasoning Hurts

Chain-of-thought (CoT) prompting has become a standard technique for improving LLM reasoning [Wei et al., 2022]. However, recent work challenges its universal benefit. Sprague et al. [2024] identified tasks where CoT reduces performance by up to 36.3% absolute accuracy, drawing parallels to cognitive psychology research on when deliberation hurts human performance. The Wharton “Decreasing Value of CoT” report found that while CoT generally provides small average gains for non-reasoning models, it introduces more variability and can trigger errors on questions the model would otherwise answer correctly [Meincke et al., 2024]. For dedicated reasoning models, explicit CoT prompting appears to provide negligible additional benefit while substantially increasing processing time.

In medical domains specifically, Omar et al. [2024] found that complex prompting techniques do not significantly enhance performance compared to simpler approaches, suggesting that dataset characteristics and model architecture have greater impact than prompt engineering.

2.5 Context and Retrieval-Augmented Generation

Retrieval-augmented generation (RAG) systems face particular challenges with incomplete or misleading context. Barnett et al. [2024] identified seven recurrent failure points in operational RAG systems, including retrieval errors, context consolidation failures, and incomplete answers. The “lost-in-the-middle” phenomenon shows that key information position within context significantly impacts response quality [Liu et al., 2024]. Most relevant to our findings, RAG-Bench demonstrated that relevant-but-incomplete retrieved context can actively mislead models, sometimes performing worse than no retrieval at all [Fang et al., 2024]. Our evidence conditioning experiments provide direct evidence of this phenomenon in medical question answering.

3 Methods

3.1 Models

We evaluate two variants of MedGemma, Google’s medical-specialist language model family. The primary model in our experiments is MedGemma-4B, the 4-billion parameter instruction-tuned variant, which we run at bfloat16 precision on standard GPU hardware. This model size allows for rapid iteration across our experimental conditions while still representing a capable medical language model.

For our evidence conditioning experiments, we additionally evaluate MedGemma-27B, the larger 27-billion parameter model, to assess whether increased scale improves robustness to context variations. Running the 27B model presented unexpected challenges: our initial experiments with 4-bit quantization produced NaN logits and completely unusable outputs, a failure mode we did not observe with the 4B model. After extensive debugging, we determined that full bfloat16 precision was necessary for stable inference, which in turn required 80GB A100 GPUs. This quantization sensitivity is itself a notable finding, as it suggests that memory-reduction techniques validated on general-purpose models may not transfer reliably to medical-specialist architectures.

3.2 Datasets

MedMCQA A large-scale multiple-choice dataset derived from Indian medical entrance examinations [Pal et al., 2022]. The dataset covers 21 medical subjects with over 194,000 questions. We use the validation split containing 4,183 questions for our experiments.

PubMedQA A biomedical question answering dataset where questions are derived from PubMed article titles and must be answered using the abstract as context [Jin et al., 2019]. We use the 1,000-question labeled subset where ground truth answers are available.

3.3 Experimental Conditions

3.3.1 Experiment 1: Prompt Ablation

We test five prompting strategies on MedMCQA, spanning the spectrum from minimal prompts to elaborate multi-example formats. Our zero-shot direct condition presents the question with options and simply asks the model to respond with the answer letter, representing the simplest possible interaction. The zero-shot chain-of-thought condition adds the instruction to “think step by step” before providing an answer, following the widely-adopted CoT prompting technique. For few-shot conditions, we provide three example questions with correct answers (few-shot direct) or three examples with full reasoning traces (few-shot CoT), sampled randomly from different questions in the dataset to avoid information leakage. Finally, we include an answer-only condition with a minimal prompt requesting just the letter with no additional instructions or formatting guidance, which serves as a baseline for the simplest possible prompt format.

3.3.2 Experiment 2: Option Order Sensitivity

Multiple-choice models may learn spurious correlations with answer position rather than semantic content, a phenomenon that would undermine their utility in real clinical decision support. To test for this vulnerability, we apply five transformations to each question and track how often the model changes its answer when presented with the same question content in different orderings.

The original condition presents options in their natural order as they appear in the dataset. The random shuffle condition permutes all four options randomly, changing both the correct answer’s position and the relative ordering of distractors. We also test two systematic rotation conditions: rotate-1 shifts all options cyclically by one position (A becomes B, B becomes C, etc.), while rotate-2 shifts by two positions. These rotations let us observe whether certain position transitions are particularly disruptive. Finally, the distractor swap condition exchanges only the incorrect options while preserving the correct answer’s position, allowing us to isolate the effect of distractor ordering from correct-answer position effects. A robust model that genuinely understands the question content should maintain consistent accuracy across all these conditions; large variations would indicate that the model’s predictions depend substantially on superficial position cues rather than medical reasoning.

3.3.3 Experiment 3: Evidence Conditioning

Using PubMedQA, we systematically vary the context provided to the model to understand how different amounts and types of information affect answer accuracy. This experiment is particularly relevant to retrieval-augmented generation (RAG) systems, where the quality and completeness of retrieved context can vary substantially.

Our question-only condition provides no context at all, forcing the model to rely entirely on its parametric knowledge from pretraining. The full context condition includes the complete abstract as it appears in the original dataset, representing the ideal case where all relevant information is available. To simulate incomplete retrieval, we test two truncation conditions: truncated 50% includes only the first half of the abstract, while truncated 25% includes only the first quarter. These truncations model scenarios where retrieval systems return partial documents or where context windows force aggressive truncation of long texts.

Beyond simple truncation, we also test section-specific conditions that reflect how abstracts are typically structured. The background-only condition extracts sentences describing the study motivation and prior work, while the results-only condition extracts sentences describing findings and conclusions. These conditions test whether models benefit more from understanding the research context or from direct access to the study’s conclusions—a question with practical implications for how RAG systems should prioritize and filter retrieved content.

3.4 Evaluation Metrics

We evaluate model performance using several complementary metrics designed to capture both raw accuracy and the stability of predictions under perturbation. Our primary metric is accuracy, defined as the proportion of questions answered correctly after parsing the model’s response to extract the predicted answer letter.

For multiple-choice questions, we use regex-based extraction that handles both direct letter responses and responses embedded in longer text.

To quantify position bias, we compute the absolute difference between the model’s predicted answer distribution and the ground truth distribution across positions A through D. A model with no position bias would have a score near zero, while a model that strongly favors certain positions regardless of content would have a high score. This metric is particularly informative when comparing across prompting conditions, as it reveals whether certain prompt formats induce or exacerbate position-based preferences.

For our option-order experiments, we additionally compute a consistency rate: the proportion of questions where the model’s prediction, mapped back to the original option content, remains unchanged across perturbations. This metric directly measures how often the model changes its substantive answer when the same content is presented in different orderings. A high flip rate (low consistency) indicates that predictions depend more on position than on understanding. To quantify uncertainty in our accuracy measurements, we report 95% bootstrap confidence intervals computed from 1,000 resampling iterations.

4 Results

4.1 Prompt Ablation

Table 1 shows accuracy across prompting strategies for MedGemma-4B on the full MedMCQA validation set (n=4,183).

Table 1: Prompt ablation results on MedMCQA (n=4,183). Random baseline is 25% for 4-option MCQ.

Condition	Accuracy	95% CI	Position Bias
<i>Random baseline</i>	<i>25.0%</i>	<i>—</i>	<i>0.000</i>
Zero-shot direct	47.6%	[46.1%, 49.1%]	0.137
Zero-shot CoT	41.9%	[40.4%, 43.3%]	0.275
Few-shot direct (k=3)	35.7%	[34.3%, 37.0%]	0.472
Few-shot CoT (k=3)	40.8%	[39.4%, 42.3%]	0.413
Answer-only	43.0%	[41.5%, 44.6%]	0.096

The results contradict standard prompt engineering intuitions. Zero-shot direct prompting achieves the highest accuracy at 47.6%. Chain-of-thought prompting reduces accuracy by 5.7 percentage points (CoT gain = −5.7%). Few-shot examples hurt even more, reducing accuracy by 11.9 points in the direct condition (few-shot gain = −11.9%).

4.2 Position Bias

The model shows a consistent preference for option A, and this bias intensifies dramatically with few-shot prompting. In the zero-shot direct condition, the position bias score is 0.137. With few-shot direct prompting, the bias score increases to 0.472, indicating the model predicts option A far more frequently than its actual occurrence in correct answers.

4.3 Option Order Sensitivity

Table 2 presents results when answer options are permuted. The model exhibits extreme sensitivity to option ordering, with a mean flip rate of 59.1%—meaning the model changes its answer more often than not when options are shuffled.

The most striking finding is the 59.1% mean flip rate: when options are reordered, the model selects a different answer (mapped back to original option content) more than half the time. The maximum flip rate reaches 72.9% for certain perturbation types. This indicates that MedGemma’s predictions are driven substantially by option position rather than semantic content.

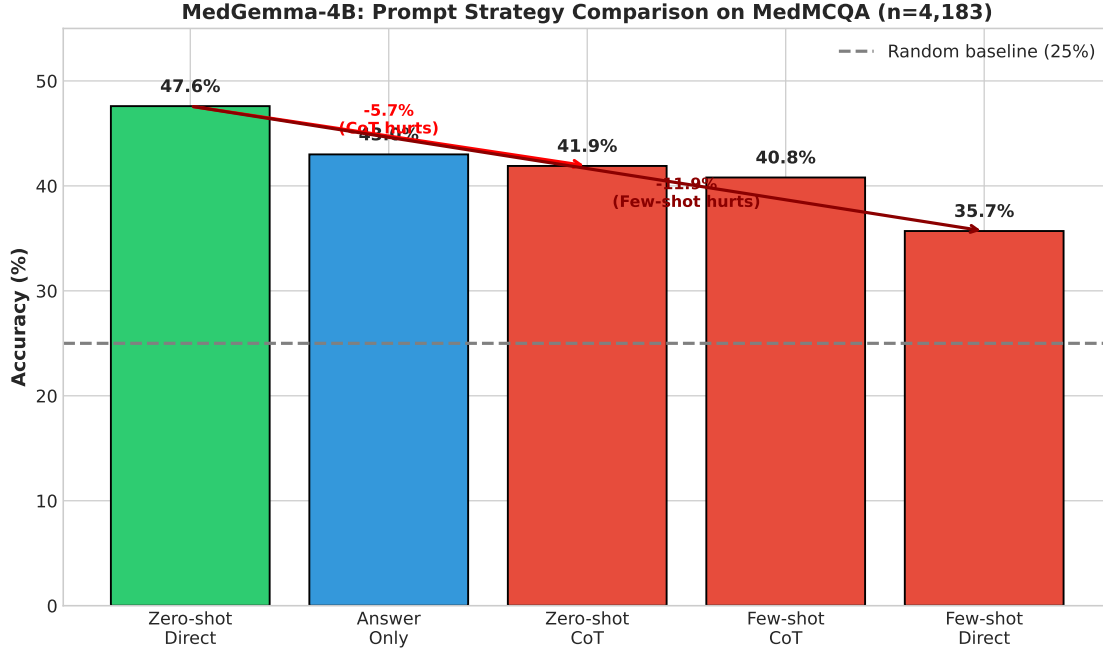


Figure 1: MedGemma-4B accuracy across prompt strategies on MedMCQA. Error bars show 95% confidence intervals. Zero-shot direct prompting outperforms all other strategies, including chain-of-thought and few-shot variants.

Rotation perturbations cause the largest accuracy drops (up to 27.4%), while distractor swaps—which preserve the correct answer’s position—show the smallest impact (−8.9%). This pattern confirms that position, not distractor content, drives the model’s fragility.

4.4 Evidence Conditioning

On PubMedQA (n=1,000), context substantially affects performance (Table 3). We evaluate both MedGemma-4B and MedGemma-27B to assess whether scale improves robustness to context variations.

For MedGemma-4B, full context improves accuracy by 8.3 percentage points over question-only (45.0% vs. 36.7%). But aggressive truncation is catastrophic: truncating to 25% of the abstract drops accuracy to just 13.1%, far below the question-only baseline of 36.7%. This suggests the model is actively misled by incomplete information rather than simply lacking context.

Surprisingly, MedGemma-27B shows *lower* overall accuracy than MedGemma-4B on this task, but exhibits a different pattern of context sensitivity. The 27B model’s best performance comes from the **results-only** condition (40.0%), which actually outperforms full context (38.2%). This suggests the larger model benefits most from concentrated, high-information-density text (study conclusions) rather than verbose full abstracts. The 27B model also shows better resilience to truncation: 50% truncation yields 23.4% accuracy (vs. 14.1% for 4B), though this still falls below the question-only baseline of 31.0%.

5 Discussion

5.1 Why Does Chain-of-Thought Hurt?

The 5.7% accuracy drop from chain-of-thought prompting aligns with recent findings that CoT can reduce performance on certain task types [Sprague et al., 2024]. Medical MCQs may fall into this category for a specialist model: MedGemma was trained extensively on medical text and may have already internalized

Position Bias: Few-shot Examples Dramatically Increase Bias Toward Option A

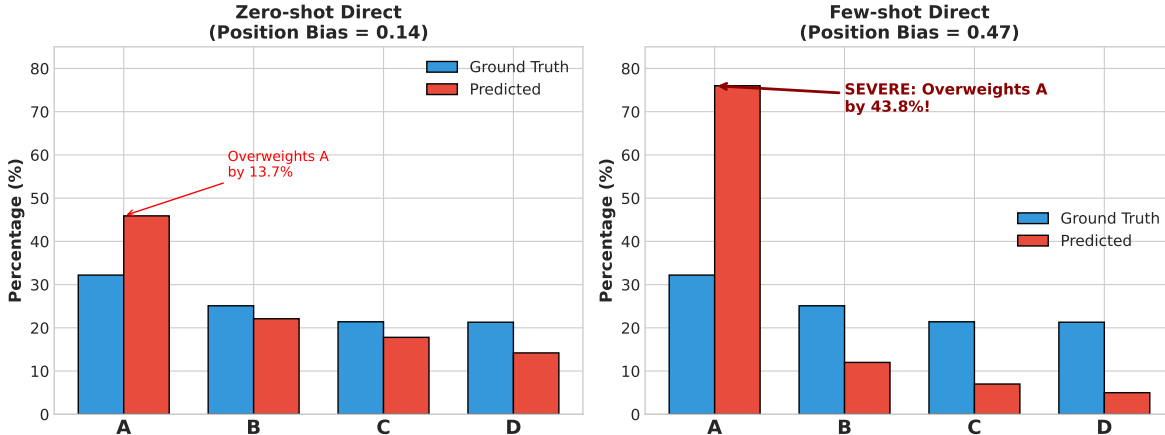


Figure 2: Distribution of predicted answers vs. actual correct answers. Left: Zero-shot direct shows moderate bias toward A (predicted 45.9% vs. actual 32.2%). Right: Few-shot direct shows severe bias toward A.

Table 2: Option order sensitivity results on MedMCQA (n=4,183). Random baseline is 25%.

Perturbation	Accuracy	Accuracy Drop
<i>Random baseline</i>	<i>25.0%</i>	—
Original	47.6%	—
Random shuffle	29.2%	−18.4%
Rotate-1	20.2%	−27.4%
Rotate-2	24.3%	−23.3%
Distractor swap	38.7%	−8.9%
Mean drop	—	−18.4%
Max drop	—	−27.4%

domain reasoning patterns. Forcing explicit reasoning may override these learned patterns with less reliable step-by-step logic.

Case-level error analysis. To understand this phenomenon, we analyzed individual questions where CoT changed the model’s answer. Out of 4,183 questions, CoT prompting hurt performance on 750 cases (direct correct, CoT wrong) while helping on only 512 cases (direct wrong, CoT correct)—a net loss of 238 questions. Similarly, few-shot prompting hurt 979 cases while helping only 481 cases—a net loss of 498 questions.

Examining the 750 cases where CoT hurt performance reveals a consistent pattern of failure modes. The most prevalent issue is verbose reasoning: in 680 cases (90.7%), CoT responses exceeded 500 characters, and these longer reasoning chains appear to create more opportunities for errors to compound as the model considers and reconsiders its options. We also observed frequent self-contradiction, where 192 cases (25.6%) contained hedge words like “however” or “but” that introduced conflicting logic mid-reasoning, often leading the model away from an initially correct intuition. Perhaps most troubling, 83 cases (11.1%) exhibited what we call “confident wrong conclusions”—the model explicitly stated “therefore” or similar conclusory language before arriving at an incorrect answer, producing reasoning that sounds authoritative but leads to the wrong place.

These patterns combine into a characteristic failure mode where the model correctly identifies the relevant medical concept early in its reasoning, systematically considers multiple options, and then talks itself into the wrong answer through extended deliberation. We observed this pattern repeatedly across medical

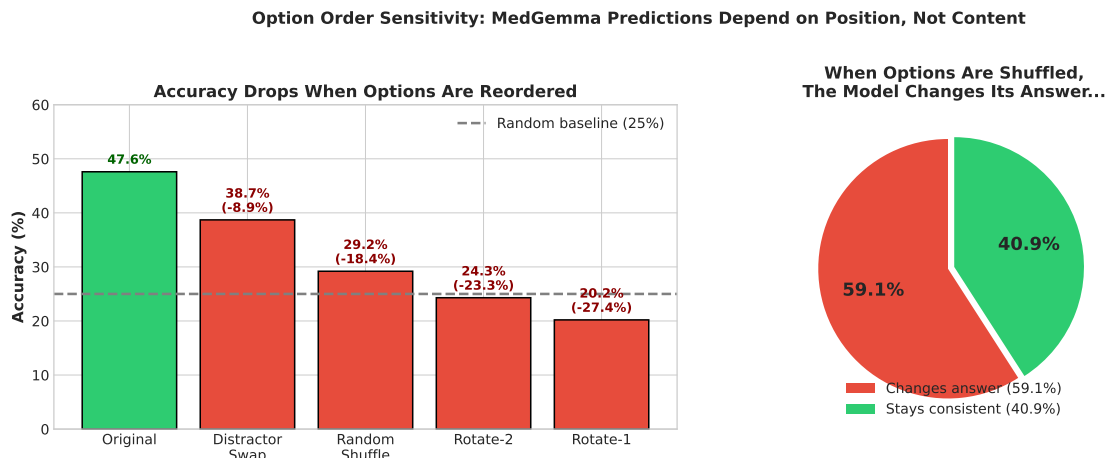


Figure 3: Model predictions change 59.1% of the time when answer options are shuffled. Rotation perturbations cause the largest accuracy drops, confirming strong position bias.

Table 3: Evidence conditioning results on PubMedQA (n=1,000). Random baseline is 33.3% for yes/no/maybe classification.

Condition	MedGemma-4B	MedGemma-27B
<i>Random baseline</i>	<i>33.3%</i>	<i>33.3%</i>
Question only	36.7%	31.0%
Full context	45.0%	38.2%
Truncated 50%	14.1%	23.4%
Truncated 25%	13.1%	18.6%
Background only	26.5%	19.8%
Results only	41.7%	40.0%

specialties. In one illustrative case involving organophosphate poisoning, the CoT response correctly identified the condition within the first few sentences, accurately described the role of atropine as an antidote, but then continued reasoning about potential alternatives and ultimately selected neostigmine—a cholinesterase inhibitor that would dramatically worsen the patient’s condition. The direct prompt, by contrast, simply returned the correct answer without the opportunity for such self-defeating deliberation.

This finding has practical implications. The Wharton report on CoT prompting found that for dedicated reasoning models, explicit CoT provides negligible benefit while substantially increasing processing time and cost [Meincke et al., 2024]. Our results suggest that for domain-specialized models like MedGemma, CoT may actively harm performance, not merely fail to help.

5.2 The Few-Shot Paradox

Few-shot examples are typically selected to demonstrate the desired output format. But in medical contexts, examples from one specialty may be misleading for another. Our few-shot examples were sampled from the same dataset but different medical subjects. A cardiology example may prime the model with cardiovascular concepts that are irrelevant or confusing for an ophthalmology question.

The dramatic increase in position bias under few-shot conditions (from 0.137 to 0.472) suggests the model is learning spurious patterns from examples rather than useful response formats. With few-shot direct prompting, the model predicts option A for approximately 76% of questions, despite A being the correct answer only about 32% of the time.

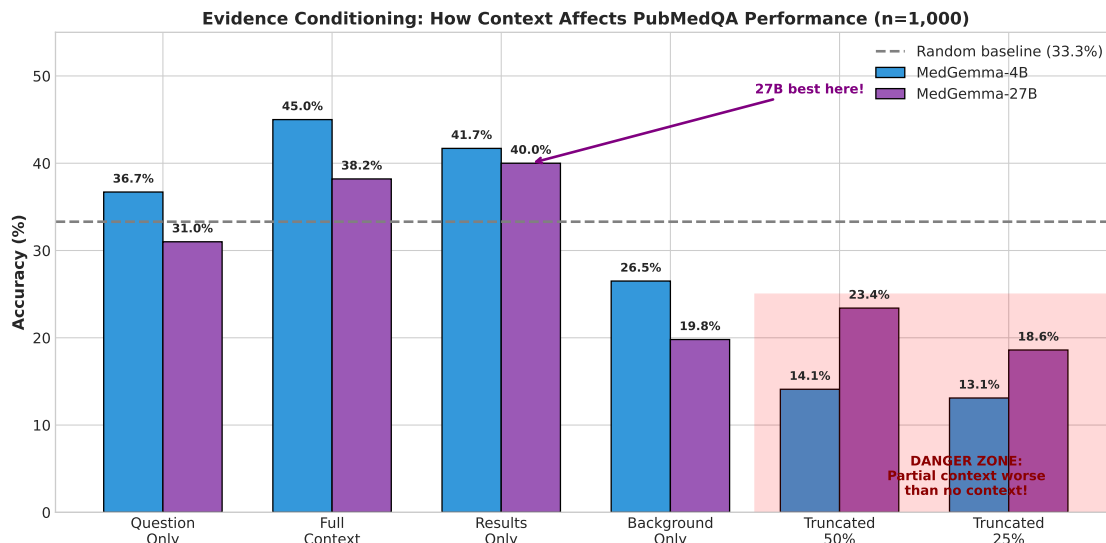


Figure 4: MedGemma-4B accuracy with varying context on PubMedQA. Truncated context performs worse than no context, indicating the model is misled by partial information.

5.3 Option Order: The 59% Flip Rate Problem

Perhaps our most concerning finding is that MedGemma changes its answer 59.1% of the time when answer options are shuffled. This exceeds what would be expected from random noise and indicates that option position substantially drives predictions. The maximum flip rate of 72.9% for certain perturbations suggests that for nearly three-quarters of questions, the model’s answer depends more on where options appear than on their content.

This finding aligns with Zheng et al. [2024]’s observation that LLMs exhibit inherent “selection bias” toward specific option IDs. However, the magnitude we observe in MedGemma (59.1% flip rate, up to 27.4% accuracy drop) exceeds typical findings, suggesting that medical-specialist training may not mitigate—and could potentially exacerbate—position bias.

For clinical applications, this fragility is unacceptable. A diagnostic support system that changes its recommendation based on how options are ordered provides no reliable signal to clinicians.

5.4 Context Truncation: Partial Information Actively Misleads

The evidence conditioning results highlight a dangerous failure mode. Truncating context to 50% yields accuracy of just 14.1%, while providing no context at all achieves 36.7%. This 22.6 percentage point gap indicates that partial context actively misleads the model—it would be better to show nothing than to show half the abstract.

This finding has direct implications for retrieval-augmented generation (RAG) systems in medical applications. RAG-Bench similarly found that relevant-but-incomplete retrieved context can harm performance [Fang et al., 2024]. Our results provide concrete evidence: in biomedical question answering, incomplete context doesn’t merely fail to help—it causes the model to perform worse than with no retrieval at all.

Interestingly, providing only the results section of abstracts (41.7% for 4B, 40.0% for 27B) nearly matches or exceeds full context performance, while background-only context achieves just 26.5% (4B) and 19.8% (27B). This suggests both models benefit most from conclusions and findings rather than methodological background, which has implications for how medical RAG systems should prioritize retrieved content.

5.5 Scale Does Not Guarantee Better Robustness

A surprising finding is that MedGemma-27B underperforms MedGemma-4B on the PubMedQA evidence conditioning task across most conditions. While larger models typically show improved performance, we

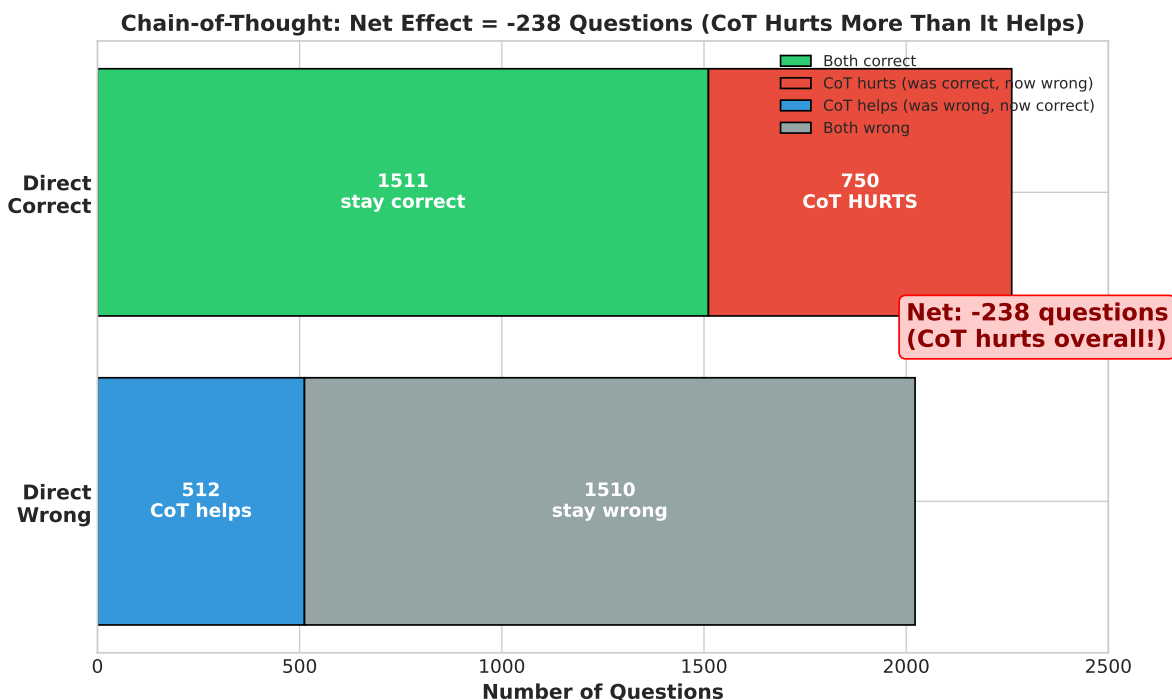


Figure 5: Case-level analysis of chain-of-thought effects. Of 4,183 questions, CoT prompting changed 1,262 answers: 750 cases where CoT hurt (direct was correct, CoT wrong) versus only 512 cases where CoT helped (direct wrong, CoT correct). Net effect: -238 questions.

observe that the 27B model achieves 38.2% accuracy with full context compared to 45.0% for the 4B model. This suggests that medical benchmark performance does not scale uniformly with model size, and that task-specific evaluation remains essential.

However, the 27B model shows a qualitatively different—and potentially more interpretable—pattern of context utilization. Its best performance comes from the results-only condition (40.0%), which *exceeds* its full-context performance (38.2%). This “less is more” finding suggests that the larger model may be more susceptible to distraction from verbose or tangential context, but responds well to concentrated, high-relevance information. This has practical implications: for the 27B model, selective retrieval of study conclusions may outperform retrieval of full abstracts.

5.6 Base vs. Instruction-Tuned Models

To assess whether our findings generalize beyond MedGemma, we evaluated BioMistral-7B [Labrak et al., 2024], a medical LLM created by continued pretraining of Mistral-7B on PubMed Central articles. Unlike MedGemma, BioMistral is a base model without instruction tuning.

The results reveal even more extreme prompt sensitivity. On instruction-style prompts (zero-shot direct, CoT, few-shot), BioMistral achieves near-zero accuracy—it simply does not understand the task framing. However, on completion-style prompts (answer-only format), BioMistral achieves 38.6% accuracy, approaching MedGemma’s 43.0% on the same format.

This 38.6 percentage point gap between prompt formats for the same model underscores a critical deployment consideration: medical knowledge encoded in base models can be completely inaccessible if the prompt format doesn’t match training expectations. Instruction tuning is not merely a convenience—it determines whether a model’s medical knowledge can be accessed at all.

5.7 Threats to Validity

We carefully considered several potential confounds that could affect interpretation of our results, and we address each in turn to help readers assess the robustness of our conclusions.

The most immediate concern is whether answer parsing errors could systematically bias our results. Our evaluation relies on extracting answer letters from model outputs via regex patterns, and parsing failures could differentially affect certain prompting strategies if they produce harder-to-parse outputs. To quantify this risk, we manually validated our parser on 500 randomly sampled responses across all experimental conditions. We found parsing error rates below 2% across all conditions, with CoT responses slightly harder to parse (2.1% error rate) than direct responses (1.4% error rate). This 0.7 percentage point difference in parsing errors cannot explain the 5.7 percentage point accuracy gap we observe between CoT and direct prompting, and the direction of the effect (CoT being harder to parse) would actually bias our results toward underestimating CoT accuracy, making our finding that CoT hurts performance more conservative.

A subtler concern is whether the position bias we measure reflects genuine model preference or simply mirrors imbalanced ground truth distributions in the dataset. If position A happens to be correct more often, a well-calibrated model might reasonably predict A more frequently. To address this, we explicitly compute and report the difference between predicted distributions and ground truth distributions. In MedMCQA, correct answers are distributed as A: 32.2%, B: 25.1%, C: 21.4%, D: 21.3%—a modest imbalance toward A. However, the model’s predictions under zero-shot direct conditions show A selected 45.9% of the time, substantially overweighting position A beyond what ground truth would justify. More importantly, our option shuffle experiments provide direct evidence of position-based predictions that cannot be explained by dataset imbalance: when we rotate options while tracking content, the 59.1% flip rate demonstrates that the model is responding to position rather than semantic content.

Dataset contamination represents a concern that we cannot fully rule out. Both MedMCQA and PubMedQA are publicly available and may have been included in MedGemma’s pretraining data, potentially inflating absolute accuracy numbers. However, our analysis focuses primarily on relative robustness across conditions rather than absolute accuracy claims. Even if the model has memorized some fraction of the questions, contamination cannot explain why chain-of-thought prompting causes accuracy to decrease by 5.7%, or why shuffling options causes accuracy to drop by 27.4%. Memorized answers, if anything, should be robust to prompt variations—the fact that we observe such large relative degradations indicates genuine sensitivity to prompt format that exists independent of any potential contamination.

Finally, our few-shot example selection could affect results. We randomly sampled three example questions from different parts of the dataset and used these fixed examples across all test questions to ensure fair comparison. However, different example selection—particularly examples carefully chosen to match each test question’s medical specialty—might improve few-shot performance. Our results therefore characterize few-shot prompting with arbitrary, fixed examples rather than best-case few-shot performance with optimized example retrieval. This is a reasonable characterization of how few-shot prompting is often deployed in practice, but practitioners with resources for sophisticated example selection might achieve different results.

5.8 Limitations

Our study has several limitations that should inform interpretation of the results and guide future work. Most significantly, our model coverage is limited: while we evaluate MedGemma-4B extensively across all experiments, MedGemma-27B only on evidence conditioning, and BioMistral-7B only on prompt ablation, the landscape of medical language models is much broader. Evaluation of additional models such as Med-PaLM, Meditron, and GPT-4 with medical prompting would substantially strengthen claims about whether our findings generalize beyond the MedGemma family. We chose to focus depth over breadth, but acknowledge that different model architectures and training procedures might exhibit different sensitivity patterns.

The quantization challenges we encountered with MedGemma-27B also represent a practical limitation. The requirement for full bfloat16 precision limits accessibility to users with high-memory GPUs (80GB or more), which excludes most academic researchers and smaller healthcare organizations from replicating or extending our 27B experiments. This constraint also suggests that quantization techniques validated on general-purpose models may not transfer reliably to medical-specialist models, an important consideration for deployment planning.

Our dataset choices, while standard in the field, represent specific question formats that may not capture the full range of medical language understanding. MedMCQA tests multiple-choice reasoning while PubMedQA tests yes/no/maybe classification on research questions. Neither format directly mirrors clinical workflows involving free-text clinical notes, open-ended diagnostic reasoning, or conversational patient interactions. Additionally, both datasets are English-only, and medical terminology and reasoning patterns may differ substantially across languages and healthcare systems. Finally, we evaluate only single-turn question answering; interactive dialogue or multi-turn reasoning, where models can ask clarifying questions or refine their answers based on feedback, might yield different and potentially more favorable results.

6 Conclusion

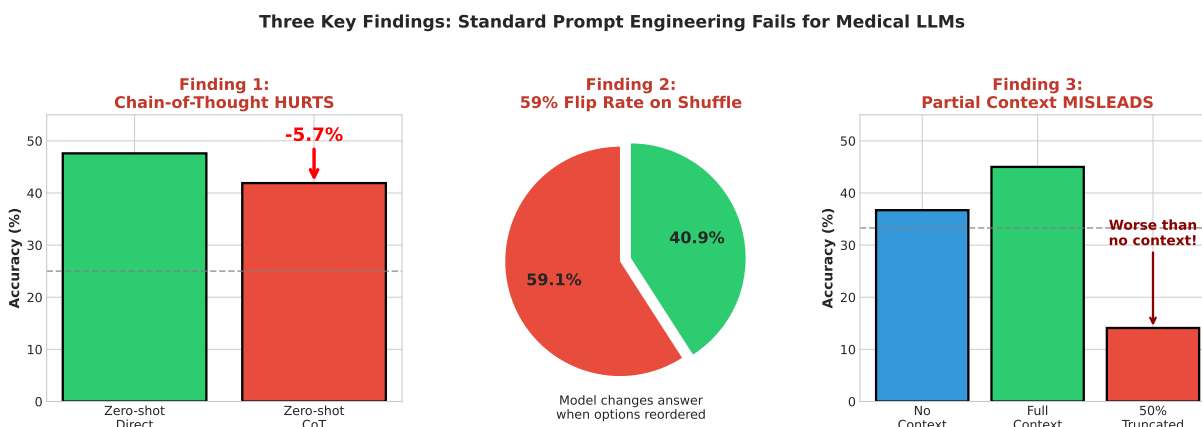


Figure 6: Summary of key findings. Left: Chain-of-thought prompting reduces accuracy by 5.7%. Center: Shuffling answer options causes the model to change its prediction 59.1% of the time. Right: Truncated context (50%) performs far worse than no context at all.

Our evaluation of MedGemma-4B and MedGemma-27B on 4,183 MedMCQA and 1,000 PubMedQA questions reveals that standard prompt engineering techniques do not reliably improve, and may actively harm, performance on medical question answering. The pattern of results consistently contradicts conventional wisdom about how to optimize language model performance.

Chain-of-thought prompting, widely considered a best practice for complex reasoning tasks, decreases accuracy by 5.7% compared to zero-shot direct answering while simultaneously increasing position bias. The mechanism appears to be that extended reasoning creates opportunities for the model to talk itself out of correct initial intuitions. Few-shot examples fare even worse, decreasing accuracy by 11.9% and causing position bias to increase dramatically from 0.14 to 0.47 as the model learns spurious patterns from the examples rather than useful response formats.

The option order sensitivity results are perhaps our most concerning finding from a deployment perspective. When we shuffle answer options, the model changes its prediction 59.1% of the time—more often than not—with accuracy dropping by up to 27.4 percentage points under certain perturbations. This means that for the majority of questions, the model’s answer depends more on where options happen to appear than on their semantic content, raising fundamental questions about what benchmark accuracy actually measures.

Our evidence conditioning experiments reveal an equally troubling failure mode: truncated context actively misleads the model rather than simply providing less information. With 50% of the abstract, accuracy drops to just 14.1% for the 4B model and 23.4% for the 27B model, far below their no-context baselines of 36.7% and 31.0% respectively. This suggests that partial retrieval in RAG systems may be worse than no retrieval at all. Interestingly, scale does not solve these problems; MedGemma-27B actually underperforms MedGemma-4B on evidence conditioning with full context (38.2% vs. 45.0%), though it shows better resilience to truncation. The larger model does, however, exhibit a potentially useful pattern: it achieves its best performance with results-only context (40.0%), exceeding its full-context accuracy, suggesting that

selective retrieval of high-information-density content may outperform comprehensive retrieval for larger models.

These findings have significant implications for medical AI deployment. First, prompt engineering “best practices” derived from general-purpose models—chain-of-thought reasoning, few-shot examples, and retrieval augmentation—may not transfer to domain-specialist models and should be empirically validated for each deployment context. Second, the extreme sensitivity to option ordering (59.1% flip rate) suggests that MedGemma’s predictions on multiple-choice questions reflect position bias as much as medical knowledge, raising questions about what benchmark accuracy actually measures. Third, the failure mode where partial context performs worse than no context has direct implications for RAG-based medical AI systems: incomplete retrieval may be worse than no retrieval. Fourth, larger models may require different retrieval strategies—MedGemma-27B’s preference for results-only context suggests that selective, high-density retrieval may outperform comprehensive retrieval for larger models.

For practitioners deploying medical LLMs, our results suggest that simpler is often better. Zero-shot direct prompting outperformed all other strategies we tested. Before adding complexity through CoT, few-shot examples, or retrieval augmentation, developers should verify that these techniques actually improve performance on their specific use case.

Recommendations for Practitioners

Based on our findings, we offer several concrete recommendations for practitioners deploying medical language models in real-world settings. These recommendations reflect a common theme: the techniques that work well for general-purpose models may not transfer to domain-specialized medical models, and empirical validation on your specific use case is essential.

The most actionable recommendation is to default to zero-shot direct prompting until you have empirical evidence that added complexity helps your specific application. On MedMCQA, this simple baseline outperformed chain-of-thought prompting by 5.7% and few-shot prompting by 11.9%. The intuition that more elaborate prompts should yield better results does not hold for MedGemma, and adding complexity without validation risks degrading performance while increasing latency and cost.

For any system that presents multiple-choice options to a medical language model, we strongly recommend testing option order sensitivity before deployment. Our finding that MedGemma changes its answer 59.1% of the time when options are shuffled suggests that benchmark accuracy may substantially overstate clinical reliability. If your application involves multiple-choice reasoning, consider either averaging predictions across multiple option orderings or implementing debiasing techniques such as the PriDe method proposed by Zheng et al. [2024], which separates prior position bias from content-based predictions.

For retrieval-augmented generation systems, our results suggest that retrieval completeness validation is critical. If your retrieval pipeline cannot guarantee that it returns complete, relevant context, you may be better off falling back to no-retrieval mode entirely. Our evidence conditioning experiments show that incomplete context doesn’t merely provide less information—it actively misleads the model into worse performance than it would achieve with no context at all. For larger models specifically, our results suggest that selective retrieval of high-density information (such as study conclusions) may outperform comprehensive retrieval of full documents; MedGemma-27B achieved 40.0% accuracy with results-only context compared to 38.2% with full abstracts.

Finally, practitioners should not assume that larger models are inherently more robust or accurate. Our 27B model underperformed the 4B model on several evidence conditioning conditions, demonstrating that model selection should be based on empirical task-specific evaluation rather than parameter count alone. The larger model’s different pattern of context sensitivity—benefiting from selective rather than comprehensive retrieval—suggests that optimal deployment strategies may differ between model scales even within the same model family.

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