

Managing Placebo and Nocebo Effects during Patient-provider interactions: A Consensus Statement and Clinical Recommendations for Healthcare Professionals

Meeuwis, S.H., Evers, A.W.M., Bagnis A., K. Mattarozzi, K. ... on behalf of the PANACEA consortium

This document represents the final output of the PANACEA project, namely the clinical recommendations for healthcare professionals on how to manage placebo and nocebo effects in clinical practice; it also constitutes a pre-peer-reviewed version that will be submitted to a scientific journal for publication. The supplementary materials referred to in the text are provided at the end of the document.



Abstract

Importance. Contextual factors, including patient-provider interaction alongside the information that is provided during consultations, significantly influence patients' expectations toward treatment and subsequent health outcomes. These expectations are pivotal determinants of placebo and nocebo effects which in turn affect treatment success. Healthcare professionals often overlook the extent of their impact in this regard and report feeling unprepared to manage placebo and nocebo effects.

Objective. Structured best practice clinical guidance on managing placebo and nocebo effects, specifically on the use of and communication about these effects during patient-provider interactions, are lacking. For these reasons, on behalf of the European cooperation partnership PANACEA, formed by an interdisciplinary group of experts, we have developed recommendations that aim to provide guidance for leveraging communication to enhance placebo effects and mitigate nocebo effects.

Evidence review. The method that was used to formulate these recommendations is based on two streams of evidence: 1) expert consensus and existing guidance, including insights from prior narrative and systematic reviews, consensus papers, and an existing eLearning module that reviewed literature until 2021, and 2) a systematic review of the current empirical evidence, substantiating the recommendations with best practice evidence. The process involved iterative rounds and consensus-building efforts.

Findings. The recommendations specifically aim to increase knowledge on how to 1) optimize the patient-provider relationship; 2) ask about a patient's expectations for a treatment; 3) discuss the treatment rationale; 4) discuss a treatment's risks and side effects with a patient and 5) explain to a patient how placebo and nocebo effects work.

Conclusions and relevance. Taken together, the recommendations are generally well supported by existing evidence, although some strategies require further study. In the future, this knowledge should be translated to regular training and educational tools to normalize its application into daily educational and clinical practice.

KEYWORDS: placebo effects, nocebo effects, clinical recommendations, expectations, communication

Introduction

Placebo and nocebo effects influence the efficacy of many different treatments^{1,2}, ultimately impacting treatment success and overall health. Placebo effects are beneficial treatment effects, that arise from contextual factors and the patient's positive expectations concerning the treatment, rather than from its active components. Nocebo effects refer to phenomena where expectations of adverse effects or harm can lead to those negative outcomes². Expectations can be shaped by learning processes³ (Figure 1), many of which are relevant during patient-provider interaction and influenced by the dynamics within the patient-provider relationship⁴. Consequently, expectations can become a focal point of therapeutic intervention, for instance by integrating contextual factors within treatment, utilizing communication to target optimal outcome expectations^{5,6}, and informing about placebo and nocebo effects^{2,5}.

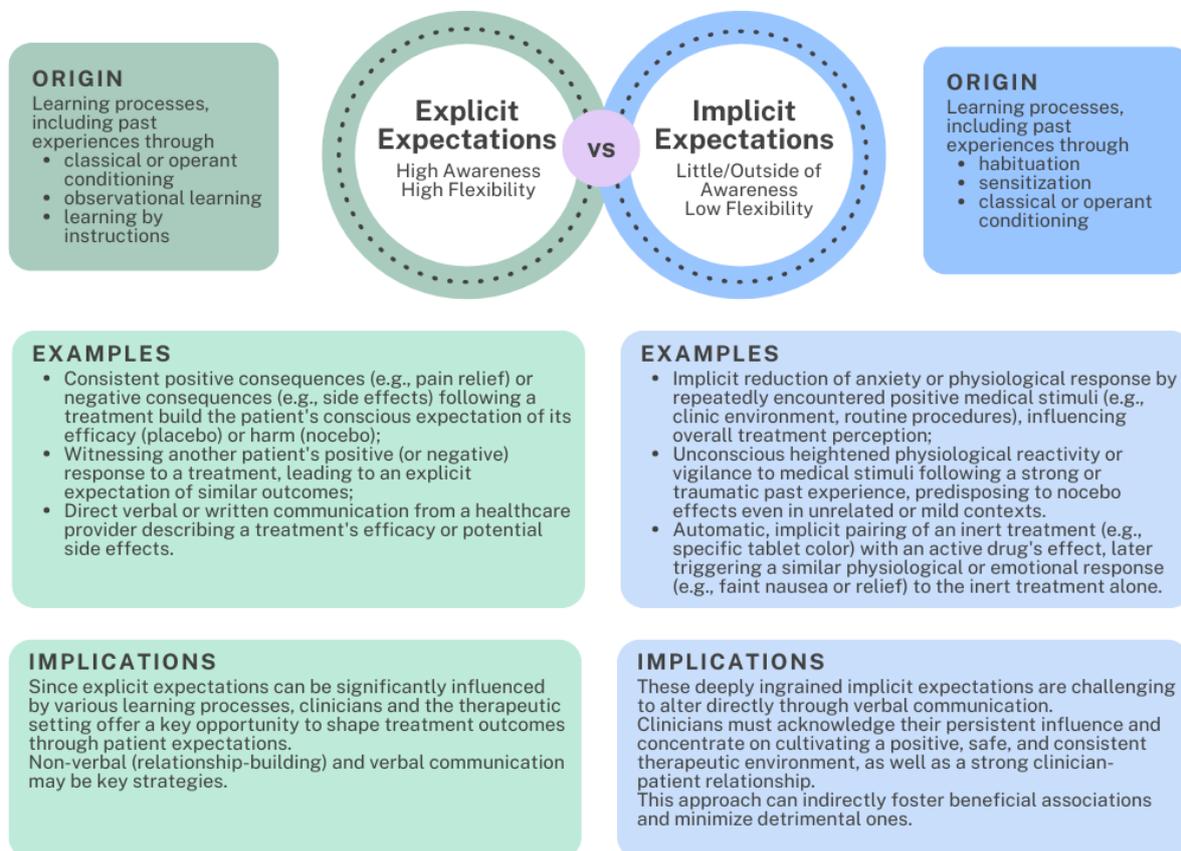


Figure 1. Understanding Expectations: Origins and Clinical Implications.

While existing work offers insights into the impact of placebo and nocebo effects in daily practice, a critical gap remains in the form of comprehensive, truly evidence-based recommendations. Such guidelines are needed to equip clinicians to amplify placebo effects and manage nocebo effects in routine care, ultimately aiming to improve patient

health. Healthcare professionals have highlighted the need for clearer training and support in this area⁷. From the necessity of an international and interdisciplinary collaboration to address this gap, the PANACEA Consortium emerges as one of the leading entities striving to bridge the gap between scientific findings on placebo/nocebo effects and their practical implementation in clinical settings.

The aim of this paper is to set forth best practice recommendations for healthcare professionals, that are rooted in existing evidence-based knowledge and can be easily implemented during patient-provider interactions.

Methods

The method that was used to formulate these recommendations is based on a comprehensive integration of two distinct yet complementary evidence streams. The first stream comprised foundational statements, derived from expert consensus and existing guidance, including insights from prior narrative reviews, consensus papers, a previous systematic review¹, and an existing eLearning module⁸, which reviewed literature until 2021. The second stream involved a systematic substantiation and enrichment of these expert-informed statements with current empirical evidence. This evidence was identified through a dedicated systematic review of empirical studies in patient populations, covering literature until May 2024. This systematic review was registered on Prospero (CRD420250642824).

For each recommendation, information from both evidence streams was meticulously synthesized into a single, cohesive statement. The recommendations are formulated based on the clinical context in which treatments are applied, as well as the limits and applications of placebo and nocebo effects specific to that domain. The interdisciplinary nature of the PANACEA consortium—including psychologists, clinicians, nurses, pharmacologists, and experts in placebo, nocebo, and health communication—enhanced the depth of our approach. The iterative process of refining these recommendations involved collecting feedback from consortium members, ensuring alignment with regular consensus-building principles and methods⁹. Details on the methodology and the resulting evidence from each stream are provided in the Supplementary Materials.

Results

Rooted in the crucial role of expectations in modulating placebo and nocebo effects, as well as the clinician's potential for intervention, several domains of medical communication were identified from the two evidence streams that can optimise placebo effects and minimise nocebo effects: i) optimising the clinician-patient relationship, ii) asking about expectations, iii) discussing the treatment rationale, iv) discussing risks and

side effects, and v) explaining placebo and nocebo effects. The quality of evidence is described in Table 1.

Table 1. Quality of evidence for the clinical recommendations.

Domain	Clinical Recommendation	Recommendation classification*	
		Class	Level of evidence
Optimizing the clinician-patient relationship	Show interest in the patient	I	B-R, B-NR
	Give reassurance about ongoing care	IIa	B-R, B-NR
	Respond to negative emotions	IIa	B-R, B-NR
	Leverage eye contact and physical presence	IIa	B-R, B-NR
	Use patient-centered communication	IIa	B-R, B-NR
	Adapt communication style to the patient	IIa	B-R, B-NR
	Leverage other behaviours to foster trust	IIa	B-R, B-NR
	<i>Overall recommendation**</i>	IIa	B-R
Asking about expectations	Ask about and identify general expectations towards treatment	IIa	C-EO
	Adjust unrealistic expectations	IIb	C-EO
	Address disappointment from previous treatments	IIb	C-EO
	<i>Overall recommendation**</i>	IIa	B-R, B-NR
Discussing the treatment rationale	Provide correct information based on evidence-based knowledge	I	B-R
	Avoid exaggeration or underestimation of treatment results	IIa	B-R, B-NR
	Clearly state the treatment rationale and expected positive outcomes in concrete terms	IIa	B-R, B-NR
	Explain treatment mechanisms of action	IIa	B-R
	Align patient expectations and beliefs with evidence-based care	IIa	B-R, B-NR
	Emphasize the purpose and advantages of treatment procedures	IIa	B-R
	Check patient comprehension	IIa	B-R, B-NR
Suggest reliable information sources	IIa	B-R	
	<i>Overall recommendation*</i>	IIa	B-R
Discussing risks and side effects	General structure of the consultation		
	Emphasize positive consequences of side effects	IIb	C-LD
	Balance side effect information with positive information	IIb	B-R, B-NR
	Investigate (sources of) nocebo effects	IIa	C-LD
	State all symptoms briefly to reduce nocebo effects	IIb	C-LD
	Information framing		
	Frame positive outcomes rather than negative outcomes	IIa	C-LD
	Discuss risks in statistical terms	IIa	B-R
Use neutral terms when performing a procedure	IIa	C-LD	
	<i>Overall recommendation*</i>	IIa	B-R
Explain placebo and nocebo effects	Placebo effects		
	Careful explanation of placebo effects	IIa	B-R
	Mechanism of action of placebo effects	IIa	B-R, B-NR
	Appropriate presentation of information	IIb	B-R
	Prevent overly optimistic expectations	IIa	C-EO
	Nocebo effects		
Attend to the purpose of explaining nocebo effects	IIa	C-EO	

Careful explanation of placebo effects	IIa	B-R, B-NR
Keep explanation simple	IIb	C-EO
Avoid unintentional negative effects	IIb	C-EO
Provide ways to manage placebo effects	IIb	C-EO
<i>Overall recommendation*</i>	IIb	B-R, C-EO

Note. An extended version of this table, including references and do's and don'ts for each recommendation, is included in the Supplementary Materials.

** Legend: 1) Classes of Evidence:*

Class I: Evidence from well-designed randomized controlled trials (RCTs)

Class II: Evidence from well-designed clinical trials without randomization or cohort studies

Class III: Evidence from non-experimental studies or expert opinion

2) Levels of Evidence:

A: High-quality evidence from multiple RCTs or systematic reviews

B: Moderate-quality evidence from RCTs or clinical trials with limitations

C: Low-quality evidence from non-randomized studies or expert opinion

D: Expert opinion or anecdotal evidence

*** With 'overall recommendation', the quality of evidence for the overall domain (e.g., 'optimizing the clinician-patient relationship' or 'asking about expectations') is rated. Grading is adjusted from the American College of Cardiology and American Heart Association "Class of Recommendations and Level (Quality) of Evidence Ratings" system¹⁰ (see Supplementary Materials).*

i) Optimizing the clinician-patient relationship

People who are confronted with illness not only need high-quality personalized information, but also a supportive healthcare provider. When clinicians are empathic and warm, this can lead to numerous benefits, including lower anxiety, better adherence and increased recall of medical information¹¹⁻¹³. It is therefore strongly recommended to invest in the patient-provider relationship. As part of the NURSE framework¹⁴⁻¹⁶ (Table 2), various empathic behaviours are recommended, as they can be powerful in fostering a good patient-provider relationship. It is important that a clinician shows interest, so that patients do not feel regarded as just a 'case'. Even minor behaviours, such as asking how things are going at home or noticing something new in their appearance, can help patients feel seen¹¹. Patient satisfaction and trust are closely tied to the perceived level of listening and personal attention from healthcare providers^{11,17}. The continuity of care should be emphasized to reduce fears of abandonment, for instance with brief reassuring statements¹⁸. Equally essential is the establishment of a cooperative clinician-patient relationship grounded in shared goals to foster trust, facilitate open communication, and enhance quality of care¹⁹. Emotional support and addressing emotional needs of patients are significant but often overlooked aspects of care¹⁷. On the long-term, the impact of emotional support on trust may surpass other supportive actions, such as information provision²⁰. Nonverbal behaviours, including attentive listening, maintaining eye contact, and adopting a patient-centered posture are recommended as these can enhance trust^{11,21,22}. Patients moreover value clinicians who appear to take time for them; when they sit rather than stand, consultations are perceived as 50% longer and more positive, despite equal duration²³. Finally, it is essential to adapt the communication style to the

patient, depending on their values and expectations, as well as their understanding and informational needs^{24,25}. Importantly, clinicians should consider whether it is always helpful to give advice, for instance when patients are highly distressed²⁶.

Table 2: NURSE framework for empathic behaviours, developed on the basis of the studies by Pollak et al.¹⁵, Back et al.¹⁴ and van Vliet & Epstein¹⁶. Note that the examples serve only as an inspiration^{13,27}.

NURSE	Explanation	Example
Naming	Name the emotion	<i>"You sound very anxious."</i>
Understanding	Show understanding of the emotion	<i>"I really understand that you're dreading it." "I can imagine that you're upset about it."</i>
Respecting	Show respect for the patient	<i>"I'm impressed with how well you're handling the situation."</i>
Supporting	Express support	<i>"Whatever happens, we'll take good care of you. You don't have to face this alone."</i>
Exploring	Explore the emotion	<i>"What are you thinking about now? You're suddenly having to face so much."</i>

ii) Asking about expectations

Patients can have beliefs about the causes of medical conditions that may contrast medical models²⁸. Investigating their general attitude towards treatment, especially pharmacology, is crucial as it can provide valuable insights into patient expectations²⁹⁻³¹. Open-ended questions can be integrated naturally in the consultation (Table 3). When timed correctly, such questions can encourage patients to articulate their expectations, facilitating a more comprehensive understanding of their perspective and meaningful discussion. Unrealistic expectations should be managed, for instance by clarifying treatment limitations, or by emphasizing positive long-term effects²⁹. This aligns with the broader literature, suggesting that managing patient expectations is essential for treatment adherence and satisfaction^{32,33}. Negative expectations shaped by previous treatment disappointments should be identified. Differences between the previous unhelpful treatment and the currently offered treatment could then be explained^{29,34}. When possible, alternative treatment routes can be offered to enhance treatment satisfaction. Such a nuanced approach underscores the value of personalized care in effectively managing patient expectations and fostering a collaborative patient-provider partnership for improved healthcare outcomes.

Table 3. Example questions and phrases to incorporate expectation assessments into a consultation.

Aim of the question	Example	Possible Follow-up
Identifying general expectations towards treatment	<i>"Today I'm going to tell you more about treatment X, but first I would like to ask if you already know anything or have found information about this treatment..."</i>	Address anxiety or misinformation; adjust unrealistic information and align them with evidence-based care; address potential misunderstandings
	<i>"Can you tell me whether you already know something about the treatment?"</i>	
	<i>"Have you ever read or heard anything about the treatment?"</i>	
Check for presence of unrealistic (i.e., overly optimistic or pessimistic) expectations	<i>"What do you think is going to happen when you start with the treatment?"</i>	Challenge and adjust unrealistic expectation, e.g., offer evidence-based treatment information, using interpersonal communication, trust and emotional support
Check history with (similar) treatment(s)	<i>"Have you already had a similar treatment in the past?"</i>	(if relevant) emphasize difference with similar treatment; provide alternate route of administration
	<i>"How would you describe your earlier experience with treatment X?"</i>	

iii) Discussing the treatment rationale

Clinicians often describe treatment outcomes as more optimistic than they themselves expect³⁵. There are risks associated with this, as it could lead to disappointments^{36,37}. On the other hand, describing treatment outcomes more negatively can reduce placebo effects. It is crucial that healthcare providers neither exaggerate nor underestimate treatment results^{29,36}. Such a balanced approach is key to managing expectations and promoting realistic optimism about treatment outcomes. The importance of providing accurate and up-to-date information about the treatment and its expected effects, based on clinical guidelines, scientific literature and professional experience^{6,38}, cannot be understated. Providing a clear rationale for the treatment and linking this to the expected positive outcomes can help to enhance patient understanding^{36,39}. Such a rationale ideally encompasses a clear reason for the treatment, and an explanation of its mechanisms of action. This can reduce anxiety and promote confidence in the treatment^{29,38,40}. Positive and gain-framed messages could improve patients' willingness to receive a treatment and its perceived effectiveness⁴¹. Patients moreover value truthful but caring explanations that foster trust and optimism⁴². It is crucial to ensure that patients fully comprehend the



information to manage the expectations that the information inherently creates⁴³⁻⁴⁵. By proactively verifying that patients understand their condition, the treatment plan and any potential outcomes, biases and misunderstandings can be prevented^{6,29,44}. Finally, clinicians should refer to reliable sources for further information, such as professional websites, as this can prevent patients from acquiring untrue information or misattributing adverse symptoms to treatment, potentially activating nocebo effects⁴⁶.

iv) Discussing risks and side effects

It is difficult to give patients the information they need to make informed decisions, without causing discomfort or increasing the risks of nocebo effects. People tend to process negative information more thoroughly, which can cause overestimation of risks, particularly when emotions are involved^{47,48}. It is therefore crucial to balance informing about side effects with positive information, for instance about health benefits or future health improvement⁴⁶, to minimize nocebo effects^{49,50}. It is crucial to investigate whether there is an increased risk of nocebo effects, for instance, due to past experiences, negative beliefs or misunderstandings. These factors should be discussed to manage nocebo effects⁴⁶.

Some studies suggest that changing patients' mindsets about side effects can reduce nocebo effects: for instance, by reinterpreting side effects of vaccines or immune desensitization as 'positive signals that the treatment is working'^{51,52}. It may also be beneficial to list side effects briefly rather than focusing on one potential symptom⁵³, as long as these are low-risk side effects that do not pose serious risks⁵⁴. Because humans tend to focus on negative information, it is recommended to use positive framing (e.g., "*9 out of 10 do not experience side effects*")^{46,55}. Risks may also be communicated in statistical terms (e.g., "*0.1%*" instead of "*1 in 1,000 people*"), without using descriptors such as "(very) common", since this appeals to the imagination less^{56,57}. Finally, medical procedures should preferably be explained omitting negative phrases (e.g., descriptors of pain or discomfort), focusing on the purpose and advantages of the procedure, to avoid inducing nocebo effects^{29,58}.

v) Explaining placebo and nocebo effects

Experts worldwide agree that placebo and nocebo effects should be explained to patients⁵. Placebo effects can still occur when they are explained, and placebos are prescribed openly⁵⁹. Because placebo effects are part of all treatments², this principle can also be applied in regular treatment, where discussing the mechanisms of placebo effects can help empower patients to maximize these effects themselves. Conversely, explaining the mechanisms of nocebo effects could help patients to minimize these effects^{5,60-62}.

It may not always be necessary to delve into the specifics of the placebo effect, however, if a healthcare provider deems it beneficial, first elucidating the treatment mechanisms of action can facilitate the patient's understanding of placebo effects and the intertwined roles of the mind and brain in treatment efficacy⁴⁰. Placebo and nocebo effects need to be explained articulately⁵, particularly when communicating with patients who face challenges understanding health-related information. It is crucial that the reason for explaining is emphasized, especially for nocebo effect. For instance, it could be mentioned that the knowledge can reduce the occurrence of side effects^{5,46,63}. Any explanation should always be carefully tailored to the individual⁶⁴. Depending on their health skills, alternative terms of phrasing could be more appropriate (Table 4).

Table 4. Example phrasing and explanations for placebo and nocebo effects.

Topic	Recommended practice	Examples
Prerequisites	Carefully consider the need for informing about placebo and nocebo effects; only do so when you think this will be beneficial for the patient	Examples include instances when explaining helps patients understand the intricate interplay between the treatment's mechanisms of action and its context, or when patients believe themselves to be generally sensitive towards medicines or have had bad experiences. An explanation could be " <i>When we use treatments, they can help trigger the release of natural substances in your body that support healing. These processes usually function well, but during a temporary illness or a longer-term condition, they can be disrupted. The medication you take works alongside these natural mechanisms to enhance their effectiveness. It's important to remember that medications work in conjunction with your body's natural processes, and your previous experiences with treatment, as well as your expectations about the outcome, can influence how these elements interact. Together, they can contribute to achieving the best possible results in your treatment.</i> "
	Consider graphical information to support the explanation of placebo and nocebo effects	Using drawings to disentangle individual components of the total treatment effect on the improvement of symptoms: specific drug effects, placebo responses and placebo effects
Terminology	Use 'placebo effect' judiciously, as alternative terms or phrasing may be more appropriate	Alternate phrasing for placebo effects include: ' <i>expectation effects</i> ', ' <i>beliefs</i> ', ' <i>optimism</i> ', ' <i>positive mindset</i> ', ' <i>trust</i> '
	Use 'nocebo effect' only if the term is explained carefully. Alternative terms may be more appropriate, as may avoidance of the term itself altogether.	Alternate phrasing for nocebo effects include effects of anxiety, fear, or: " <i>Negative effects, such as side effects, that can arise or be strengthened if you pay close attention to whether they are occurring or not.</i> "



Mechanisms of action	Explain that placebo effects arise due to positive expectations about a treatment and are responses of the body that can alleviate symptoms	Emphasize, for instance, the effects of learning on expectations: <i>"If one of your family members had this treatment once, and you heard or saw what it did for her, this will unconsciously influence your expectation for this treatment"</i> , or <i>"If you have confidence in the treatment, this can strengthen its effects."</i>
	Emphasize that placebo effects are still efficacious when patients are informed about these effects	For example: <i>"Studies show that when patients are informed about placebo effects, these effects still occur. This is because they are automatic responses to the treatment and the context in which treatment occurs"</i> .
	Keep the explanations of placebo and especially nocebo effects simple	<i>"You could become more alert to symptoms if you have read the entire contents of an information leaflet, because I (the clinician) have put more attention on them"</i> .
Considerations	Avoid placing responsibility for the treatment's success or failure on the patient. Avoid people being made to feel personally responsible for having side effects	<i>"A positive mindset may be helpful but is not necessary"</i> ⁶⁵ or pointing out the automatic nature of conditioning. In case of nocebo effects: <i>"the body learns to react to a certain situation with a symptom, and this happens completely automatic. This is like rejecting a specific food taste when you have had food poisoning"</i>
	Prevent overly optimistic expectations of placebo effects	For example: <i>"in the treatment of serious illnesses, they can alleviate pain symptoms but cannot cure the illness"</i> ⁵ .
	Emphasize that not all side effects are nocebo effects, and that these need to be discussed with the clinician irrespective of whether side effects could plausibly be nocebo effects.	Highlight, for instance, that if side effects do occur, you will decide together what the next follow-up steps will be, and which treatment options there are.
Managing effects	Provide the patient with strategies to manage nocebo effects. These could include ways of coping such as relaxation or finding distractions	For example: <i>"If you are afraid of having side effects and are very alert to them, then side effects can increase; these are nocebo effects. There are multiple methods that can help you to reduce nocebo effects and to be more relaxed; for example, by doing relaxation exercises, or finding a distraction to keep your attention away from the side effects."</i> ²⁹

Discussion

In this paper, we present recommendations for healthcare professionals aimed at optimizing patient outcomes by strategically modulating patient expectations, which are recognized as robust determinants of both placebo and nocebo effects. This optimization is primarily achieved through contextual factors, specifically the relational and communicative aspects inherent to clinician-patient interactions. The recommendations

are intended for all healthcare professionals and aim to provide guidance for leveraging communication to enhance placebo effects and mitigate nocebo effects. They were developed by the PANACEA consortium and are grounded in previous existing materials and recommendations, supplemented by a systematic review of the empirical evidence in patient groups. Resultantly, several domains were identified to optimize communication behaviours during clinician-patient interactions. The results of the systematic review showed that, while most of these recommendations were backed by empirical evidence, others remain relatively under-studied. Here, we discuss the combined results of each of the identified domains.

The empirical evidence amply supports the recommendations for optimizing clinician-patient interactions and the clinician-patient relationship. These findings not only validate the importance of personal interest, reassurance, fostering trust, empathic communication, and non-verbal cues in patient care but also offer practical strategies for clinicians to improve the quality of their interactions with patients. Future research could further explore the long-term effects of these recommendations on patient outcomes and overall healthcare satisfaction.

A structured approach to identify and adjust patient expectations could significantly enhance patient satisfaction, treatment adherence, and overall treatment outcomes^{6,29}. Note that while the evidence collected in the systematic review underlines the general importance of asking about expectations, the identified studies did not specifically investigate the recommended strategies, such as adjusting unrealistic expectations. Personalized and empathic communication adapted to the individual situation is key to establishing trust and maintaining a positive patient-physician relationship. Future research could focus on evaluating the long-term effects of such strategies on patient satisfaction, treatment adherence, and health outcomes to further refine and validate these recommendations.

The recommendations for discussing the treatment rationale and the expected positive outcomes are aligned with existing literature on effective patient-physician communication^{6,16,39,40}. Providing accurate and balanced information, explaining treatment mechanisms, defining outcomes concretely, and recommending existing information sources are key strategies to enhance patient confidence, manage expectations, and improve treatment outcomes. The crucial role of providing balanced treatment information should be highlighted, that is, combining negative information with positive messages to mitigate its impact on expectancy and patients' welfare. Future research could evaluate the effectiveness of optimized treatment rationale communication on patient satisfaction, treatment adherence, and health outcomes to further refine and validate these recommendations.



The recommendations for structuring the consultation around and formulating precise information about to-be-expected side effects are empirically supported and aligned with existing literature on reducing nocebo side effects^{2,6,46,55}. For emphasizing the positive consequences of side effects, caution is needed, as it may need to be limited to situations where side effects are a natural and to-be-expected consequence of the type of treatment (e.g., desensitization therapies⁵²). Likewise, caution is needed when discussing possible advantages of side effects, as in some cases, they could be a sign of drug toxicity or intolerance, and changes to the treatment regimen are needed instead⁶⁶. Healthcare professionals should always carefully weigh each strategy and its potential use for mitigating nocebo effects depending on the treatment, its context and the individual that is being treated. Moreover, investigating and managing nocebo effects, and using neutral terms when discussing risks and procedures are key strategies to manage expectations and reduce nocebo effects.

Carefully explaining the mechanisms and implications of placebo and nocebo effects, presenting this information in an appropriate manner, and providing ways to manage nocebo effects could be key strategies to enhance patient understanding and improve treatment outcomes^{5,6,46}. Note that, while experts agree that placebo and nocebo effects should be explained⁵, most empirical evidence is preliminary, or stemming from open-label placebo studies. Future research could focus on evaluating the effectiveness of these communication strategies in regular treatments. Moreover, the effects of education about nocebo effects are still under-studied but could prove a powerful tool to mitigate nocebo effects and increase treatment adherence in daily practice.

There are several strengths of this recommendation paper, including the diverse and international interdisciplinary team within the PANACEA consortium, and the first systematic review on this topic as a first step to develop formal clinical guidelines while simultaneously identifying existing knowledge gaps and research needs. The inclusion of both quantitative and qualitative studies allows for a comprehensive overview of the breadth and depth of knowledge across all available evidence levels. Overall, these recommendations follow the AGREE II criteria⁶⁷, including a clear description of the scope and purpose, and have considered stakeholders' involvement. We also strived into the rigor of development and clarity of presentation, while considered possible applicability and editorial independence.

Several limitations need to be addressed. The search strategy was centered around communication and placebo and nocebo effects specifically. As a result, other papers, for instance, in the field of communication but not focused on placebo or nocebo, were not included. These can also provide effective strategies to optimize patient-provider

relationships and other communication aspects^{e.g.68,69} but are beyond the scope of the current work. In addition, other strategies to leverage placebo effects or mitigate nocebo effects, including counterconditioning^{70,71} or demonstrating the presence of nocebo effects by personalized N=1 trials⁷⁰⁻⁷², were not included.

Some of the strategies are expert-recommended but have only been studied preliminary in patient groups, such as the strategies to explain placebo and nocebo effects. The first studies in patient groups are promising^{60,73-75}, yet support mostly still stems from research with healthy volunteers (for review⁶²). Similarly, other communication strategies are frequently studied in healthy participants, such as framing (for review⁵⁵). Finally, the recommendations offer general advice and tools to communicate. It is crucial to carefully weigh the (dis)advantages of employing a strategy for each individual patient. Communication is always person- and context-dependent, and one strategy may not be equally beneficial for everyone.

CONCLUSIONS

In summary, this paper discusses multiple recommendations for optimizing placebo effects and managing nocebo effects to improve treatment efficacy in clinical practice. By leveraging effective communication strategies during patient-provider interactions, we aim to ensure that healthcare professionals are well-equipped to shape patient expectations and enhance therapeutic outcomes. As we move forward, these recommendations will be integrated into training programs designed to spread this knowledge as widely as possible within daily clinical practice



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SUPPLEMENTARY FILES.

Belonging to

"Managing Placebo and Nocebo Effects during Patient-provider interactions: A Consensus Statement and Clinical Recommendations for Healthcare Professionals"

Methods

First Stream (prior work synthesis & expert consensus)

Evidence for the first stream stems from expert consensus within the PANACEA Consortium, existing guidance and insights from prior narrative and systematic reviews^{e.g.-1²}, consensus papers^{13,14}, and a previous review of the literature, that informed the development of an existing eLearning module that reviewed literature until 2021¹⁵.

Second stream (systematic review of empirical evidence)

Eligibility criteria

Studies were included if they concerned placebo or nocebo effects in the context of medical communication and patient-provider interaction, were available full-text, and were published in peer-reviewed journals. To illustrate, studies assessing strategies to enhance placebo effects or minimize nocebo effects in patients during medical communication were included, whereas RCTs with placebo as comparator or experimental studies in healthy volunteers were excluded. Studies on open-label placebo were also included, as they concern communicating about placebo effects. The search was not limited to specific health conditions or methodologies.

Search strategy

The search strategy was developed based on the PRISMA guidelines¹⁶ in collaboration with an information specialist. It was piloted to ensure its validity and to ensure that a set of core articles were included in the search results. We strategically searched the following five databases from January 1, 2000, until May 15, 2024: PubMed, Embase (Ovid version), Web of Science, Cochrane Library, and PsycINFO (ABSCO host). The search strategy comprised a combination of keywords and controlled vocabulary terms relevant to clinician-patient communication and placebo and nocebo effects in patient populations (Appendix A in this document). The search strings were tailored to each database's syntax and subject headings.

Selection of evidence

After extracting all records and removing duplicates, one reviewer screened the records' titles, abstracts, and keywords. Next, full texts were evaluated against the in- and exclusion criteria. Doubts about study eligibility were resolved in discussion with the research group. Data from the included articles were extracted by 2 reviewers using a piloted form. The Quality of evidence was assessed based on the American College of Cardiology and

American Heart Association "Class of Recommendations and Level (Quality) of Evidence Ratings" system¹⁷ (Appendix B).

Results

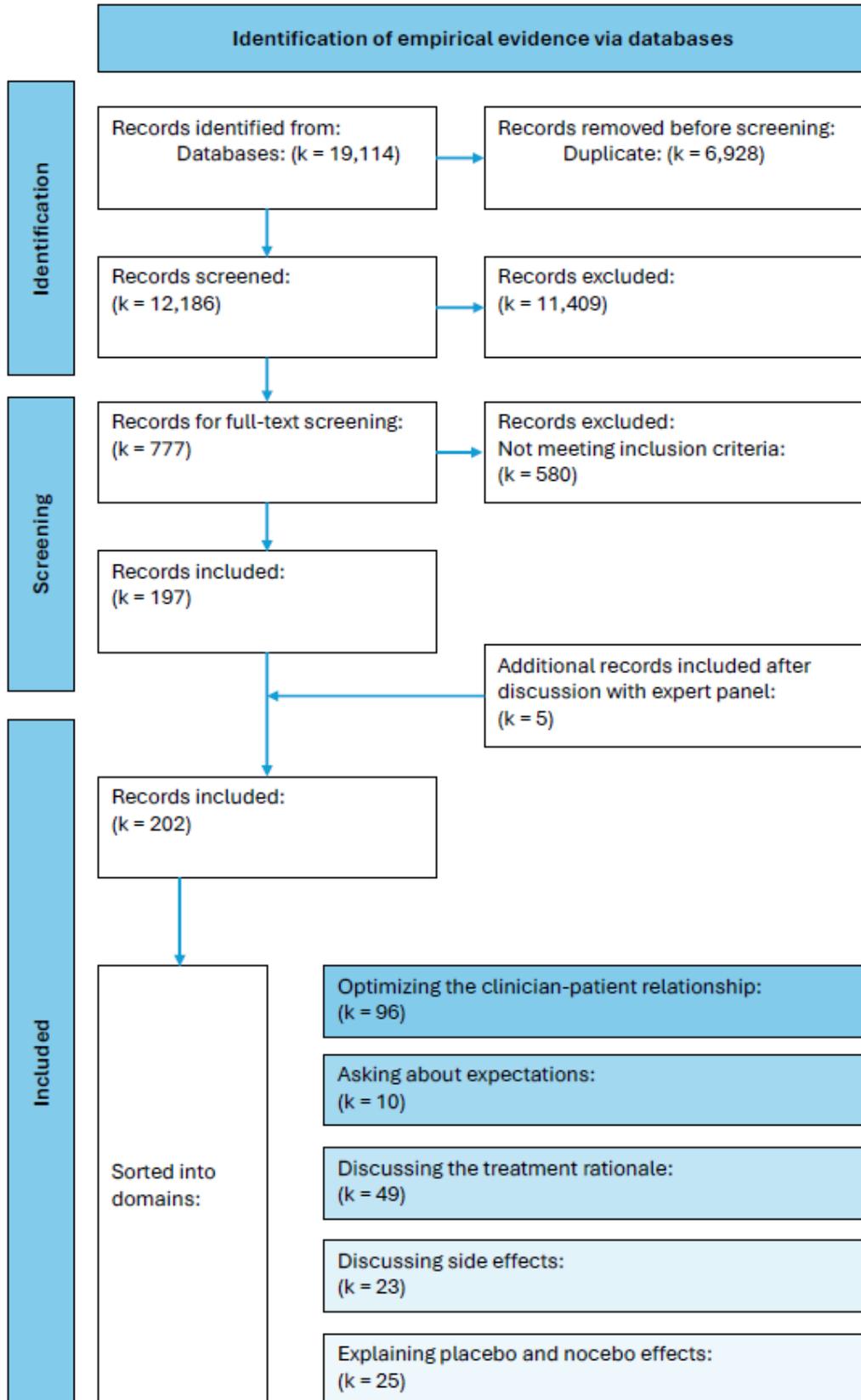
An overview of the literature backing each of the clinical recommendations from each information stream is provided in [Supplementary Table S1 \(Appendix C\)](#). For each domain, an information box is provided to outline the relevant theoretical background.

First stream (prior work synthesis & expert consensus)

Evidence for the first stream stems from expert consensus within the PANACEA Consortium, existing guidance and insights from prior narrative and systematic reviews^{e.g.1-12}, consensus papers^{13,14}, and a previous review of the literature, that informed the development of an existing eLearning module that reviewed literature until 2021¹⁵.

Systematic review (systematic review of empirical evidence)

We extracted 19,114 records from the databases. Duplicates were identified using Rayyan¹⁸, resulting in 12,186 unique records. After screening the titles and abstracts, 197 articles were included in the guidelines. Five studies were additionally included after discussion with an expert panel. A flowchart is provided in [Supplementary Figure S1](#). For an extensive summary of each individual study's findings and subsequently extracted recommendations for clinical practice, see [Supplementary Table S2 \(Appendix C\)](#).



Supplementary Figure 1. Flow chart for the inclusion of second-stream evidence



i) Optimizing the clinician-patient relationship

Systematic search: evidence in patient groups

Ninety-six studies relating to the patient-provider relationship were included in the systematic review (total $n = 133.608$). Most studies concerned cross-sectional survey, observational or similar design types ($k = 57$), followed by (semi-structured) interview and focus group studies ($k = 15$). Eight randomized controlled trials/studies were included, as were four experiments. Other design types included vignette ($k = 4$), mixed methods ($k = 2$), prospective or observational cohort ($k = 2$), and other studies ($k = 4$).

Regarding sample descriptors, most did not disclose specific information or included patients with various diagnoses ($k = 21$). Some samples were described using general terms (i.e., inpatients $k = 6$, outpatients $k = 5$, patients seeking GP or family doctor care $k = 10$, scheduled for radiotherapy $k = 2$, elective knee surgery $k = 1$, other specialties $k = 9$). When specific diagnoses were described, these mostly concerned patients with cancer ($k = 21$), diabetes ($k = 4$), HIV ($k = 4$), (chronic) pain ($k = 3$) or other diagnoses, including irritable bowel syndrome, tuberculosis, hypertension and pulmonary conditions ($k = 7$).

Info box i. Background

When people are confronted with a potential illness, they need high-quality information that is understandable, personalized, and backed by scientific evidence and clinical expertise, coupled with emotional support^{19,20}. Studies demonstrate that such an approach can result in placebo effects²⁰. Investing in the patient-provider relationship can help achieve this – among other, it is important that clinicians are empathic and warm, as this has positive effects on health outcomes¹⁹⁻²¹. When a clinician is empathic, patients are less anxious and insecure, and more satisfied^{19,20}, they can recall more medical information^{22,23}, and clinical outcomes can be improved through better treatment adherence^{20,24}. Additionally, empathy is important for physicians themselves: when physicians are more empathic, they experience greater job satisfaction²⁵. Thus, empathy is important, not only because patients regard it as such, but also through improvements of clinical outcomes and healthcare professionals' job satisfaction. As part of the NURSE framework for empathic behaviours^{12,26,27} (Supplementary Table S3, Appendix C), both verbal and non-verbal behaviours are recommended, as they can be powerful in fostering empathy, can contribute to a good clinician-patient relationship and can improve health outcomes.

Clinical recommendations

Interest in the Patient

Patients value a physician's interest in them as a person, preferring not to be regarded as merely a "body," "number," or "case." Even minor behaviours, such as inquiring about their



home life or noting changes in their appearance, can significantly contribute to patients feeling "seen" and valued¹⁹. This perspective, rooted in expert recommendations based on prior reviews and consensus papers, highlights that patient satisfaction and trust are closely tied to the perceived level of listening and personal attention from healthcare providers^{19,28,29}, and is further substantiated by systematic evidence from patient studies. These studies indicate that clinicians' attentiveness and engagement—through active listening and genuine interpersonal connection—are fundamental in enhancing the clinician-patient relationship³⁰⁻³². While some research reports limited effects of patient-centered communication, suggesting nuances in its impact³³, it is clear that misjudging patients' unmet needs can reduce perceived empathy and satisfaction^{34,35}. Overall, respectful and caring behaviours, including honesty, respect, and demonstrated competence, are consistently found to be central to building patient trust and fostering positive interactions³⁶⁻⁴⁶.

Give Reassurance about Ongoing Care

Patients who are confronted with serious illness frequently experience fears, for instance about abandonment, the progression of their illness, or its incurability. Brief reassuring statements from clinicians, such as "we're going to take good care of you", can reduce insecurity and fear, while enhancing memory recall after consultation by up to 8%⁴⁷. This finding underscores the therapeutic significance of empathy and reassurance in clinical encounters. Equally essential is the establishment of a cooperative clinician-patient relationship grounded in shared goals. Such collaboration fosters trust, facilitates open communication, and ultimately enhances the quality of care⁴⁸. These practices, rooted in expert recommendations, are supported by empirical evidence in patient groups, as evidenced from the systematic review: patients list continuity of care as one of the key actions to take to improve trust in the clinician, and a priority, alongside with being treated as a partner and person⁴⁹⁻⁵¹. One study finds that blending interpersonal and digital interventions could facilitate the availability of healthcare to patients and further reassure patients that continuous care will be provided⁵².

Respond to Negative Emotions

Patients frequently express negative emotions, such as grief and anger. Clinicians don't always respond to these emotions well. When clinicians do respond to negative emotions, patients are more satisfied⁵³, while the consultation does not necessarily last longer⁵⁴. Concrete examples of how to respond with empathy to patients' negative emotions are described using the NURSE model in Supplementary Table S3. Emotional support remains a significant but often overlooked aspect of patient care. These strategies, recommended by experts, are consistent with previous work²⁸ and offer practical guidance for clinicians to effectively address patients' emotional needs. From the systematic review, there is ample evidence that , among other, paying attention to emotional needs and support,

addressing and responding to emotions, and empathic and affective communication are central to fostering trust and building rapport with patients across disciplines⁵⁵⁻⁶⁷. On long-term outcomes, the impact of emotional support on trust may even surpass other supportive actions, such as information provision⁵⁸. Nonverbal communication in response to emotions and clinician demeanours, such as kindness and approachability, were also found relevant to building trust⁶⁵.

Eye Contact and Physical Presence

Nonverbal behaviours can be used to optimize the clinician-patient relationship in a number of ways and experts agree that these should be leveraged in daily practice. For instance, increased eye contact enhances patient trust and information recall²¹. Patients value clinicians who appear to take time for them; when clinicians sit rather than stand, consultations are perceived as 50% longer and more positive, despite equal duration⁶⁸. Such findings highlight the therapeutic importance of non-verbal cues and physical presence. Clinicians' facial expressions and their demeanour moreover influence perceived trustworthiness and competence, shaping satisfaction and modulating placebo effects, including pain reduction⁶⁹⁻⁷¹. In addition, the presence of supportive individuals during invasive procedures can induce hypoalgesic effects and lower anxiety⁷², underscoring the broader value of emotional support and trusted presence in promoting patient comfort and well-being.

This perspective on the critical role of certain nonverbal behaviours in fostering trust and strengthening the clinician-patient relationship is backed by evidence stemming from the systematic review. Behaviours such as attentive listening, maintaining eye contact, and adopting a patient-centered posture enhance trust^{19,21,73}. However, excessive eye contact or expressions of uncertainty can also diminish trust^{74,75}, highlighting the need for natural body language and behaviours that are adapted to the situation at hand. Several studies confirmed that patients preferred clinicians who sit during consultations, especially when receiving bad news, and perceived these encounters as longer and more engaging^{68,76,77}. Facial expressions contributed to trust and help address patients' emotional needs⁷⁸. In contrast, clinician attire appeared to have little effect on trust or perceived empathy^{79,80}.

Adapt communication style to the patient

It is essential to adapt the communication style to the patient, depending on their values and expectations, as well as their understanding and information needs⁸¹⁻⁸⁷. Importantly, clinicians should consider whether it is always helpful to give advice, for instance when patients are highly distressed⁸⁸. These findings underline the need for personalized communication, and understanding of the patient as a person, to foster trust and optimize the clinician-patient relationship.

Patient-centered communication

Results from the systematic search highlight that patient-centered communication is highly valued by patients, and is regarded as a core component of empathy, enhancing decision making and the clinician–patient relationship⁸⁹⁻¹⁰⁰. It could also reduce information avoidance by fostering trust and improving health literacy of patients¹⁰¹. Enhanced clinician empathy is linked to greater care quality, treatment satisfaction, and stronger clinician–patient relations¹⁰²⁻¹⁰⁵. Overall, effective, patient-centered, and shared decision-making communication could improve self-management, adherence, and treatment outcomes¹⁰⁶⁻¹¹³.

Other behaviours

While not explicitly mentioned in earlier evidence (i.e., the first stream of evidence), our systematic review identified several studies that recommended other behaviours to leverage the clinician-patient relationship to foster placebo effects. Of note are: augmenting contextual factors¹¹⁴, shared decision making and providing clear explanations^{115,116}; addressing prior negative encounters¹¹⁷; optimizing affective¹¹⁸, positive¹¹⁹, effective or direct communication¹²⁰⁻¹²⁶; addressing language barriers¹²⁷; fostering a sense of control and autonomy¹²⁸; fostering trust in benevolence¹²⁹; and improving cultural understanding^{130,131}.

ii) Asking about expectations

Systematic search: evidence in patient groups

Ten studies relating to the prior expectations of patients (total n = 1.478.350) were included. Study types comprised randomized controlled trials (k = 3), observational (cross-sectional) studies or surveys (k = 3), structured interviews (k = 2), focus groups (k = 1), and a prospective longitudinal survey-based study (k = 1). Patients with diverse diagnoses (k = 1), iron deficiency anaemia (k = 1), terminal diagnoses (k = 1), chronic pain (k = 1), cancer (k = 1), hypertension (k = 1), acute respiratory tract symptoms (k = 1), and those scheduled for surgery (k = 1), vaccination (k = 1), or registered with general practitioner offices (k = 1) were included.

Info box ii. Background

Patients naturally bring their own expectations to the consultation, shaped, for instance, by their treatment history, or by experiences of their family members. Such factors can contribute to patient beliefs about a treatment, irrespective of whether the treatment itself is identical or shares similar characteristics of administration. Pre-existing beliefs play a significant role in shaping patients' expectations regarding the treatment process and outcomes¹³². When patients expect a treatment to help them, it generally results in better outcomes^{2,133,134}. On the contrary, when patients have little confidence in a treatment, this can result in the treatment being less effective¹³⁵.

Patients may have unrealistic expectations about a treatment, for example that symptoms will be cured immediately, or they may underestimate the length of the treatment. These beliefs could impede the treatment or result in nonadherence. If patients' expectations are not met, they may be disappointed and lose their motivation for a treatment, which increases the risk of treatment dropout^{136,137}. Detecting unrealistically positive expectations and adjusting them can help to prevent such outcomes. It is therefore crucial to ask about a patient's expectations before the start of a certain treatment or medication^{136,138}.

Clinical recommendations

Ask About and Identify General Expectations Towards Treatment

Incorporating questions about expectations in the consultation with a patient is crucial. Investigating the general attitude towards treatment, especially drug treatments, based on previous experiences can provide valuable insights into patient expectations^{2,3,139}. These questions can be integrated seamlessly just before a specific treatment part is explained. Several studies^{140,141} highlight the importance of incorporating questions about patient expectations into consultations. Open-ended questions about patients' prior knowledge and experiences with a treatment can provide valuable insights into their expectations and concerns. Such questions encourage patients to articulate their expectations and can



facilitate a more comprehensive understanding of their perspective. Timing these questions just before introducing a specific treatment, as suggested, can enhance patient engagement and facilitate more meaningful discussions.

The value of understanding patient expectations prior to treatment is built upon the vast literature and expert consensus that highlights the importance of expectations to treatment over several years^{e.g.,13,14}. Little empirical evidence exists for the immediate value of asking about expectations, however, the systematic review did underline the role of pre-existing beliefs in treatment. For instance, pre-vaccination expectations mediated the relation between different information sources (e.g., social media exposure) and vaccine side effects¹⁴². Observation of positive treatment effects in others affected disability reported in chronic pain, underlining the importance of these social processes and the potential to leverage them to enhance placebo effects¹⁴³. Focusing on expectations and addressing them before treatment moreover improved clinical outcomes in several conditions¹⁴⁴⁻¹⁴⁶. In palliative care, using hypothetical scenarios to guide patients in aligning their care plans with realistic outcomes could help informed decisions, including considering of alternate options¹⁴⁷.

Adjust Unrealistic Expectations

Next to asking patients about their general treatment expectations, it is strongly recommended to manage overly optimistic or pessimistic expectations². For instance, when patients express overly optimistic expectations, it is important to clarify the limitations of the treatment. Conversely, for those with overly negative expectations, focusing on potential positive long-term effects can help to recalibrate their outlook². This aligns with the broader literature, suggesting that managing patient expectations is essential for treatment adherence and satisfaction^{132,148}. Patients may also expect certain tests or treatment based on their complaints, that could or could not conflict with existing treatment protocols¹⁴¹. An open and honest attitude of healthcare professionals can help patients understand treatment suggestions and form realistic expectations¹⁴⁹.

From our systematic review, qualitative research highlights that patients can have varying beliefs about the causes of medical conditions, that may contrast medical models¹⁵⁰. These beliefs need to be taken seriously. To illustrate, a survey study conducted in a large group of UK GP patients (n = 1.476.252) demonstrated that taking patient problems seriously was the strongest predictor of trust and confidence in clinicians¹⁵¹. Patients moreover appear value proactive involvement of clinicians to manage expectations and reassure them in the face of diagnostic uncertainty¹⁵². Expectations are continuously shaped¹⁵³, and addressing them during treatment has the potential to leverage placebo effects.

Address Disappointment from Previous Treatments

Bingel² and Schemer et al.¹⁵⁴ suggest identifying negative expectations that may be due to previous treatments and their outcomes, to then emphasize the differences and potential effectiveness of the current treatment. For patients who have experienced disappointing treatment results multiple times, offering alternative routes of administration that resonate with the patient's preferences can play a significant role in enhancing treatment satisfaction and adherence. This nuanced approach underscores the value of personalized care in effectively managing patient expectations and fostering a collaborative patient-provider partnership for improved healthcare outcomes.



iii) Discussing the treatment rationale

Systematic search: evidence from patient groups

Forty-nine studies were included that concerned discussing the treatment rationale. These comprised $n = 10.032$ patients, $n = 14.549$ individuals from the general population or diverse patient groups, and $n = 1.230$ HCPs and $n = 95$ healthy volunteers (who were included alongside patients in several instances). Studies included randomized controlled trials ($k = 17$), observational studies ($k = 7$), cross-sectional surveys ($k = 8$), semi-structured interviews ($k = 5$), experiments ($k = 4$), multicentre or longitudinal surveys ($k = 2$), and a prospective observational ($k = 1$), a quasi-experimental ($k = 1$), a feasibility ($k = 1$) and an experimental vignette study ($k = 1$). One study comprised two parts: a cross-sectional survey followed by an experimental study. One study combined a survey with structured interviews.

Studies were conducted in diverse patient groups or the general population ($k = 8$), in oncology ($k = 8$), primary care ($k = 4$), gynaecology ($k = 3$), Parkinson's disease ($k = 3$), chronic pain conditions ($k = 3$), MDD ($k = 2$), medically unexplained symptoms ($k = 2$), or other (isolated) medical conditions ($k = 11$). Some studies included children ($k = 3$) or adults visiting a medical centre ($k = 2$).

Info box iii. Background

While the placebo effect is best known as a positive reaction to non-active treatment, the power of expectations perhaps applies even more for recognized, active treatments¹⁵⁵. In these instances, earlier experiences with a treatment can facilitate placebo effects based on conditioning^{6,156}. When someone expects that a treatment will help, this can significantly increase the treatment's effectiveness¹⁰ and alleviate fear and uncertainty¹⁵⁷. Vastly contributing to such expectations is the rationale that a clinician provides: for instance, the reasons for the treatment, its working mechanisms, and the to-be-expected benefits. A recent study found that most general practitioners regularly describe the expected treatment outcomes as more positive than they themselves expect, possibly to increase their effectiveness. However, a considerable number of practitioners presents the envisaged treatment results sometimes as more negative, possibly to avoid disappointment¹⁵⁸. There are risks associated with presenting treatments as more effective than they really are, because this can lead to disappointment and damage trust^{159,160}. On the other hand, presenting the treatments as less effective can impair their effectiveness and may confront patients with unnecessary fears. It is therefore important to balance information to create positive, but realistic expectations, that are grounded in the patient's individual situation^{159,161}.

Clinical recommendations

Provide Correct Information Based on Evidence-Based Knowledge

The importance of providing accurate and up-to-date information about a treatment and its expected effects cannot be understated¹⁶¹. It is recommended that this information is grounded in the most recent medical knowledge in the clinical guidelines and scientific literature, but also the clinicians' professional experience^{138,161}. This recommendation, derived from expert consensus and multiple resources in the placebo literature, aligns with the broader literature, suggesting that patient trust and confidence are closely tied to the credibility of the information provided by healthcare providers. It is also supported by evidence stemming from the systematic review. Multiple studies were identified that investigated the way information is provided during clinician-patient interaction, frequently focusing on the message's content and delivery method. Patients appear to prefer explanatory information and the discussion of biomedical issues over psychosocial issues or empathizing with their emotions¹⁶²⁻¹⁶⁴, except for when psychosocial problems are present¹⁶⁴. Sensitive topics are preferably approached indirectly¹⁶². Certain medical terms may also induce negative expectations: patients exposed to the term 'heart failure', for instance, believed the illness to have more serious consequences, and felt more anxious and depressed relative to those for whom a euphemism was used¹⁶⁵. This underlines the need for a sensitive approach in communicating, among other, medical diagnoses. Effective communication could balance optimism with a clear discussion of pessimistic aspects and being emotionally sensitive¹⁶⁶, while providing clear and comprehensive explanations can improve shared decision making¹⁶⁷. Taken together, these results underline the critical need for careful and credible explanations of treatments.

Avoid Exaggeration or Underestimation of Treatment Results

An accurate presentation of treatment outcomes is essential to maintain credibility and trust with patients. Clinicians should neither exaggerate nor underestimate treatment results^{2,159}. Such a balanced approach is crucial for managing expectations and promoting realistic optimism about treatment outcomes. This recommendation for balance is underlined by the empirical evidence, where it was shown that positive phrasing and optimistic communication can be effective for optimizing outcomes^{156,168-177}, but also that in some instances, interventions to boost placebo effects were counterproductive (i.e., in weight loss)¹⁷⁸. Such, at a first glance, contradictory study findings could be indicative of different approaches being necessary, depending on the specific medical condition and the treatment that is being recommended.

Empirical evidence further shows that, if the to-be-provided information is negative, indirect messages could be used to reduce anxiety¹⁷⁹. In case of uncertainty (e.g., of illness trajectory), patients prefer transparent communication^{180,181}. Implicit statements of



uncertainty (e.g., "*most likely diagnosis*") led to better outcomes compared with explicit statements of uncertainty, that negatively affected trust, perceived competence, and adherence¹⁸⁰. These findings underline the need for open discussion of treatment results, without exaggeration or underestimation of potential outcomes. Finally, clear communication, for instance about placebo rates in clinical trials, can improve evaluation of drug accuracy by patients, according to a single study¹⁸².

Clearly State Treatment Rationale and Expected Positive Outcomes in Concrete Terms

Providing a clear rationale for prescribing a treatment and explicitly stating the expected positive outcomes can enhance patient understanding and confidence in the treatment^{159,183}. Example phrases are: "*We know that the treatment I'm now prescribing for you can significantly reduce the pain for many patients*" or "*Medical research has shown that most patients experience a strong improvement in the symptoms shortly after taking it.*"

Supplying tangible details about the predicted outcomes, grounded in evidence-based knowledge, helps to align patients' expectations with realistic treatment outcomes, and fosters understanding and confidence in the treatment. This perspective is supported by the systematic review: patients value truthful but caring explanations that foster trust and optimism¹⁸⁴. Positive and gain-framed messages can improve the willingness to receive a treatment and elevate its perceived effectiveness^{185,186}, while positive affect-oriented speech can reduce anxiety and optimize self-efficacy¹⁸⁷. The way that numerical information is presented affects illness experience¹⁸⁸. Finally, discussion of biomedical information is highly valued, however, physicians usually do not meet expectations regarding biomedical details like symptom causes, severity, and test results¹⁸⁶. This finding underlines the importance of clinical recommendations on and training in communication about the treatment rationale.

Explain Treatment Mechanisms of Action

Kleinstäuber et al.¹⁶¹ suggest that explaining the mechanisms of action of the treatment (e.g. what it does in the patient's body) and explaining this clearly and vividly can increase patient confidence in its effectiveness. This recommendation is supported by the broader literature, suggesting that a better understanding of the treatment mechanism can optimize placebo effects and enhance treatment outcomes²⁸, and is backed by empirical evidence that shows that explaining procedural information can reduce anxiety¹⁸⁹, while explaining about the drug approval process can foster confidence in the safety and quality of generic medicines¹⁶¹. Lin et al.¹⁹⁰ show that considering a combination of the patient's (medical) history, physical examination, and prior research, in explaining the treatment is frequently indicated as important for patients. Taken together, these findings highlight the

importance of explaining the treatment mechanisms in detail to foster placebo effects in patients.

Aligning Patient Expectations and Beliefs with Evidence-Based Care

To create realistic optimistic expectations, that are grounded in the patient's individual situation, it is recommended to first ask what the patient personally expects from a treatment and try to align this with evidence-based care by addressing potential misunderstandings^{161,191}. This can help reassure patients and reduce fears¹⁰. This can also be done when no treatment is being prescribed, by expressing positive expectations about the course of the symptoms, for instance, that the symptoms will usually fade by themselves within a few days. Potentially negative expectations, that could arise because patients are referred to another clinician, could be mitigated, for instance, by explicitly expressing confidence in the new clinician¹⁶⁰.

Empirical evidence underlines that prioritizing and meeting patient expectations can enhance satisfaction and improve outcomes^{192,193}. Others found that, while raising positive and realistic expectations did not affect clinical outcomes, this practice did improve perceived care quality¹⁹⁴. These findings suggest that even when there are limits to the strength of placebo effects, positive effects may be sought elsewhere. Identified strategies to handle unrealistic beliefs include balancing information with either acceptance or denial of these beliefs and using medical expertise to address challenges around unrealistic beliefs¹⁹⁵. Negative beliefs, including beliefs about medication overuse and medication harm, can be addressed through interpersonal communication, emphasizing trust and emotional support¹⁹⁶. Taken together, various strategies are possible to align expectations with realistic outcomes, and it is prudent that the clinician identifies the best possible option given the patient's individual situation.

Emphasize the Purpose and Advantages of Treatment Procedures

For some treatment procedures, focusing on the purpose and advantages of the procedure rather than explicitly mentioning potential pain or discomfort can help to alleviate patient anxiety and enhance cooperation². For example: "*I'm going to numb the area with this, so that you will be comfortable during the procedure.*"¹⁹⁷. This recommendation is consistent with the broader literature, suggesting that framing information in a positive and reassuring manner can improve patient satisfaction and experience^{5,198}. Empirical evidence supports focusing on the potential benefits when explaining procedures, including case descriptions, storytelling, reassurance, clear explanations and favourable prognoses^{199,200}. Patients could moreover be more receptive to tailored communication strategies, that account for individual characteristics and emotional states^{200,201}. To improve satisfaction, informal talk may be best restricted to history-taking and not be done, for instance, during physical examinations²⁰².

Check Patient Comprehension

Ensuring that patients fully comprehend the information provided during medical consultations is essential for managing the expectation effects that this information inherently creates^{156,203,204}. By verifying that patients have a solid understanding of the information that is being provided, clinicians can address any misconceptions, elucidate complex concepts, and confirm the patient's grasp of their condition, treatment plan, and potential outcomes^{2,138,203}. Such a proactive approach allows for prompt corrections and adjustments, and ensures that patients are well-informed and actively engaged in decisions regarding their health²⁰³. While such practices are well supported through consensus and existing literature, the systematic review only identified three studies, whose findings suggest that checking patient comprehension and taking individual preferences into account may have beneficial effects²⁰⁵⁻²⁰⁷. Moreover, even when patients tend to prefer a specific communication type (i.e., qualitative probability expressions), there is a wide variability in information interpretation²⁰⁵, underlining the need for checking patient comprehension.

Suggest Reliable Information Sources

Providing patients with additional resources, including leaflets or reliable websites, could address patients' need for additional information and alleviate doubts, ultimately enhancing their understanding of the treatment and its expected outcomes. This recommendation is supported by the broader literature, suggesting that patient education and access to reliable information can improve treatment adherence and outcomes. For instance, studies on health literacy find that increasing patients' knowledge and understanding of treatments can improve treatment engagement, facilitate communication and shared decision making, and improve quality of life^{208,209}. Suggesting reliable sources could moreover prevent patients from acquiring untrue information or misattributing adverse symptoms to their treatment, potentially activating nocebo effects²¹⁰. Findings from the systematic review elucidate an interaction between patients' health information seeking behaviours and improvements in communication and the clinician-patient relationship²¹¹⁻²¹⁵. Taken together, these findings underline that clinicians should encourage patients to seek credible health information and provide them with reliable resources.



iv) Discussing risks and side effects

Systematic search: evidence in patient groups

Twenty-two studies were included. Fourteen were conducted in patient groups (n = 3.435), one in patients + healthy controls (n = 69), and three in the general population, including patients (n = 2.598). Study types included (pilot) randomized controlled or comparative trials (k = 12), cross-sectional or randomized surveys (k = 3), (semi-structured) interviews (k = 1), randomized prospective (longitudinal) studies (k = 2), experiments (k = 2), and case reports (k = 1). Studies were conducted in the general population or diverse patient groups (k = 5), oncology (k = 3), depressive or sleeping disorder (k = 3), cardiology (k = 2), or other singular conditions (k = 9).

Info box iv. Background

Research show that mentioning risks and side effects can elicit nocebo effects²¹⁰. Clinicians routinely explain risks and possible side effects of a treatment to their patients, but major restrictions to do this adequately include limited time²¹⁶. Only briefly mentioning certain serious side effects or complications can make a patient apprehensive or anxious and can result in the patient experiencing those symptoms more strongly⁷. People tend to focus on and process negative information more thoroughly, which makes it likely that they will overestimate risks²¹⁷. Nonexperts' perception of risk for an event hinges even more on irrational and emotional factors than evidence-based information²¹⁸. It is difficult to give patients the information to which they are entitled, and that they need for informed decisions, without causing unnecessary discomfort or increasing risks of experiencing nocebo effects. It is additionally important to realize that, if the patient experiences side effects or complications without being properly warned about them, this in turn could negatively reflect on the clinician-patient relationship.

Clinical recommendations

General structure of the consultation

Emphasize Positive Consequences of Side Effects

In some situations, changing patients' mindsets about side effects may reduce nocebo effects. By reinterpreting minor, non-harmful side effects as positive signals that the treatment may be working, patients may be able to deal with side effects better, reducing anxiety and the risk of nocebo effects²¹⁹. Evidence shows that such a strategy could be relevant when side effects are inherent to the treatment being given, as for vaccinations²²⁰ or allergy desensitization therapies²²¹.

Balance Side Effect Information with Positive Information

Petrie & Rief²¹⁰ recommend that information about side effects can be balanced by other, positive information, for instance about future health changes or treatment benefits: "the

side effects may be experienced as unpleasant, but the treatment can help you to feel better later". This approach aligns with the broader literature on information framing, suggesting that framing information positively can enhance patient confidence and treatment adherence. Other forms of balance may be sought in addressing anxiety before providing the information on side effects, or by improving support for patients should any effects occur²¹⁰. This perspective is also supported by empirical evidence, that shows that informing about side effects, while beneficial, can also lead to nocebo effects and misattribution^{222,223}, the risk of which can be minimized by balancing information about side effects with positive information²²⁴⁻²²⁶. Balanced information could moreover promote informed decision making²²⁷. Qualitative studies further highlight the need for balanced and transparent information on risks and benefits among patients²²⁸, although other evidence suggests limited effects^{229,230}. This could reflect that patients may have different information needs that the clinician should address. Finally, balancing negative information with other positive interventions, such as self-affirmation, may be helpful to mitigate nocebo effects^{231,232}.

Investigate Nocebo Effects

It is advisable to investigate whether a patient has a greater risk to develop nocebo effects, for instance due to past negative experiences, or the belief that they are generally highly susceptible to side effects²¹⁰. Leading questions or words should be avoided (e.g. "*Do you think the treatment will be difficult?*" or "*Do you think the side effects will be unpleasant?*") and instead an open formulation should be used (e.g. "*What do you expect about the side effects of this treatment?*"). It is crucial that the patient knows and feels that there are no wrong answers to establish an open and trusting clinician-patient relationship. Understanding the patient's perspective on side effects is crucial to manage nocebo effects²¹⁰. This notion is underlined by evidence stemming from the systematic review: patients can have difficulties with interpreting numerical information²³³, underscoring the need to investigate nocebo effects that could be caused by patients' understanding of medical information. Health anxiety and negative beliefs about medicines can further increased negative expectations about side effects²³⁴. Finally, delayed responses to negative information were found in patients²³⁵, suggesting that nocebo effects may also occur at different timeframes.

State All Symptoms Briefly to Reduce Nocebo Effects

Research has shown that people may remember a smaller number of symptoms better and more vividly, which could increase the risk that they will experience those symptoms via nocebo effects²³⁶. Based on these findings, it may be beneficial to name all symptoms briefly rather than focus on a few. However, this obviously applies primarily to low-risk side effects; it is essential to clearly communicate side effects that pose serious risks²³⁷. Empirical evidence shows that most patients prefer brief, targeted explanations of



adverse drug reactions, with details only being provided by the clinician upon request²³⁸. Other work shows no effects of detailed side effect briefings on number of side effects reported²²⁹. This underlines the need for further study of this topic.

Framing of Information

Frame Positive Outcomes Rather Than Negative Outcomes

Since humans are more prone to negative than positive information²¹⁷, it is recommended to frame information about side effects positively and focus on those who do not experience side effects, such as stating that 90% of patients do not experience complications, rather than saying that 10% do. This strategy aligns with recommendations provided in prior literature^{198,210} and is supported by our systematic review, in which several studies demonstrated effects of positive framing on negative expectations and side effects^{239,240}.

Discuss Risks in Statistical Terms

Planès et al.²⁴¹ suggest discussing risks in statistical terms (i.e., 0.1%), rather than referring to "1 in 1,000 people" (or 100, or 10,000), as this approach appeals more to the imagination and reduces the likelihood of anxiety. The second way of risk phrasing appeals more to the imagination, therefore potentially enhancing anxiety. Similar evidence was identified through the systematic review: Simplified and risk-neutral information may reduce nocebo side effects and anxiety²⁴². In serious illnesses, such as cancer, absolute mortality risk framing may be preferred over relative mortality risk framing, to reduce bias and ensure better informed decision making²⁴³. Using descriptors such as "very common" and "common" can also result in negative expectations for side effects, even when probabilities of these effects are low²³⁴. Taken together, there are various ways to phrase risks, and discussing them in statistical terms could be most helpful to mitigate nocebo effects.

Use Neutral Terms When Performing a Procedure

Varelmann et al.¹⁹⁷ show that the phrasing used to describe medical procedures (i.e., either positively, "*we will give you a local anaesthetic that will make you comfortable*", or negatively, "*you will feel a sting, this will be the worst part of the procedure*"¹⁹⁷) can affect pain experience following a local anaesthetic injection. Avoiding negative phrasing is therefore recommended to prevent inadvertently evoking nocebo effects.

v) Explain placebo and nocebo effects to patients

Systematic search results: evidence from studies in patient groups

Twenty-five studies were included. Most were randomized controlled (clinical) trials (k = 16) or online or laboratory-based experiments (k = 4). Five employed other study designs (e.g., interviews, mixed methods). Most studies (k = 18) were focused on open-label placebo effects, where placebo interventions were provided along with a rationale explaining their potentially beneficial effects. Five studies focused on education about nocebo effects, and two on education about placebo effects.

All studies included patients (n = 2.171). Specific medical conditions were pain (k = 5), allergic rhinitis (k = 4), oncology (k = 4), gastroenterology (k = 3), surgery (k = 2), ADHD (k = 3), (risk for) depressive disorders (k = 3), or primary insomnia (k = 1).

Info box v. Background

Recent evidence shows that placebo effects can occur when patients know about them^{244,245}, contradicting the traditional view that these effects only occur under unaware conditions. With "open-label placebos", patients are clearly and openly informed about how and why placebo effects work¹⁴. This principle can also be applied in regular treatment, because – as we saw earlier – placebo effects are part of all treatments¹³. Discussing the mechanisms of placebo effects could provide patients a feeling of control, or "empowerment": when they know how placebo effects work, they can take steps themselves to increase them. Importantly, feeling in control over a situation has health benefits²⁴⁶.

In contrast, explaining the mechanisms of nocebo effects could help to mitigate them^{11,14,247-249}. To illustrate, demonstrating that some side effects are attributable to nocebo effects has helped patients successfully restart statins treatment, that they had discontinued²⁵⁰. It is important to make careful judgements about whether explaining nocebo effects can help the patient. For instance, 1 in 5 people say that they are "very sensitive to medicines" or are afraid of side effects²⁵¹, which increases risks of side effects²¹⁰. In such a case, it could be useful to explain the mechanisms of nocebo effects.

Clinical recommendations

Careful Explanation of Placebo Effects

While it may not always be necessary to delve into the specifics of the placebo effect, if a healthcare provider deems it beneficial to explain this phenomenon for the patient's benefit, first elucidating the treatment mechanisms of action can facilitate the patient's understanding of placebo effects and the intertwined roles of the mind and brain in treatment efficacy⁸. It is essential to also communicate that treatments can still be efficacious even when patients are informed about placebo responses²⁴⁴.



An example of an explanation that highlights the symbiosis between the treatment's mechanisms of action and placebo effects could be as follows: *"When we use treatments, they can help trigger the release of natural substances in your body that support healing. These processes usually function well, but during a temporary illness or a longer-term condition, they can be disrupted. The medication you take works alongside these natural mechanisms to enhance their effectiveness. It's important to remember that medications work in conjunction with your body's natural processes, and your previous experiences with treatment, as well as your expectations about the outcome, can influence how these elements interact. Together, they can contribute to achieving the best possible results in your treatment."*

Charlesworth et al.²⁴⁴ and Evers et al.¹⁴ emphasize the importance of articulately explaining the concept of "placebo effects", stressing the need to use the term "placebo effects" judiciously, particularly when communicating with patients with limited health literacy skills. This perspective is underlined by the systematic review, wherein ample studies show that providing a placebo with a rationale on placebo effects is beneficial in multiple health conditions^{e.g.252-260}, even though others report no or limited effects²⁶¹⁻²⁶³. In hallmark studies, the rationale often concerns four topics: 1) the powerful placebo effect; 2) explaining classical conditioning; 3) positive attitudes can help but are not necessary, and 4) adherence is necessary²⁵³. Such explanations highlight potentially beneficial effects and simultaneously avoid placing disproportionate accountability for the outcomes with the patient (i.e., by emphasizing that *"a positive attitude helps but is not necessary"*).

Mechanism of Action of Placebo Effects

It is recommended to emphasize that placebo effects arise due to positive expectations about a treatment and that they are real responses of the body that can alleviate symptoms¹⁴. Providing examples of how learning affects expectations, can help patients understand the mechanisms of placebo effects²⁶⁴. For example: *"If one of your family members had this treatment once, and you heard or saw what it did for her, this will unconsciously influence your expectation for this treatment."* Several studies were identified that demonstrate that such an explanation could be helpful²⁶⁵⁻²⁶⁸.

Appropriate Presentation of Information

Any explanation given should always be carefully tailored to the individual²⁶⁴. It is possible to use additional terminology or euphemisms to explain the mechanisms of placebo effects, such as the effect of confidence and the clinician-patient relationship, as recommended by experts in a Delphi study¹⁴. For example: *"If you have confidence in the treatment, this can strengthen the effects of the treatment."* It is not necessary to use the



terms 'placebo effects' or 'nocebo effects' since patient providers and patients might prefer other terminology or phrasing¹⁴.

There is little empirical evidence yet for how information about placebo effects can be appropriately presented, but some evidence suggests that patients who already have more favourable attitudes towards placebo are more receptive to such information²⁶⁹. Other studies find that using standardized educational tools on placebo effects could be helpful to increase understanding of these effects, although one of these trials found that the information mitigated placebo effects^{270,271}. This indicates that the direction of the effects could depend on the type of information that is provided as well as the medium (i.e., via a website instead of personally by healthcare provider, who is able to adjust the appropriateness of the information).

Prevent Overly Optimistic Expectations

While not explicitly backed by empirical evidence yet, it is crucial to prevent *overly* optimistic expectations of placebo effects. Experts emphasize the importance of preventing overly optimistic expectations by clarifying that placebo effects can alleviate symptoms but cannot cure them, particularly in the treatment of serious illnesses. For example: in the treatment of serious illnesses, they can alleviate pain symptoms but cannot cure the illness¹⁴.

Attending to the Purpose of Explaining Nocebo Effects

When explaining the nocebo effect, it can be important to emphasize to the patient that this is explained because the knowledge can reduce the occurrence of side effects^{14,210,272}. For example: if the patient had negative experiences with a treatment in the past, then knowledge about nocebo effects can help to mitigate these earlier experiences. Experts on placebo effects have placed stipulations on informing about particularly nocebo effects; they caution that the need for informing about nocebo effects should be carefully considered¹⁴.

Careful Explanation of Nocebo Effects

Evers et al.¹⁴ recommend that if the term "nocebo effects" is used, it should be explained carefully. For instance, these effects can be described as negative effects, such as side effects, that can arise or be strengthened if you pay close attention to whether they are occurring or not. Petrie and Rief²¹⁰ also recommend providing a lay definition of the concept. Depending on the specific patient and their health literacy skills, using alternative or related terms may be more appropriate, including anxiety or fear. This perspective is backed by empirical evidence examining the effects of educating patients about nocebo effects^{247,273-275}, finding that this could have beneficial effects, for instance on side effects and adverse event reporting^{247,274}. Some studies demonstrated limited effects^{247,274}, again highlighting the need for an individualized approach, in which the benefits of informing

about nocebo effects are carefully weighed against potential adverse effects. Identified beneficial effects included improved side-effect expectations and coping expectations²⁷³, and in one study, patients who had received it preferred general information over information about all potential side effects²⁷⁵.

Keep Explanation Simple

The explanation of nocebo effects should preferably be kept short and simple¹⁴; for example, that the patient can become *"more alert to symptoms if they have read the entire contents of an information leaflet, because the clinician has put more attention on them"*.

Avoid Unintentional Negative Effects

When explaining the mechanisms of nocebo effects, it is important to do this in such a way that no iatrogenic or unintentional negative effects are generated, for example the feeling of being personally responsible for treatment success or side effects¹⁴. It is crucial that the healthcare provider ensures that the patient understands that not all side effects are nocebo effects: sometimes they are real side effects, and it is important to report them to the clinician. In case side effects are experienced, it is important that the healthcare provider and the patient together discuss and decide about the best possible follow-up steps and possible treatments options, irrespective of whether side effects could plausibly be nocebo effects.

Provide Ways to Manage Nocebo Effects

Additional suggestions to deal with nocebo effects could be offered, for instance, asking a family member to read the information leaflet instead of the patients themselves. If the patient then has any symptoms, the family member can be asked whether these are included in the side effects. Bingel² highlights that strategies, such as relaxation or finding distractions to keep the attention away from side effects, could be combined with explaining nocebo effects (mechanisms). For example: *"If you are afraid of having side effects and are very alert to them, then side effects can increase; these are nocebo effects. There are multiple methods that can help you to reduce nocebo effects; for example, by doing relaxation exercises, or finding a distraction to keep your attention away from the side effects"*. In the systematic review, one study was identified that underlines this recommendation: Von Blanckenburg et al.²⁷⁶ describe two cases where psychoeducation on nocebo effects is provided and patients are taught strategies to deal with side effects of cancer treatment. These cases provide preliminary evidence that this approach could reduce harmful side effect expectations and empower patients.

Appendix A. Search strings for the systematic review

Automatic search string translation:

<https://sr-accelerator.com/#/polyglot>

Databases:

PubMed

<http://www.ncbi.nlm.nih.gov/pubmed?otool=leiden>

11-part strategy (excluding animal only studies, time limit: 2000-.): 4751 references, including 22 of the 22 known items

((("Placebo Effect"[majr] OR "Placebo Effect"[tiab] OR "Placebo Effects"[tiab] OR "Placebo Effect*" [tiab] OR "Placebo Response"[tiab] OR "Placebo Responses"[tiab] OR "Placebo Respon*" [tiab] OR "Nocebo Effect"[majr] OR "Nocebo Effect"[tiab] OR "Nocebo Effects"[tiab] OR "Nocebo Effect*" [tiab] OR "Nocebo Response"[tiab] OR "Nocebo Responses"[tiab] OR "Nocebo Respon*" [tiab] OR "nocebo"[tiab] OR "nocebo*" [tiab] OR "placebo induced"[tiab] OR "bogus"[tiab] OR "null treatment"[tiab] OR "inactive treatment"[tiab] OR "Placebos/adverse effects"[majr]) AND ((("Communication"[majr] AND "Physician-Patient Relations"[majr]) OR ("Communication"[majr] AND "Physicians"[majr] AND "Patients"[majr]) OR "Communicate Patient"[title/abstract:~5] OR "Communicate Patients"[title/abstract:~5] OR "Communicated Patient"[title/abstract:~5] OR "Communicated Patients"[title/abstract:~5] OR "Communicating Patient"[title/abstract:~5] OR "Communicating Patients"[title/abstract:~5] OR "Communication Patient"[title/abstract:~5] OR "Communication Patients"[title/abstract:~5] OR "Inform Patient"[title/abstract:~5] OR "Inform Patients"[title/abstract:~5] OR "Informed Patient"[title/abstract:~5] OR "Informed Patients"[title/abstract:~5] OR "Information Patient"[title/abstract:~5] OR "Information Patients"[title/abstract:~5] OR "Informing Patient"[title/abstract:~5] OR "Informing Patients"[title/abstract:~5] OR "Communicate Individual"[title/abstract:~5] OR "Communicate Individuals"[title/abstract:~5] OR "Communicated Individual"[title/abstract:~5] OR "Communicated Individuals"[title/abstract:~5] OR "Communicating Individual"[title/abstract:~5] OR "Communicating Individuals"[title/abstract:~5] OR "Communication Individual"[title/abstract:~5] OR "Communication Individuals"[title/abstract:~5] OR "Inform Individual"[title/abstract:~5] OR "Inform Individuals"[title/abstract:~5] OR "Informed Individual"[title/abstract:~5] OR "Informed Individuals"[title/abstract:~5] OR "Information Individual"[title/abstract:~5] OR "Information Individuals"[title/abstract:~5] OR "Informing Individual"[title/abstract:~5] OR "Informing Individuals"[title/abstract:~5] OR "Explain Patient"[title/abstract:~5] OR "Explain Patients"[title/abstract:~5] OR "Explaining Patient"[title/abstract:~5] OR "Explaining Patients"[title/abstract:~5] OR "Explanation Patient"[title/abstract:~5] OR "Explanation Patients"[title/abstract:~5] OR "Explanations Patient"[title/abstract:~5] OR "Explanations Patients"[title/abstract:~5] OR "Explain Individual"[title/abstract:~5] OR "Explain Individuals"[title/abstract:~5] OR "Explaining Individual"[title/abstract:~5] OR "Explaining Individuals"[title/abstract:~5] OR "Explanation Individual"[title/abstract:~5] OR "Explanation Individuals"[title/abstract:~5] OR "Explanations Individual"[title/abstract:~5] OR "Explanations Individuals"[title/abstract:~5] OR "Language"[Mesh] OR "Communications Media"[Mesh])) OR ((("Anticipation, Psychological"[majr] OR "beliefs about medicines"[ti] OR "expectation*" [ti] OR "improvement expect*" [ti] OR "Medicine-related belief*" [ti] OR "Medicine-related beliefs"[ti] OR "mindset"[ti] OR "negative belief*" [ti] OR "negative beliefs"[ti] OR "negative expectat*" [ti] OR

"patient belief*" (ti) OR "patient beliefs" (ti) OR "patient expectation*" (ti) OR "patient perspective" (ti) OR "patients' belief*" (ti) OR "patients' beliefs" (ti) OR "patients expectation*" (ti) OR "patients' expectation*" (ti) OR "patients perspective" (ti) OR "patients' perspective" (ti) OR "patient's perspective" (ti) OR "patients trust" (ti) OR "patients' trust" (ti) OR "patient's trust" (ti) OR "perceived effectiveness" (ti) OR "perceived efficacy" (ti) OR "positive belief" (ti) OR "positive belief*" (ti) OR "positive expectat*" (ti) OR "respondent expectation*" (ti) OR "respondents expectation*" (ti) OR "respondents' expectation*" (ti) OR "suggestibility" (ti) OR "treatment expect*" (ti) OR "beliefs about medicines" (title:~3) OR "improvement expectation" (title:~3) OR "improvement expectations" (title:~3) OR "Medicine-related belief" (title:~3) OR "Medicine-related beliefs" (title:~3) OR "negative belief" (title:~3) OR "negative beliefs" (title:~3) OR "negative expectation" 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"respondents expectations" (title:~3) OR "treatment expectation" (title:~3) OR "treatment expectations" (title:~3) OR suggestion*(tiab) OR (("expectancy" (ti) OR "expectancies" (ti)) NOT ("Life Expectancy" (Mesh) OR "Life Expectancy" (tiab) OR "Life Expectan*" (tiab)))) AND (("Communication" (majr) AND "Physician-Patient Relations" (majr)) OR ("Communication" (majr) AND "Physicians" (majr) AND "Patients" (majr)) OR "Communicate Patient" (title:~5) OR "Communicate Patients" (title:~5) OR "Communicated Patient" (title:~5) OR "Communicated Patients" (title:~5) OR "Communicating Patient" (title:~5) OR "Communicating Patients" (title:~5) OR "Communication Patient" (title:~5) OR "Communication Patients" (title:~5) OR "Inform Patient" (title:~5) OR "Inform Patients" (title:~5) OR "Informed Patient" (title:~5) OR "Informed Patients" (title:~5) OR "Information Patient" (title:~5) OR "Information Patients" (title:~5) OR "Informing Patient" (title:~5) OR "Informing Patients" (title:~5) OR 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(title:~5) OR "Explaining Individuals" (title:~5) OR "Explanation Individual" (title:~5) OR "Explanation Individuals" (title:~5) OR "Explanations Individual" (title:~5) OR "Explanations Individuals" (title:~5) OR "treatment dropout" (tiab) OR "treatment dropouts" (tiab) OR "therapy dropout" (tiab) OR "therapy dropouts" (tiab) OR "nonverbal communication" (mesh) OR "health communication" (mesh))) OR (("Placebo Effect" (majr) OR "Placebo Effect" (tiab) OR "Placebo Effects" (tiab) OR "Placebo Effect*" (tiab) OR "Placebo Response" (tiab) OR "Placebo Responses" (tiab) OR "Placebo Respon*" (tiab) OR "Nocebo Effect" (majr) OR "Nocebo Effect" (tiab) OR "Nocebo Effects" (tiab) OR "Nocebo Effect*" (tiab) OR "Nocebo Response" (tiab) OR "Nocebo Responses" (tiab) OR "Nocebo Respon*" (tiab) OR "nocebo" (tiab) OR "nocebo*" (tiab) OR "placebo induced" (tiab) OR "bogus" (tiab) OR "null treatment" (tiab) OR "inactive treatment" (tiab) OR "Placebos/adverse effects" (majr) OR "positive expectat*" (tiab) OR "negative expectat*" (tiab) OR "treatment



expect** (tiab) OR "patients' expectation** (tiab) OR "patients expectation** (tiab) OR "patient expectation** (tiab) OR "respondents' expectation** (tiab) OR "respondents expectation** (tiab) OR "respondent expectation** (tiab) OR "improvement expect** (tiab) OR "mindset" (tiab) OR "patient's trust" (tiab) OR "patients trust" (tiab) OR "patients' trust" (tiab) OR "perceived efficacy" (tiab) OR "perceived effectiveness" (tiab) OR "suggestibility" (tiab) OR "patient beliefs" (tiab) OR "patient belief** (tiab) OR "patients' beliefs" (tiab) OR "patients' belief** (tiab) OR "negative beliefs" (tiab) OR "negative belief** (tiab) OR "positive belief" (tiab) OR "positive belief** (tiab) OR "beliefs about medicines" (tiab) OR "Medicine-related beliefs" (tiab) OR "Medicine-related belief** (tiab) OR "patient's perspective" (tiab) OR "patients perspective" (tiab) OR "patient perspective" (tiab) OR "patients' perspective" (tiab) OR "Anticipation, Psychological" (majr) OR "expectation** (ti) OR ("expectancy" (tiab) OR "expectancies" (tiab)) NOT ("Life Expectancy" (Mesh) OR "Life Expectancy" (tiab) OR "Life Expectan** (tiab)))) AND ("information frame" (tiab) OR "information frames" (tiab) OR "information framing" (tiab) OR "information fram** (tiab) OR "message frame" (tiab) OR "message frames" (tiab) OR "message framing" (tiab) OR "message fram** (tiab) OR "frame effect" (tiab) OR "frame effects" (tiab) OR "positive frame" (tiab) OR "positive frames" (tiab) OR "positive framing" (tiab) OR "positive fram** (tiab) OR "negative frame" (tiab) OR "negative frames" (tiab) OR "negative framing" (tiab) OR "negative fram** (tiab) OR "positively framed" (tiab) OR "positively fram** (tiab) OR "negatively framed" (tiab) OR "negatively fram** (tiab) OR "information frame" [title/abstract:~6] OR "information frames" [title/abstract:~6] OR "information framing" [title/abstract:~6] OR "message frame" [title/abstract:~6] OR "message frames" [title/abstract:~6] OR "message framing" [title/abstract:~6] OR "frame effect" [title/abstract:~6] OR "frame effects" [title/abstract:~6] OR "positive frame" [title/abstract:~6] OR "positive frames" [title/abstract:~6] OR "positive framing" [title/abstract:~6] OR "negative frame" [title/abstract:~6] OR "negative frames" [title/abstract:~6] OR "negative framing" [title/abstract:~6] OR "positively framed" [title/abstract:~6] OR "negatively framed" [title/abstract:~6])) OR (("Placebo Effect" (majr) OR "Placebo Effect" (tiab) OR "Placebo Effects" (tiab) OR "Placebo Effect** (tiab) OR "Placebo Response" (tiab) OR "Placebo Responses" (tiab) OR "Placebo Respon** (tiab) OR "Nocebo Effect" (majr) OR "Nocebo Effect" (tiab) OR "Nocebo Effects" (tiab) OR "Nocebo Effect** (tiab) OR "Nocebo Response" (tiab) OR "Nocebo Responses" (tiab) OR "Nocebo Respon** (tiab) OR "nocebo" (tiab) OR "nocebo** (tiab) OR "placebo induced" (tiab) OR "bogus" (tiab) OR "null treatment" (tiab) OR "inactive treatment" (tiab) OR "Placebos/adverse effects" (majr) OR "expectation interventions" (tiab) OR "expectation intervention" (tiab) OR "positive expectat** (tiab) OR "negative expectat** (tiab) OR "treatment expect** (tiab) OR "patients' expectation** (tiab) OR "patients expectation** (tiab) OR "patient expectation** (tiab) OR "respondents' expectation** (tiab) OR "respondents expectation** (tiab) OR "respondent expectation** (tiab) OR "improvement expect** (tiab) OR "mindset" (tiab) OR "patient's trust" (tiab) OR "patients trust" (tiab) OR "patients' trust" (tiab) OR "perceived efficacy" (tiab) OR "perceived effectiveness" (tiab) OR "suggestibility" (tiab) OR "patient beliefs" (tiab) OR "patient belief** (tiab) OR "patients' beliefs" (tiab) OR "patients' belief** (tiab) OR "negative beliefs" (tiab) OR "negative belief** (tiab) OR "positive belief" (tiab) OR "positive belief** (tiab) OR "beliefs about medicines" (tiab) OR "Medicine-related beliefs" (tiab) OR "Medicine-related belief** (tiab) OR "patient's perspective" (tiab) OR "patients perspective" (tiab) OR "patient perspective" (tiab) OR "patients' perspective" (tiab) OR "Anticipation, Psychological" (majr) OR "expectation** (ti) OR ("expectancy" (tiab) OR "expectancies" (tiab)) NOT ("Life Expectancy" (Mesh) OR "Life Expectancy" (tiab) OR "Life Expectan** (tiab)))) AND ("Deception" (mesh) OR "deception" (tiab) OR "deceptive" (tiab) OR "nondeceptive" (tiab) OR "nondecept** (tiab))) OR (("Placebo Effect" (majr) OR "Placebo Effect" (tiab) OR "Placebo Effects" (tiab) OR "Placebo Effect** (tiab) OR "Placebo Response" (tiab) OR "Placebo Responses" (tiab) OR "Placebo Respon** (tiab) OR "Nocebo Effect" (majr) OR "Nocebo Effect" (tiab) OR "Nocebo Effects" (tiab) OR "Nocebo Effect** (tiab) OR "Nocebo Response" (tiab) OR "Nocebo Responses" (tiab) OR "Nocebo Respon** (tiab) OR "nocebo" (tiab) OR "nocebo** (tiab) OR "placebo induced" (tiab) OR "bogus" (tiab) OR "null treatment" (tiab) OR "inactive

treatment"[tiab] OR "Placebos/adverse effects"[majr]) AND "Attitude to Health"[majr]) OR ("Placebo Effect"[majr] OR "Placebo Effect"[tiab] OR "Placebo Effects"[tiab] OR "Placebo Effect*"[tiab] OR "Placebo Response"[tiab] OR "Placebo Responses"[tiab] OR "Placebo Respon*"[tiab] OR "Nocebo Effect"[majr] OR "Nocebo Effect"[tiab] OR "Nocebo Effects"[tiab] OR "Nocebo Effect*"[tiab] OR "Nocebo Response"[tiab] OR "Nocebo Responses"[tiab] OR "Nocebo Respon*"[tiab] OR "nocebo"[tiab] OR "nocebo*"[tiab] OR "placebo induced"[tiab] OR "bogus"[tiab] OR "null treatment"[tiab] OR "inactive treatment"[tiab] OR "Placebos/adverse effects"[majr]) AND ("expectation interventions"[tiab] OR "expectation intervention"[tiab] OR "positive expectat*"[tiab] OR "negative expectat*"[tiab] OR "treatment expect*"[tiab] OR "patients' expectation*"[tiab] OR "patients expectation*"[tiab] OR "patient expectation*"[tiab] OR "respondents' expectation*"[tiab] OR "respondents expectation*"[tiab] OR "respondent expectation*"[tiab] OR "improvement expect*"[tiab] OR "mindset"[tiab] OR "patient's trust"[tiab] OR "patients trust"[tiab] OR "patients' trust"[tiab] OR "perceived efficacy"[tiab] OR "perceived effectiveness"[tiab] OR "suggestibility"[tiab] OR "patient beliefs"[tiab] OR "patient belief*"[tiab] OR "patients' beliefs"[tiab] OR "patients' belief*"[tiab] OR "negative beliefs"[tiab] OR "negative belief*"[tiab] OR "positive belief"[tiab] OR "positive belief*"[tiab] OR "beliefs about medicines"[tiab] OR "Medicine-related beliefs"[tiab] OR "Medicine-related belief*"[tiab] OR "patient's perspective"[tiab] OR "patients perspective"[tiab] OR "patient perspective"[tiab] OR "patients' perspective"[tiab] OR "Anticipation, Psychological"[majr] OR "expectation*"[ti] OR ("expectancy"[tiab] OR "expectancies"[tiab]) NOT ("Life Expectancy"[Mesh] OR "Life Expectancy"[tiab] OR "Life Expectan*"[tiab])))) OR ("Placebo Effect belief"[title/abstract:~6] OR "Placebo Effects belief"[title/abstract:~6] OR "Placebo Response belief"[title/abstract:~6] OR "Placebo Responses belief"[title/abstract:~6] OR "nocebo belief"[title/abstract:~6] OR "placebo induced belief"[title/abstract:~6] OR "bogus belief"[title/abstract:~6] OR "null treatment belief"[title/abstract:~6] OR "inactive treatment belief"[title/abstract:~6] OR "Placebo Effect beliefs"[title/abstract:~6] OR "Placebo Effects beliefs"[title/abstract:~6] OR "Placebo Response beliefs"[title/abstract:~6] OR "Placebo Responses beliefs"[title/abstract:~6] OR "nocebo beliefs"[title/abstract:~6] OR "placebo induced beliefs"[title/abstract:~6] OR "bogus beliefs"[title/abstract:~6] OR "null treatment beliefs"[title/abstract:~6] OR "inactive treatment beliefs"[title/abstract:~6]) OR ("Communication"[mesh] AND "Physician-Patient Relations"[mesh] AND "Physicians"[majr]) OR ("patient mindset"[tiab] OR "patients mindset"[tiab] OR "patient's mindset"[tiab] OR "patient mindsets"[tiab]) OR "Nocebo Effect"[majr] OR ("Placebo Effect"[majr] OR "Placebo"[ti] OR "nocebo"[ti]) AND ("side effect"[ti] OR "side effects"[ti] OR "analges*"[ti] OR "hyperalges*"[ti]) OR ("Pain Perception"[mesh] AND "Patient Education as Topic"[mesh])) NOT ("Animals"[mesh] NOT "Humans"[mesh])

Embase

<http://ovidsp.ovid.com/ovidweb.cgi?T=JS&PAGE=main&MODE=ovid&D=oemezd>

((exp *Placebo Effect"/ OR "Placebo Effect".tw. OR "Placebo Effects".tw. OR "Placebo Effect*".tw. OR "Placebo Response".tw. OR "Placebo Responses".tw. OR "Placebo Respon*".tw. OR exp *Nocebo Effect"/ OR "Nocebo Effect".tw. OR "Nocebo Effects".tw. OR "Nocebo Effect*".tw. OR "Nocebo Response".tw. OR "Nocebo Responses".tw. OR "Nocebo Respon*".tw. OR nocebo.tw. OR nocebo*.tw. OR "placebo induced".tw. OR bogus.tw. OR "null treatment".tw. OR "inactive treatment".tw. OR exp *Placebos/adverse effects"/) AND ((exp *Communication/ AND exp *Physician-Patient Relations"/) OR (exp *Communication/ AND exp *Physicians/ AND exp *Patients/) OR "Communicate Patient".tw. OR "Communicate Patients".tw. OR "Communicated Patient".tw. OR "Communicated Patients".tw. OR "Communicating Patient".tw. OR "Communicating Patients".tw. OR "Communication Patient".tw. OR "Communication Patients".tw. OR "Inform Patient".tw. OR "Inform Patients".tw. OR "Informed Patient".tw. OR "Informed Patients".tw. OR "Information Patient".tw. OR "Information Patients".tw. OR "Informing

Patient".tw. OR "Informing Patients".tw. OR "Communicate Individual".tw. OR "Communicate Individuals".tw. OR "Communicated Individual".tw. OR "Communicated Individuals".tw. OR "Communicating Individual".tw. OR "Communicating Individuals".tw. OR "Communication Individual".tw. OR "Communication Individuals".tw. OR "Inform Individual".tw. OR "Inform Individuals".tw. OR "Informed Individual".tw. OR "Informed Individuals".tw. OR "Information Individual".tw. OR "Information Individuals".tw. OR "Informing Individual".tw. OR "Informing Individuals".tw. OR "Explain Patient".tw. OR "Explain Patients".tw. OR "Explaining Patient".tw. OR "Explaining Patients".tw. OR "Explanation Patient".tw. OR "Explanation Patients".tw. OR "Explanations Patient".tw. OR "Explanations Patients".tw. OR "Explain Individual".tw. OR "Explain Individuals".tw. OR "Explaining Individual".tw. OR "Explaining Individuals".tw. OR "Explanation Individual".tw. OR "Explanation Individuals".tw. OR "Explanations Individual".tw. OR "Explanations Individuals".tw. OR exp "Communications Media"/) OR ((exp **Anticipation, Psychological"/ OR "beliefs about medicines".ti. OR expectation*.ti. OR "improvement expect*.ti. OR "Medicine-related belief*.ti. OR "Medicine-related beliefs".ti. OR mindset.ti. OR "negative belief*.ti. OR "negative beliefs".ti. OR "negative expectat*.ti. OR "patient belief*.ti. OR "patient beliefs".ti. OR "patient expectation*.ti. OR "patient perspective".ti. OR "patients' belief*.ti. OR "patients' beliefs".ti. OR "patients expectation*.ti. OR "patients' expectation*.ti. OR "patients perspective".ti. OR "patients' perspective".ti. OR "patient's perspective".ti. OR "patients trust".ti. OR "patients' trust".ti. OR "patient's trust".ti. OR "perceived effectiveness".ti. OR "perceived efficacy".ti. OR "positive belief".ti. OR "positive belief*.ti. OR "positive expectat*.ti. OR "respondent expectation*.ti. OR "respondents expectation*.ti. OR "respondents' expectation*.ti. 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OR "treatment expectation".ti. OR "treatment expectations".ti. OR suggestion*.ti. OR ((expectancy.ti. OR expectancies.ti.) NOT (exp "Life Expectancy"/ OR "Life Expectancy".tw. OR "Life Expectan*.tw.))) AND ((exp *Communication/ AND exp **Physician-Patient Relations"/) OR (exp *Communication/ AND exp *Physicians/ AND exp *Patients/) OR "Communicate Patient".ti. OR "Communicate Patients".ti. OR "Communicated Patient".ti. OR "Communicated Patients".ti. OR "Communicating Patient".ti. OR "Communicating Patients".ti. OR "Communication Patient".ti. OR "Communication Patients".ti. OR "Inform Patient".ti. OR "Inform Patients".ti. OR "Informed Patient".ti. OR "Informed Patients".ti. OR "Information Patient".ti. OR "Information Patients".ti. OR "Informing Patient".ti. OR "Informing Patients".ti. OR "Communicate Individual".ti. OR "Communicate Individuals".ti. OR "Communicating Individual".ti. OR "Communicating Individuals".ti. OR "Communication Individual".ti. OR "Communication Individuals".ti. 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OR exp **nonverbal communication"/ OR exp **health communication"/) OR ((exp **Placebo Effect"/ OR "Placebo Effect".tw. OR "Placebo Effects".tw. OR "Placebo Effect*.tw. OR "Placebo Response".tw. OR "Placebo Responses".tw. OR "Placebo Respon*.tw. OR exp **Nocebo Effect"/ OR "Nocebo Effect".tw. OR "Nocebo Effects".tw. OR "Nocebo Effect*.tw. OR "Nocebo Response".tw. OR "Nocebo Responses".tw. OR "Nocebo Respon*.tw. OR nocebo.tw. OR nocebo*.tw. OR "placebo induced".tw. OR bogus.tw. OR "null treatment".tw. OR "inactive treatment".tw. OR exp **Placebos/adverse effects"/ OR "positive expectat*.tw. OR "negative expectat*.tw. OR "treatment expect*.tw. OR "patients' expectation*.tw. OR "patients expectation*.tw. OR "patient expectation*.tw. OR "respondents' expectation*.tw. OR "respondents expectation*.tw. OR "respondent expectation*.tw. OR "improvement expect*.tw. OR mindset.ti. OR "patient's trust".tw. OR "patients trust".tw. OR "patients' trust".tw. OR "perceived efficacy".tw. OR "perceived effectiveness".tw. OR suggestibility.tw. OR "patient beliefs".tw. OR "patient belief*.tw. OR "patients' beliefs".tw. OR "patients' belief*.tw. OR "negative beliefs".tw. OR "negative belief*.tw. OR "positive belief".tw. OR "positive belief*.tw. OR "beliefs about medicines".tw. OR "Medicine-related beliefs".tw. OR "Medicine-related belief*.tw. OR "patient's perspective".tw. OR "patients perspective".tw. OR "patient perspective".tw. OR "patients' perspective".tw. OR exp **Anticipation, Psychological"/ OR expectation*.ti. OR ((expectancy.tw. OR expectancies.tw.) NOT (exp "Life Expectancy"/ OR "Life Expectancy".tw. OR "Life Expectan*.tw.))) AND ("information frame".tw. OR "information frames".tw. OR "information framing".tw. OR "information fram*.tw. OR "message frame".tw. OR "message frames".tw. OR "message framing".tw. OR "message fram*.tw. OR "frame effect".tw. OR "frame effects".tw. OR "positive frame".tw. OR "positive

frames".tw. OR "positive framing".tw. OR "positive fram*".tw. OR "negative frame".tw. OR "negative frames".tw. OR "negative framing".tw. OR "negative fram*".tw. OR "positively framed".tw. OR "positively fram*".tw. OR "negatively framed".tw. OR "negatively fram*".tw. OR "information frame".tw. OR "information frames".tw. OR "information framing".tw. OR "message frame".tw. OR "message frames".tw. OR "message framing".tw. OR "frame effect".tw. OR "frame effects".tw. OR "positive frame".tw. OR "positive frames".tw. OR "positive framing".tw. OR "negative frame".tw. OR "negative frames".tw. OR "negative framing".tw. OR "positively framed".tw. OR "negatively framed".tw.)) OR ((exp "Placebo Responses".tw. OR "Placebo Effect".tw. OR "Placebo Effects".tw. OR "Placebo Effect".tw. OR "Placebo Response".tw. OR "Placebo Responses".tw. OR "Placebo Respon*".tw. OR exp "Nocebo Effect"/ OR "Nocebo Effect".tw. OR "Nocebo Effects".tw. OR "Nocebo Effect".tw. OR "Nocebo Response".tw. 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Web of Science

<http://isiknowledge.com/wos>

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Effect**) OR (TI="Nocebo Response" OR AB="Nocebo Response") OR (TI="Nocebo Responses" OR AB="Nocebo Responses") OR (TI="Nocebo Respon** OR AB="Nocebo Respon**") OR (TI=nocebo OR AB=nocebo) OR (TI=nocebo* OR AB=nocebo*) OR (TI="placebo induced" OR AB="placebo induced") OR (TI=bogus OR AB=bogus) OR (TI="null treatment" OR AB="null treatment") OR (TI="inactive treatment" OR AB="inactive treatment") OR TS="Placebos/adverse effects" OR (TI="expectation interventions" OR AB="expectation interventions") OR (TI="expectation intervention" OR AB="expectation intervention") OR (TI="positive expectat** OR AB="positive expectat**") OR (TI="negative expectat** OR AB="negative expectat**") OR (TI="treatment expect** OR AB="treatment expect**") OR (TI="patients' expectation** OR AB="patients' expectation**") OR (TI="patients expectation** OR AB="patients expectation**") OR (TI="patient expectation** OR AB="patient expectation**") OR (TI="respondents' expectation** OR AB="respondents' expectation**") 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Cochrane Library

<https://www.cochranelibrary.com/advanced-search/search-manager>

((("Placebo Effect" OR ("Placebo Effect" OR "Placebo Effect") OR ("Placebo Effects" OR "Placebo Effects") OR ("Placebo
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perspective" OR "patient's perspective" OR "patients trust" OR "patients' trust" OR "patient's trust" OR "perceived effectiveness" OR "perceived efficacy" OR "positive belief" OR "positive belief**" OR "positive expectat**" OR "respondent expectation**" OR "respondents expectation**" OR "respondents' expectation**" OR suggestibility OR "treatment expect**" OR "beliefs about medicines" OR "improvement expectation" OR "improvement expectations" OR "Medicine-related belief" OR "Medicine-related beliefs" OR "negative belief" OR "negative beliefs" OR "negative expectation" OR "negative expectations" OR "patient belief" OR "patient beliefs" OR "patient expectation" OR "patient expectations" OR "patient perspective" OR "patient perspectives" OR "patients belief" OR "patients beliefs" OR "patients expectation" OR "patients expectations" OR "patients perspective" OR "patients perspectives" OR "patients trust" OR "patients trust" OR "perceived effectiveness" OR "perceived efficacy" OR "positive 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"Informing Patients" OR "Communicate Individual" OR "Communicate Individuals" OR "Communicated Individual" OR "Communicated Individuals" OR "Communicating Individual" OR "Communicating Individuals" OR "Communication Individual" OR "Communication Individuals" OR "Inform Individual" OR "Inform Individuals" OR "Informed Individual" OR "Informed Individuals" OR "Information Individual" OR "Information Individuals" OR "Informing Individual" OR "Informing Individuals" OR "Explain Patient" OR "Explain Patients" OR "Explaining Patient" OR "Explaining Patients" OR "Explanation Patient" OR "Explanation Patients" OR "Explanations Patient" OR "Explanations Patients" OR "Explain Individual" OR "Explain Individuals" OR "Explaining Individual" OR "Explaining Individuals" OR "Explanation Individual" OR "Explanation Individuals" OR "Explanations Individual" OR "Explanations Individuals" OR ("treatment dropout" OR "treatment dropout") OR ("treatment dropouts" OR "treatment dropouts") OR ("therapy dropout" OR "therapy dropout") OR ("therapy dropouts" OR "therapy dropouts" OR "nonverbal communication" OR "health communication")))) OR (("Placebo Effect" OR ("Placebo Effect" OR "Placebo Effect") OR ("Placebo Effects" OR "Placebo Effects") OR ("Placebo Effect**" OR "Placebo Effect**") OR ("Placebo Response" OR "Placebo Response") OR ("Placebo Responses" OR "Placebo Responses") OR ("Placebo Respon**" OR "Placebo Respon**") OR "Nocebo Effect" OR ("Nocebo Effect" OR "Nocebo Effect") OR ("Nocebo Effects" OR "Nocebo Effects") OR ("Nocebo Effect**" OR "Nocebo Effect**") OR ("Nocebo Response" OR "Nocebo Response") OR ("Nocebo Responses" OR "Nocebo Responses") OR ("Nocebo Respon**" OR "Nocebo Respon**") OR (nocebo OR nocebo) OR (nocebo* OR nocebo*) OR ("placebo induced" OR "placebo induced") OR (bogus OR bogus) OR ("null treatment" OR "null treatment") OR ("inactive treatment" OR "inactive treatment") OR ("Placebos/adverse effects" OR ("positive expectat**" OR "positive expectat**") OR ("negative expectat**" OR "negative expectat**") OR ("treatment expect**" OR "treatment expect**") OR ("patients' expectation**" OR "patients' expectation**") OR ("patients expectation**" OR "patients expectation**") OR ("patient expectation**" OR "patient expectation**") OR ("respondents' expectation**" OR "respondents' expectation**") OR ("respondents expectation**" OR "respondents expectation**") OR ("respondent expectation**" OR "respondent expectation**") OR ("improvement expect**" OR "improvement expect**") OR (mindset OR mindset) OR ("patient's trust" OR "patient's trust") OR ("patients trust" OR "patients trust") OR ("patients' trust" OR "patients' trust") OR ("perceived efficacy" OR "perceived efficacy") OR ("perceived effectiveness" OR "perceived effectiveness") OR (suggestibility OR suggestibility) OR ("patient beliefs" OR "patient beliefs") OR ("patient belief**" OR "patient belief**") OR ("patients' beliefs" OR "patients' beliefs") OR ("patients' belief**" OR "patients' belief**") OR ("negative beliefs" OR "negative beliefs") OR ("negative belief**" OR "negative belief**") OR ("positive belief" OR "positive belief") OR ("positive belief**" OR "positive belief**") OR ("beliefs about medicines" OR "beliefs about medicines") OR ("Medicine-related beliefs" OR "Medicine-related beliefs") OR ("Medicine-related belief**" OR "Medicine-related belief**") OR ("patient's perspective" OR "patient's perspective") OR ("patients perspective" OR "patients perspective") OR ("patient perspective" OR "patient perspective") OR ("patients' perspective" OR "patients' perspective") OR "Anticipation, Psychological" OR expectation* OR (((expectancy OR expectancy) OR (expectancies OR expectancies)) NOT ("Life Expectancy" OR ("Life Expectancy" OR "Life Expectancy") OR ("Life Expectan**" OR "Life Expectan**")))) AND ((("information frame" OR "information frame") OR ("information frames" OR "information frames") OR ("information framing" OR "information framing") OR ("information fram**" OR "information fram**") OR ("message frame" OR "message frame") OR ("message frames" OR "message frames") OR ("message framing" OR "message framing") OR ("message fram**" OR "message fram**") OR ("frame effect" OR "frame effect") OR ("frame effects" OR "frame effects") OR ("positive frame" OR "positive frame") OR ("positive frames" OR "positive frames") OR ("positive framing" OR "positive framing") OR ("positive fram**" OR "positive fram**") OR ("negative frame" OR "negative frame") OR ("negative frames" OR "negative frames") OR ("negative framing" OR "negative framing") OR ("negative fram**" OR "negative fram**") OR ("positively framed" OR "positively framed") OR ("positively fram**" OR "positively fram**") OR ("negatively framed" OR "negatively framed") OR ("negatively fram**" OR "negatively fram**") OR ("information frame" OR "information

frame") OR ("information frames" OR "information frames") OR ("information framing" OR "information framing") OR ("message frame" OR "message frame") OR ("message frames" OR "message frames") OR ("message framing" OR "message framing") OR ("frame effect" OR "frame effect") OR ("frame effects" OR "frame effects") OR ("positive frame" OR "positive frame") OR ("positive frames" OR "positive frames") OR ("positive framing" OR "positive framing") OR ("negative frame" OR "negative frame") OR ("negative frames" OR "negative frames") OR ("negative framing" OR "negative framing") OR ("positively framed" OR "positively framed") OR ("negatively framed" OR "negatively framed")) OR ("Placebo Effect" OR "Placebo Effect") OR ("Placebo Effect" OR "Placebo Effects" OR "Placebo Effects") OR ("Placebo Effect**" OR "Placebo Effect**") OR ("Placebo Response" OR "Placebo Response") OR ("Placebo Responses" OR "Placebo Responses") OR ("Placebo Respon**" OR "Placebo Respon**") OR "Nocebo Effect" OR ("Nocebo Effect" OR "Nocebo Effect") OR ("Nocebo Effects" OR "Nocebo Effects") OR ("Nocebo Effect**" OR "Nocebo Effect**") OR ("Nocebo Response" OR "Nocebo Response") OR ("Nocebo Responses" OR "Nocebo Responses") OR ("Nocebo Respon**" OR "Nocebo Respon**") OR (nocebo OR nocebo) OR (nocebo* OR nocebo*) OR ("placebo induced" OR "placebo induced") OR (bogus OR bogus) OR ("null treatment" OR "null treatment") OR ("inactive treatment" OR "inactive treatment") OR "Placebos/adverse effects" OR ("expectation interventions" OR "expectation interventions") OR ("expectation intervention" OR "expectation intervention") OR ("positive expectat**" OR "positive expectat**") OR ("negative expectat**" OR "negative expectat**") OR ("treatment expect**" OR "treatment expect**") OR ("patients' expectation**" OR "patients' expectation**") OR ("patients' expectation**" OR "patients' expectation**") OR ("patient expectation**" OR "patient expectation**") OR ("respondents' expectation**" OR "respondents' expectation**") OR ("respondents' expectation**" OR "respondents' expectation**") OR ("respondent expectation**" OR "respondent expectation**") OR ("improvement expect**" OR "improvement expect**") OR (mindset OR mindset) OR ("patient's trust" OR "patient's trust") OR ("patients trust" OR "patients trust") OR ("patients' trust" OR "patients' trust") OR ("perceived efficacy" OR "perceived efficacy") OR ("perceived effectiveness" OR "perceived effectiveness") OR (suggestibility OR suggestibility) OR ("patient beliefs" OR "patient beliefs") OR ("patient belief**" OR "patient belief**") OR ("patients' beliefs" OR "patients' beliefs") OR ("patients' belief**" OR "patients' belief**") OR ("negative beliefs" OR "negative beliefs") OR ("negative belief**" OR "negative belief**") OR ("positive belief" OR "positive belief") OR ("positive belief**" OR "positive belief**") OR ("beliefs about medicines" OR "beliefs about medicines") OR ("Medicine-related beliefs" OR "Medicine-related beliefs") OR ("Medicine-related belief**" OR "Medicine-related belief**") OR ("patient's perspective" OR "patient's perspective") OR ("patients perspective" OR "patients perspective") OR ("patient perspective" OR "patient perspective") OR ("patients' perspective" OR "patients' perspective") OR "Anticipation, Psychological" OR expectation* OR (((expectancy OR expectancy) OR (expectancies OR expectancies)) NOT ("Life Expectancy" OR "Life Expectancy" OR "Life Expectancy") OR ("Life Expectan**" OR "Life Expectan**")) AND (Deception OR (deception OR deception) OR (deceptive OR deceptive) OR (nondeceptive OR nondeceptive) OR (nondecept** OR nondecept**)) OR ("Placebo Effect" OR "Placebo Effect" OR "Placebo Effect") OR ("Placebo Effects" OR "Placebo Effects") OR ("Placebo Effect**" OR "Placebo Effect**") OR ("Placebo Response" OR "Placebo Response") OR ("Placebo Responses" OR "Placebo Responses") OR ("Placebo Respon**" OR "Placebo Respon**") OR "Nocebo Effect" OR ("Nocebo Effect" OR "Nocebo Effect") OR ("Nocebo Effects" OR "Nocebo Effects") OR ("Nocebo Effect**" OR "Nocebo Effect**") OR ("Nocebo Response" OR "Nocebo Response") OR ("Nocebo Responses" OR "Nocebo Responses") OR ("Nocebo Respon**" OR "Nocebo Respon**") OR (nocebo OR nocebo) OR (nocebo* OR nocebo*) OR ("placebo induced" OR "placebo induced") OR (bogus OR bogus) OR ("null treatment" OR "null treatment") OR ("inactive treatment" OR "inactive treatment") OR "Placebos/adverse effects") AND ("Attitude to Health" OR ("Placebo Effect" OR "Placebo Effect" OR "Placebo Effect") OR ("Placebo Effects" OR "Placebo Effects") OR ("Placebo Effect**" OR "Placebo Effect**") OR ("Placebo Response" OR "Placebo Response") OR ("Placebo Responses" OR "Placebo Responses") OR ("Placebo Respon**" OR "Placebo Respon**") OR "Nocebo Effect" OR ("Nocebo Effect" OR "Nocebo Effect") OR ("Nocebo Effects" OR "Nocebo Effects") OR ("Nocebo Effect**" OR "Nocebo Effect**") OR ("Nocebo Response" OR "Nocebo Response") OR ("Nocebo Responses" OR "Nocebo Responses") OR ("Nocebo Respon**" OR "Nocebo Respon**") OR (nocebo OR nocebo) OR (nocebo* OR nocebo*) OR ("placebo induced" OR "placebo induced") OR (bogus OR bogus) OR ("null treatment" OR "null treatment") OR ("inactive treatment" OR "inactive treatment") OR "Placebos/adverse effects") AND ("expectation interventions" OR "expectation interventions") OR ("expectation intervention" OR "expectation intervention") OR ("positive expectat**" OR "positive expectat**") OR ("negative expectat**" OR "negative expectat**") OR ("treatment expect**" OR "treatment expect**") OR ("patients' expectation**" OR "patients' expectation**") OR ("patients' expectation**" OR "patients' expectation**") OR ("patient expectation**" OR "patient expectation**") OR ("respondents' expectation**" OR "respondents' expectation**") OR ("respondents' expectation**" OR "respondents' expectation**") OR ("respondent expectation**" OR "respondent expectation**") OR ("improvement expect**" OR "improvement expect**") OR (mindset OR mindset) OR ("patient's trust" OR "patient's trust") OR ("patients trust" OR "patients trust") OR ("patients' trust" OR "patients' trust") OR ("perceived efficacy" OR "perceived efficacy") OR ("perceived effectiveness" OR "perceived effectiveness") OR (suggestibility OR suggestibility) OR ("patient beliefs" OR "patient beliefs") OR ("patient belief**" OR "patient belief**") OR ("patients' beliefs" OR "patients' beliefs") OR ("patients' belief**" OR "patients' belief**") OR ("negative beliefs" OR "negative beliefs") OR ("negative belief**" OR "negative belief**") OR ("positive belief" OR "positive belief") OR ("positive belief**" OR "positive belief**") OR ("beliefs about medicines" OR "beliefs about medicines") OR ("Medicine-related beliefs" OR "Medicine-related beliefs") OR ("Medicine-related belief**" OR "Medicine-related belief**") OR ("patient's perspective" OR "patient's perspective") OR ("patients perspective" OR "patients perspective") OR ("patient perspective" OR "patient perspective") OR ("patients' perspective" OR "patients' perspective")

perspective*) OR "Anticipation,
Psychological" OR expectation* OR (((expectancy OR expectancy) OR (expectancies OR expectancies)) NOT ("Life
Expectancy" OR ("Life Expectancy" OR "Life Expectancy") OR ("Life Expectan**" OR "Life Expectan**")))) OR (("Placebo Effect
belief" OR "Placebo Effect belief") OR ("Placebo Effects belief" OR "Placebo Effects belief") OR ("Placebo Response
belief" OR "Placebo Response belief") OR ("Placebo Responses belief" OR "Placebo Responses belief") OR ("nocebo
belief" OR "nocebo belief") OR ("placebo induced belief" OR "placebo induced belief") OR ("bogus belief" OR "bogus
belief") OR ("null treatment belief" OR "null treatment belief") OR ("inactive treatment belief" OR "inactive treatment
belief") OR ("Placebo Effect beliefs" OR "Placebo Effect beliefs") OR ("Placebo Effects beliefs" OR "Placebo Effects
beliefs") OR ("Placebo Response beliefs" OR "Placebo Response beliefs") OR ("Placebo Responses beliefs" OR "Placebo
Responses beliefs") OR ("nocebo beliefs" OR "nocebo beliefs") OR ("placebo induced beliefs" OR "placebo induced
beliefs") OR ("bogus beliefs" OR "bogus beliefs") OR ("null treatment beliefs" OR "null treatment beliefs") OR ("inactive
treatment beliefs" OR "inactive treatment beliefs")) OR (Communication AND "Physician-Patient
Relations" AND Physicians) OR (("patient mindset" OR "patient mindset") OR ("patients mindset" OR "patients
mindset") OR ("patient's mindset" OR "patient's mindset") OR ("patient mindsets" OR "patient mindsets")) OR "Nocebo
Effect" OR ("Placebo Effect" OR Placebo OR nocebo) AND ("side effect" OR "side
effects" OR analges* OR hyperalges*)) OR ("Pain Perception" AND "Patient Education as Topic"));ti

(conference abstract OR meeting abstract OR conference proceeding OR conference proceedings):pt

PsycINFO

<http://search.ebscohost.com/login.aspx?authtype=ip.uid&profile=lumc&defaultdb=psyh>

((DE"Placebo Effect" OR (TI"Placebo Effect" OR DE"Placebo Effect") OR (TI"Placebo Effects" OR DE"Placebo
Effects") OR (TI"Placebo Effect*" OR DE"Placebo Effect*)) OR (TI"Placebo Response" OR DE"Placebo
Response") OR (TI"Placebo Responses" OR DE"Placebo Responses") OR (TI"Placebo Respon**" OR DE"Placebo
Respon**") OR DE"Placebo Effect" OR (TI"Placebo Effect" OR DE"Placebo Effect") OR (TI"Placebo Effects" OR DE"Placebo
Effects") OR (TI"Placebo Effect*" OR DE"Placebo Effect*)) OR (TI"Placebo Response" OR DE"Placebo Response") OR (TI"Placebo
Responses" OR DE"Placebo Responses") OR (TI"Placebo Respon**" OR DE"Placebo
Respon**") OR (TI"nocebo OR ABnocebo) OR (TI"nocebo*" OR ABnocebo*) OR (TI"placebo induced" OR DE"placebo
induced") OR (TI"bogus OR ABbogus) OR (TI"null treatment" OR DE"null treatment") OR (TI"inactive treatment" OR DE"inactive
treatment")) AND ((DE(Communication) AND DE"Physician-Patient
Relations" OR (DE(Communication) AND DE(Physicians) AND DE(Patients)) OR (TI"Communicate
Patient" OR DE"Communicate Patient") OR (TI"Communicate Patients" OR DE"Communicate Patients") OR (TI"Communicated
Patient" OR DE"Communicated Patient") OR (TI"Communicated Patients" OR DE"Communicated
Patients") OR (TI"Communicating Patient" OR DE"Communicating Patient") OR (TI"Communicating
Patients" OR DE"Communicating Patients") OR (TI"Communication Patient" OR DE"Communication
Patient") OR (TI"Communication Patients" OR DE"Communication Patients") OR (TI"Inform Patient" OR DE"Inform
Patient") OR (TI"Inform Patients" OR DE"Inform Patients") OR (TI"Information Patient" OR DE"Information Patient") OR (TI"Information
Patients" OR DE"Information Patients") OR (TI"Informing Patient" OR DE"Informing Patient") OR (TI"Informing
Patients" OR DE"Informing Patients") OR (TI"Communicate Individual" OR DE"Communicate Individual") OR (TI"Communicate
Individuals" OR DE"Communicate Individuals") OR (TI"Communicated Individual" OR DE"Communicated
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Individuals") OR (TI"Communication Individual" OR DE"Communication Individual") OR (TI"Communication
Individuals" OR DE"Communication Individuals") OR (TI"Inform Individual" OR DE"Inform Individual") OR (TI"Inform
Individuals" OR DE"Inform Individuals") OR (TI"Information Individual" OR DE"Information Individual") OR (TI"Information
Individuals" OR DE"Information Individuals") OR (TI"Informing Individual" OR DE"Informing Individual") OR (TI"Informing
Individuals" OR DE"Informing Individuals") OR (TI"Explain Patient" OR DE"Explain Patient") OR (TI"Explain
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Patients" OR DE"Explaining Patients") OR (TI"Explanation Patient" OR DE"Explanation Patient") OR (TI"Explanation
Patients" OR DE"Explanation Patients") OR (TI"Explanations Patient" OR DE"Explanations Patient") OR (TI"Explanations
Patients" OR DE"Explanations Patients") OR (TI"Explain Individual" OR DE"Explain Individual") OR (TI"Explain
Individuals" OR DE"Explain Individuals") OR (TI"Explaining Individual" OR DE"Explaining Individual") OR (TI"Explaining
Individuals" OR DE"Explaining Individuals") OR (TI"Explanation Individual" OR DE"Explanation Individual") OR (TI"Explanation
Individuals" OR DE"Explanation Individuals") OR (TI"Explanations Individual" OR DE"Explanations
Individuals" OR (TI"Explanations Individuals" OR DE"Explanations Individuals" OR TI" language" OR DE" language" OR TI"
Communications Media" OR DE" Communications Media"))) OR ((TI"beliefs about

medicines" OR TI"expectation*" OR TI"improvement expect**" OR TI"Medicine-related belief**" OR TI"Medicine-related beliefs" OR TI"mindset OR TI"negative belief**" OR TI"negative beliefs" OR TI"negative expectat**" OR TI"patient belief**" OR TI"patient beliefs" OR TI"patient expectation**" OR TI"patient perspective" OR TI"patients' belief**" OR TI"patients' beliefs" OR TI"patients expectation**" OR TI"patients' expectation**" OR TI"patients perspective" OR TI"patients' perspective" OR TI"patient's perspective" OR TI"patients trust" OR TI"patients' trust" OR TI"patient's trust" OR TI"perceived effectiveness" OR TI"perceived efficacy" OR TI"positive belief" OR TI"positive belief**" OR TI"positive expectat**" OR TI"respondent expectation**" OR TI"respondents expectation**" OR TI"respondents' expectation**" OR TI"respondents' expectation**" OR TI"(suggestibility) OR TI"treatment expect**" OR TI"beliefs about medicines" OR TI"improvement expectation" OR TI"improvement expectations" OR TI"Medicine-related belief" OR TI"Medicine-related beliefs" OR TI"negative belief" OR TI"negative beliefs" OR TI"negative expectation" OR TI"negative expectations" OR TI"patient belief" OR TI"patient beliefs" OR TI"patient expectation" OR TI"patient expectations" OR TI"patient perspective" OR TI"patient perspectives" OR TI"patients belief" OR TI"patients beliefs" OR TI"patients expectation" OR TI"patients expectations" OR TI"patients perspective" OR TI"patients perspectives" OR TI"patients trust" OR TI"patient trust" OR TI"perceived effectiveness" OR TI"perceived efficacy" OR TI"positive belief" OR TI"positive beliefs" OR TI"positive expectation" OR TI"positive expectations" OR TI"respondent expectations" OR TI"respondent expectation" OR TI"respondents expectations" OR TI"respondents expectation" OR TI"positive expectations" OR TI"respondent expectations" OR TI"respondents expectations" OR TI"treatment expectation" OR TI"treatment expectations" OR TI"suggestion**" OR DE"suggestion**" OR ((TI"expectancy OR TI"expectancies) NOT (DE"Life Expectancy" OR (TI"Life Expectancy" OR DE"Life Expectancy") OR (TI"Life Expectan**" OR DE"Life Expectan**")))) AND ((DE"(Communication) AND DE"Physician-Patient Relations") OR (DE"(Communication) AND DE"(Physicians) AND DE"(Patients))) OR TI"Communicate Patient" OR TI"Communicate Patients" OR TI"Communicated Patient" OR TI"Communicated Patients" OR TI"Communicating Patient" OR TI"Communicating Patients" OR TI"Communication Patient" OR TI"Communication Patients" OR TI"Inform Patient" OR TI"Inform Patients" OR TI"Informed Patient" OR TI"Informed Patients" OR TI"Information Patient" OR TI"Information Patients" OR TI"Informing Patient" OR TI"Informing Patients" OR TI"Communicate Individual" OR TI"Communicate Individuals" OR TI"Communicated Individual" OR TI"Communicated Individuals" OR TI"Communicating Individual" OR TI"Communicating Individuals" OR TI"Communication Individual" OR TI"Communication Individuals" OR TI"Inform Individual" OR TI"Inform Individuals" OR TI"Informed Individual" OR TI"Informed Individuals" OR TI"Information Individual" OR TI"Information Individuals" OR TI"Informing Individual" OR TI"Informing Individuals" OR TI"Explain Patient" OR TI"Explain Patients" OR TI"Explaining Patient" OR TI"Explaining Patients" OR TI"Explanation Patient" OR TI"Explanation Patients" OR TI"Explanations Patient" OR TI"Explanations Patients" OR TI"Explain Individual" OR TI"Explain Individuals" OR TI"Explaining Individual" OR TI"Explaining Individuals" OR TI"Explanation Individual" OR TI"Explanation Individuals" OR TI"Explanations Individual" OR TI"Explanations Individuals" OR (TI"treatment dropout" OR DE"treatment dropout") OR (TI"treatment dropouts" OR DE"treatment dropouts") OR (TI"therapy dropout" OR DE"therapy dropout") OR (TI"therapy dropouts" OR DE"therapy dropouts" OR DE"nonverbal communication" OR DE"health communication" OR TI"nonverbal communication" OR TI"health communication")) OR ((DE"Placebo Effect" OR (TI"Placebo Effect" OR DE"Placebo Effect") OR (TI"Placebo Effects" OR DE"Placebo Effects") OR (TI"Placebo Effect**" OR DE"Placebo Effect**") OR (TI"Placebo Response" OR DE"Placebo Response") OR (TI"Placebo Responses" OR DE"Placebo Responses") OR (TI"Placebo Respon**" OR DE"Placebo Respon**") OR DE"Nocebo Effect" OR (TI"Nocebo Effect" OR DE"Nocebo Effect") OR (TI"Nocebo Effects" OR DE"Nocebo Effects") OR (TI"Nocebo Effect**" OR DE"Nocebo Effect**") OR (TI"Nocebo Response" OR DE"Nocebo Response") OR (TI"Nocebo Responses" OR DE"Nocebo Responses") OR (TI"Nocebo Respon**" OR DE"Nocebo Respon**") OR (TI"nocebo OR ABnocebo) OR (TI"nocebo*" OR ABnocebo*) OR (TI"placebo induced" OR DE"placebo induced") OR (TI"bogus OR ABbogus) OR (TI>null treatment" OR DE>null treatment") OR (TI"inactive treatment" OR DE"inactive treatment") OR (TI"positive expectat**" OR DE"positive expectat**") OR (TI"negative expectat**" OR DE"negative expectat**") OR (TI"treatment expect**" OR DE"treatment expect**") OR (TI"patients' expectation**" OR DE"patients' expectation**") OR (TI"patients expectation**" OR DE"patients expectation**") OR (TI"patient expectation**" OR DE"patient expectation**") OR (TI"respondents' expectation**" OR DE"respondents' expectation**") OR (TI"respondents expectation**" OR DE"respondents expectation**") OR (TI"respondent expectation**" OR DE"respondent expectation**") OR (TI"improvement expect**" OR DE"improvement expect**") OR (TI"mindset OR ABmindset) OR (TI"patient's trust" OR DE"patient's trust") OR (TI"patients trust" OR DE"patients trust") OR (TI"patients' trust" OR DE"patients' trust") OR (TI"perceived efficacy" OR DE"perceived efficacy") OR (TI"perceived effectiveness" OR DE"perceived effectiveness") OR (TI"suggestibility OR ABSuggestibility) OR (TI"patient beliefs" OR DE"patient beliefs") OR (TI"patient belief**" OR DE"patient belief**") OR (TI"patients' beliefs" OR DE"patients' beliefs") OR (TI"patients' belief**" OR DE"patients' belief**") OR (TI"negative beliefs" OR DE"negative beliefs") OR (TI"negative belief**" OR DE"negative belief**") OR (TI"positive belief" OR DE"positive belief") OR (TI"positive belief**" OR DE"positive belief**") OR (TI"beliefs about medicines" OR DE"beliefs about medicines") OR (TI"Medicine-related beliefs" OR DE"Medicine-related beliefs") OR (TI"Medicine-related belief**" OR DE"Medicine-related belief**") OR (TI"patient's perspective" OR DE"patient's perspective") OR (TI"patients perspective" OR DE"patients perspective") OR (TI"patient perspective" OR DE"patient perspective") OR (TI"patients' perspective" OR DE"patients' perspective")) NOT (DE"Life Expectancy" OR (TI"Life Expectancy" OR DE"Life Expectancy") OR (TI"Life Expectan**" OR DE"Life Expectan**")) AND ((TI"information frame" OR DE"information frame") OR (TI"information frames" OR DE"information

frames*) OR (TI'information framing* OR DE'information framing*) OR (TI'information fram** OR DE'information fram**) OR (TI'message frame* OR DE'message frame*) OR (TI'message frames* OR DE'message frames*) OR (TI'message framing* OR DE'message framing*) OR (TI'message fram** OR DE'message fram**) OR (TI'frame effect* OR DE'frame effect*) OR (TI'frame effects* OR DE'frame effects*) OR (TI'positive frame* OR DE'positive frame*) OR (TI'positive frames* OR DE'positive frames*) OR (TI'positive framing* OR DE'positive framing*) OR (TI'positive fram** OR DE'positive fram**) OR (TI'negative frame* OR DE'negative frame*) OR (TI'negative frames* OR DE'negative frames*) OR (TI'negative framing* OR DE'negative framing*) OR (TI'negative fram** OR DE'negative fram**) OR (TI'positively framed* OR DE'positively framed*) OR (TI'positively fram** OR DE'positively fram**) OR (TI'negatively framed* OR DE'negatively framed*) OR (TI'negatively fram** OR DE'negatively fram**) OR (TI'information frame* OR 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DE'Placebo Responses*) OR (TI'Placebo Respon** OR DE'Placebo Respon**) OR DE'Nocebo Effect* OR (TI'Nocebo Effect* OR DE'Nocebo Effect*) OR (TI'Nocebo Effects* OR DE'Nocebo Effects*) OR (TI'Nocebo Effect** OR DE'Nocebo Effect**) OR (TI'Nocebo Response* OR DE'Nocebo Response*) OR (TI'Nocebo Responses* OR DE'Nocebo Responses*) OR (TI'Nocebo Respon** OR DE'Nocebo Respon**) OR (TI'nocebo OR ABnocebo) OR (TI'nocebo* OR ABnocebo*) OR (TI'placebo induced* OR DE'placebo induced*) OR (TI'bogus OR ABbogus) OR (TI'null treatment* OR DE>null treatment*) OR (TI'inactive treatment* OR DE'inactive treatment*) OR (TI'expectation interventions* OR DE'expectation interventions*) OR (TI'expectation intervention* OR DE'expectation intervention*) OR (TI'positive expectat** OR DE'positive expectat**) OR (TI'negative expectat** OR DE'negative expectat**) OR (TI'treatment expect** OR DE'treatment expect**) OR (TI'patients' expectation** OR DE'patients' expectation**) OR (TI'patients expectation** OR 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Appendix B. Adjusted Grading Criteria.

			Class of Recommendation (COR)			
			Class I	Class IIa	Class IIb	Class III
			Benefit >>> Risk	Benefit >> Risk	Benefit ≥ Risk	No benefit or harmful
			<i>Should be done / is useful</i>	<i>Reasonable to do / can be useful</i>	<i>May be considered / unknown usefulness</i>	<i>Not helpful or harmful</i>
Level (Quality) Of Evidence (LOE)	A	<i>High quality data, > 1 RCT</i>	+++++	+++	+++	----
	B-R (randomized)	<i>Moderate quality data, >1 RCT</i>	++++	++	++	---
	B-NR (non-randomized)	<i>Moderate quality evidence from > 1 high quality nonrandomized/observational study</i>	+++	+	+	--
	C-LD (limited data)	<i>(non)randomized studies with limitations in design, mechanistic studies</i>	++	+	0	-
	C-EO (expert opinion)	<i>Consensus opinion based on clinical experience and expertise</i>	+	+	0	-

Note. Grading is adjusted from the American College of Cardiology and American Heart Association "Class of Recommendations and Level (Quality) of Evidence Ratings" system¹⁷. Because the current recommendation paper did not systematically search for systematic reviews and meta-analyses, these criteria were omitted from the evaluation of the level of evidence.

Appendix C. Supplementary Tables

Supplementary Table S1. Overview of the literature backing each clinical recommendation.

Category	Recommendation	Do's and don'ts	References		Systematic review Study type ratio (quantitative / qualitative / mixed)
			First stream	Second stream	
Optimizing the clinician-patient relationship	Show interest in the patient	<ul style="list-style-type: none"> ✓ Use active listening and personal attention to foster trust and enhance patient satisfaction ✓ Ensure that a patient doesn't feel regarded as just a "number", a "body" or a "case" ✓ Use respectful and caring behaviours, displaying honesty, respect and competence to build trust and foster positive interactions ✗ Avoid misjudgements of patients' unmet needs 	19,28,29	30-46	10 / 5 / 2
	Give reassurance about ongoing care	<ul style="list-style-type: none"> ✓ Use brief reassuring statements to reduce insecurity and fears of abandonment, e.g. "we're going to take good care of you" ✓ Establish a cooperative clinician-patient relationship grounded in shared goals 	47,48	49-52	2 / 2 / 0
	Respond to negative emotions	<ul style="list-style-type: none"> ✓ Respond to negative emotions, e.g., by naming and understanding them ✓ Provide emotional support and address the emotional needs of patients 	28,53,54	55-67	8 / 5 / 0
	Leverage eye contact and physical presence	<ul style="list-style-type: none"> ✓ Use attentive listening and maintain a patient-oriented posture to foster trust ✓ Make eye contact with the patient while listening or providing information ✓ Sit rather than stand during consultations, as this will cause patients to perceive as longer and more positive ✓ Use appropriate facial expressions to increase perceived trustworthiness and competence ✗ Avoid excessive eye contact, as this can diminish trust 	21,68-72	19,21,68,73 -80	9 / 2 / 0

Use patient-centered
communication

89-113

22 / 3 / 0

*Note. * With 'overall recommendation', we refer to the evidence for the overall domain (e.g., 'optimizing the clinician-patient relationship' or 'asking about expectations').*

Supplementary Table S1. *Continued*

Category	Recommendation	Do's and don'ts	References		Systematic review
			First stream	Second stream	Study type ratio (quantitative / qualitative / mixed)
Optimizing the clinician-patient relationship	Adapt communication style to the patient	<ul style="list-style-type: none"> ✓ Adapt your communication to the patient's values and expectations, comprehension levels and information needs ✓ Consider whether it is always helpful to give advice (e.g., when patients are highly distressed) 		81-88	6 / 2 / 0
	Leverage other behaviours to foster trust	<ul style="list-style-type: none"> ✓ Provide clear explanations and use shared decision making ✓ Address prior negative encounters that the patient may have had ✓ Foster trust, a sense of control and autonomy, and improve cultural understanding 		114-131	15 / 2 / 1
<i>Overall recommendation*</i>			12,19- 27,277		72 / 21 / 3

Supplementary Table S1. *Continued*

Category	Recommendation	Do's and don'ts	References		Systematic review Study type ratio (quantitative / qualitative / mixed)
			First stream	Second stream	
Asking about expectations	Ask about and identify general expectations towards treatment	<ul style="list-style-type: none"> ✓ Incorporate questions about expectations in the natural flow of the conversation, e.g., integrate them before explaining a specific treatment part ✓ Use open-ended questions to ask about patients' prior knowledge and experiences ✓ Encourage patients to articulate their expectations ✗ Avoid leading questions and filling in the blanks for patients when they have trouble expressing what they expect 	2,3,139-141		
	Adjust unrealistic expectations	<ul style="list-style-type: none"> ✓ If patients are overly optimistic, clarify the limitations of the treatment ✓ If patients are overly pessimistic, focus on potential long-term effects to recalibrate their outlook ✓ Check whether patients expect any tests or treatments that are not aligned with existing guidelines and therapies ✓ Take patient beliefs seriously, even if they appear unrealistic or unfounded ✓ Have an open and honest attitude towards patients' expectations, to help them understand treatment suggestions and form realistic expectations 	2,132,141,148,149		
	Address disappointment from previous treatments	<ul style="list-style-type: none"> ✓ Identify negative expectations that may be due to previous disappointments with treatment ✓ Emphasize the differences between the current treatment option and those prior negative experiences ✓ Emphasize the potential effectiveness of the current treatment ✓ If possible, offer alternate routes of administration that resonate with the patient's preferences 	2,154		



*Overall recommendation**

2,132-138

142-147,150-

153

6 / 4 / 0

Supplementary Table S1. *Continued*

Category	Recommendation	Do's and don'ts	References		Systematic review
			First stream	Second stream	Study type ratio (quantitative / qualitative / mixed)
Discussing the treatment rationale	Provide correct information based on evidence-based knowledge	<ul style="list-style-type: none"> ✓ Provide accurate and up-to-date information about the treatment and its expected effects, that is grounded in medical knowledge and the clinician's experience ✓ Provide clear and comprehensive explanations to improve decision making ✓ Use explanatory information and discuss biomedical information ✓ Balance optimism with a clear discussion of pessimistic aspects (e.g., a bad prognosis), while being emotionally sensitive ✓ Approach sensitive topics more indirectly ✗ Do not avoid talking about psychosocial issues and offering emotional support if psychosocial problems are present ✗ Avoid medical terms that may induce negative expectations. If possible, use euphemisms instead. 	138,161	162-167	6 / 0 / 0
	Avoid exaggeration or underestimation of treatment results	<ul style="list-style-type: none"> ✓ Give an accurate presentation of treatment outcomes to maintain credibility and trust ✓ Transparently communicate about uncertainty, but use implicit phrasing (e.g., "most likely diagnoses") rather than direct statements of uncertainty to maintain trust ✗ Do not exaggerate or underestimate treatment results 	2,159	156,168-182,278	15 / 1 / 1
	Clearly state the treatment rationale and expected positive outcomes in concrete terms	<ul style="list-style-type: none"> ✓ Explicitly state the treatment rationale and the expected positive benefits ✓ Supply this statement with tangible details about the predicted outcomes, to help align patient expectations with realistic treatment outcomes ✓ Use positive and gain-framed messages, and use positive affect-oriented speech to reduce anxiety and improve the willingness to receive a treatment 	159,183	184-188	2 / 2 / 1

- ✘ Do not omit important biomedical details, e.g. on symptom causes and severity, and test results

Supplementary Table S1. *Continued*

Category	Recommendation	Do's and don'ts	References		Systematic review
			First stream	Second stream	Study type ratio (quantitative / qualitative / mixed)
Discussing the treatment rationale	Explain treatment mechanisms of action	<ul style="list-style-type: none"> ✓ Explain what the treatment does in the body clearly and vividly to increase confidence in its effectiveness ✓ Increase patient understanding by explaining mechanisms of treatment, e.g. on procedural information to reduce anxiety or on drug approval processes to foster confidence in (generic) medicine safety and quality ✓ Consider the patient's (medical) history, physical examinations, and prior research in explaining the mechanisms of action 	2,8,161	161,189,190	3 / 0 / 0
	Align patient expectations and beliefs with evidence-based care	<ul style="list-style-type: none"> ✓ Ask what a patient personally expects from a treatment and align this with realistic outcomes and evidence-based care ✓ If no treatment is prescribed, express confidence and positive expectations about the course of the symptoms or the patient's ability to cope with them ✓ Balance information with acceptance or denial of unrealistic expectations, using medical knowledge and expertise to challenge these expectations 	10,160,161,191	192-196	4 / 1 / 0
	Emphasize the purpose and advantages of treatment procedures	<ul style="list-style-type: none"> ✓ Focus on the purpose and advantages of medical procedures, rather than explicitly mentioning potential pain or discomfort ✓ Use case descriptions, storytelling, reassurance, clear explanations and describing favourable prognoses to highlight potential benefits of treatment and medical procedures 	2,5,197,198	199-202	4 / 0 / 0
	Check patient comprehension	<ul style="list-style-type: none"> ✓ Ensure that patients fully comprehend the information provided during medical consultations and verify that they have a solid understanding of their condition and the treatment that they will receive ✓ Promptly correct and adjust any apparent misconceptions or misunderstanding 	2,138,156,203,204	205-207	1 / 1 / 1



Supplementary Table S1. *Continued*

Category	Recommendation	Do's and don'ts	References		Systematic review
			First stream	Second stream	Study type ratio (quantitative / qualitative / mixed)
Discussing the treatment rationale	Suggest reliable information sources	<ul style="list-style-type: none"> ✓ Provide patients with additional resources, including leaflets and reliable websites ✓ Encourage health seeking behaviours and support patients with recommending credible sources, or gently correct them if unreliable sources are used 	208-210	211-215	6 / 0 / 0
<i>Overall recommendation*</i>			6,10,155-161,197	.	41 / 5 / 3

Supplementary Table S1. *Continued*

Category	Recommendation	Do's and don'ts	References		Systematic review
			First stream	Second stream	Study type ratio (quantitative / qualitative / mixed)
Discussing side effects	Emphasize positive consequences of side effects	<ul style="list-style-type: none"> ✓ Change patients' mindsets about side effects, e.g. encourage them to reinterpret minor, non-harmful side effects as signals that the treatment may be working (e.g., for vaccinations, allergen desensitization) ✗ Do not encourage patients to reinterpret side effects as positive when they may be harmful, signal treatment intolerance or if alternatives exist 	219	220,221	2 / 0 / 0
	General structure of the consultation	<ul style="list-style-type: none"> ✓ If not harmful, indicate that side effects may be unpleasant but that the treatment will have benefits on the longer term ✓ Frame information positively to enhance patient confidence and treatment adherence ✓ Address anxiety before providing information about treatment side effects ✓ Provide a support system for patients should side effects occur ✓ Balance negative information with other positive interventions, including but not limited to self-affirmation, or relaxation 	210	222-232	9 / 1 / 0**
	Investigate (sources of) nocebo effects	<ul style="list-style-type: none"> ✓ Carefully check whether the patient had an increased risk of developing nocebo effects, e.g., due to past negative experiences or beliefs about increased sensitivity towards side effects ✓ Openly and without judgement, ask about what a patient expects to experience ✓ Create an atmosphere in which the patient understands that there are no wrong answers to such questions ✓ Help patients to understand numerical information regarding side effects ✗ Avoid leading questions or words to ask about side effect expectations 	210	233-235	3 / 0 / 0

Supplementary Table S1. *Continued*

Category	Recommendation	Do's and don'ts	References		Systematic review
			First stream	Second stream	Study type ratio (quantitative / qualitative / mixed)
	State all symptoms briefly to reduce nocebo effects	<ul style="list-style-type: none"> ✓ State all symptoms briefly to reduce the chance of side effects, particularly for low-risk side effects ✗ Do not focus or highlight specific side effects, unless they are high-risk effects for which actions should be taken 	236,237	229,238	2 / 0 / 0
	Frame positive outcomes rather than negative outcomes	<ul style="list-style-type: none"> ✓ Frame information about side effects positively (i.e., 9 out of 10 do not experience side effects) 	198,210,217	239,240	2 / 0 / 0
Information framing	Discuss risks in statistical terms	<ul style="list-style-type: none"> ✓ Use statistical terms to describe risks, as these do not appeal to the imagination as much as personal terms (e.g., 1 in 10 people) ✓ Use simplified and risk-neutral phrasing where possible ✓ In serious illness, absolute mortality risk framing may be preferred to reduce bias and improve informed decision making ✗ Avoid descriptors such as 'very common' and 'common' as these can result in negative expectations 	241	234,242,243	3 / 0 / 0
	Use neutral terms when performing a procedure	<ul style="list-style-type: none"> ✓ When performing a procedure, use neutral terms rather than terms that can indicate discomfort or pain evoked by the procedure 		197	1 / 0 / 0
<i>Overall recommendation*</i>			7,210,216-218,237		22 / 1 / 0

Supplementary Table S1. *Continued*

Category	Recommendation	Do's and don'ts	References		Systematic review
			First stream	Second stream	Study type ratio (quantitative / qualitative / mixed)
Explain placebo and nocebo effects	Careful explanation of placebo effects	<ul style="list-style-type: none"> ✓ First explain the treatment mechanisms of action to increase understanding of the intertwined roles of the mind and brain in treatment efficacy ✓ Communicate clearly that treatments can still be efficacious even when patients are informed about placebo effects ✓ Carefully explain the concept of "placebo effects", using this term judiciously especially for patients with limited health literacy skills 	14,244	252-263	12 / 0 / 1
	Mechanism of action of placebo effects	<ul style="list-style-type: none"> ✓ Explain that the placebo effect arises due to positive expectations and that this is a real response of the body that can alleviate symptoms ✓ Provide examples that help patients gain a better understanding of how placebo effects work 	14,264	265-268	3 / 1 / 0
	Appropriate presentation of information	<ul style="list-style-type: none"> ✓ Tailor the explanation to the individual that receives it ✓ Use additional terminology or euphemisms to explain the mechanisms of placebo effects 	14,264	269-271	3 / 0 / 0
	Prevent overly optimistic expectations	<ul style="list-style-type: none"> ✓ Prevent the shaping of overly optimistic expectations about placebo effects ✓ Clarify that placebo effects can alleviate symptoms, but that they cannot cure illness ✗ Do not give the impression that placebo is preferable over medical treatment or that it can replace conventional treatment 	14		

Supplementary Table S1. *Continued*

Category	Recommendation	Do's and don'ts	References		Systematic review
			First stream	Second stream	Study type ratio (quantitative / qualitative / mixed)
Explain placebo and nocebo effects	Attend to the purpose of explaining nocebo effects	<ul style="list-style-type: none"> ✓ Carefully consider if a patient should be informed about nocebo effects ✓ Emphasize the reasons for why information about nocebo effects is provided, for instance, to reduce them from occurring or to further patient understanding 	14,210,272	.	.
	Careful explanation of nocebo effects	<ul style="list-style-type: none"> ✓ Carefully explain nocebo effects, using alternate or related terms if deemed more appropriate, such as anxiety or fear 	14,210	247,273-275	4 / 0 / 0
	Keep explanation simple	<ul style="list-style-type: none"> ✓ Keep the explanation of nocebo effects short and simple, if possible, for instance, explaining that side effects or symptoms may be more readily noticed if a patient focuses their attention on them 	14	.	.
	Nocebo effects	<ul style="list-style-type: none"> ✓ Ensure that it is understood that not all side effects are nocebo effects and that the patient should report them; sometimes they are real side effects that require prompt action from the clinician 			
	Avoid unintentional negative effects	<ul style="list-style-type: none"> ✓ If (nocebo) side effects are experienced, discuss and decide about follow-up steps together with the patient ✗ When explaining nocebo effects, avoid unintentionally placing blame or responsibility for experiencing nocebo effects on the patient 	14	.	.
	Provide ways to manage nocebo effects	<ul style="list-style-type: none"> ✓ Suggest that family members read the information leaflet to keep an eye out for side effects ✓ Suggest relaxation or distraction techniques to keep attention away from side effects ✓ Combine relaxation and distraction techniques with nocebo education to manage these effects and empower patients 	2	276	0 / 1 / 0



*Overall recommendation**

11,13,14,210,2

44-251

22 / 2 / 1

Total: 163 / 33 / 7

Supplementary Table S2

Please go to this link:

<https://docs.google.com/spreadsheets/d/1qa2PPjFWOV1j6sflkcojz96DGodbts2z/edit?usp=sharing&oid=107410726990848575949&rtpof=true&sd=tru>
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Supplementary Table S3 (also presented in main manuscript as Table 2): NURSE framework for empathic behaviours, developed on the basis of the studies by Pollak et al.²⁷, Back et al.²⁶ and van Vliet & Epstein¹². Note that the examples serve only as an inspiration^{23,277}.

NURSE	Explanation	Example
Naming	Name the emotion	You sound very anxious.
Understanding	Show understanding of the emotion	I really understand that you're dreading it. I can imagine that you're upset about it.
Respecting	Show respect for the patient	I'm impressed with how well you're handling the situation.
Supporting	Express support	And whatever happens, we'll take good care of you. You don't have to face this alone.
Exploring	Explore the emotion	What are you thinking about now? You're suddenly having to face so much.



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