

NEJM BEMPEDOIC ACID

NEJM BEMPEDOIC ACID HAS EMERGED AS A SIGNIFICANT TOPIC WITHIN THE REALM OF CARDIOVASCULAR PHARMACOTHERAPY, PARTICULARLY IN THE TREATMENT OF HYPERCHOLESTEROLEMIA. THIS ARTICLE EXPLORES THE COMPREHENSIVE DETAILS SURROUNDING BEMPEDOIC ACID, ITS CLINICAL IMPLICATIONS AS REPORTED IN THE NEW ENGLAND JOURNAL OF MEDICINE (NEJM), AND ITS MECHANISMS OF ACTION. AS A NOVEL CHOLESTEROL-LOWERING AGENT, BEMPEDOIC ACID HAS SHOWN PROMISE IN MANAGING PATIENTS WHO ARE INTOLERANT TO STATINS OR REQUIRE ADDITIONAL LDL CHOLESTEROL REDUCTION. THE DISCUSSION INCLUDES CLINICAL TRIAL EVIDENCE, SAFETY PROFILES, AND ITS ROLE IN CONTEMPORARY LIPID MANAGEMENT STRATEGIES. ADDITIONALLY, THIS ARTICLE WILL EXAMINE THE PHARMACOLOGICAL PROPERTIES AND POTENTIAL BENEFITS OF BEMPEDOIC ACID, SUPPORTED BY NEJM FINDINGS. TO PROVIDE A STRUCTURED OVERVIEW, A TABLE OF CONTENTS FOLLOWS TO GUIDE THE READER THROUGH THE DETAILED SECTIONS.

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- COMPARATIVE EFFECTIVENESS WITH OTHER LIPID-LOWERING AGENTS

CLINICAL BACKGROUND OF BEMPEDOIC ACID

BEMPEDOIC ACID IS AN ORAL LIPID-LOWERING AGENT DEVELOPED TO ADDRESS UNMET NEEDS IN CHOLESTEROL MANAGEMENT, PARTICULARLY FOR PATIENTS WHO DO NOT ACHIEVE TARGET LDL CHOLESTEROL LEVELS WITH STATINS ALONE OR FOR THOSE WHO ARE STATIN INTOLERANT. THE DEVELOPMENT OF BEMPEDOIC ACID REFLECTS ADVANCES IN CARDIOVASCULAR MEDICINE AIMED AT REDUCING ATHEROSCLEROTIC CARDIOVASCULAR DISEASE (ASCVD) RISK THROUGH NOVEL MECHANISMS. SINCE HYPERCHOLESTEROLEMIA REMAINS A MAJOR MODIFIABLE RISK FACTOR FOR CARDIOVASCULAR EVENTS, NEW THERAPIES LIKE BEMPEDOIC ACID HAVE GAINED ATTENTION FOLLOWING PUBLICATION IN AUTHORITATIVE JOURNALS SUCH AS THE NEW ENGLAND JOURNAL OF MEDICINE (NEJM).

HISTORICAL CONTEXT AND FDA APPROVAL

THE INTRODUCTION OF BEMPEDOIC ACID INTO CLINICAL PRACTICE FOLLOWED COMPREHENSIVE RESEARCH AND REGULATORY EVALUATION. IT RECEIVED FDA APPROVAL BASED ON PIVOTAL PHASE 3 CLINICAL TRIALS THAT DEMONSTRATED ITS EFFICACY IN LOWERING LDL CHOLESTEROL WHEN USED ALONE OR IN COMBINATION WITH OTHER LIPID-LOWERING THERAPIES. THE NEJM PUBLISHED KEY STUDIES HIGHLIGHTING ITS CARDIOVASCULAR BENEFITS AND SAFETY PROFILE. THIS APPROVAL PROVIDED AN IMPORTANT ALTERNATIVE FOR PATIENTS INADEQUATELY MANAGED BY EXISTING THERAPIES.

PREVALENCE OF HYPERCHOLESTEROLEMIA AND NEED FOR NEW TREATMENTS

DESPITE WIDESPREAD USE OF STATINS, A SIGNIFICANT PROPORTION OF PATIENTS WITH HYPERCHOLESTEROLEMIA REMAIN AT ELEVATED CARDIOVASCULAR RISK DUE TO INSUFFICIENT LDL CHOLESTEROL REDUCTION OR INTOLERANCE TO TREATMENT. BEMPEDOIC ACID ADDRESSES THESE CHALLENGES BY OFFERING AN ADDITIONAL MECHANISM TO DECREASE LDL CHOLESTEROL LEVELS. THIS COMPLEMENTS EXISTING THERAPIES AND BROADENS TREATMENT OPTIONS FOR CLINICIANS MANAGING DYSLIPIDEMIA.

MECHANISM OF ACTION AND PHARMACOLOGY

BEMPEDOIC ACID FUNCTIONS THROUGH A DISTINCT PATHWAY THAT DIFFERENTIATES IT FROM STATINS AND OTHER LIPID-LOWERING AGENTS. UNDERSTANDING ITS MECHANISM OF ACTION IS CRUCIAL FOR APPRECIATING ITS THERAPEUTIC ROLE AND

POTENTIAL ADVANTAGES.

INHIBITION OF ATP CITRATE LYASE

THE PRIMARY MECHANISM OF BEMPEDOIC ACID IS INHIBITION OF ATP CITRATE LYASE (ACL), AN ENZYME UPSTREAM OF HMG-CoA REDUCTASE IN THE CHOLESTEROL BIOSYNTHESIS PATHWAY. BY TARGETING ACL, BEMPEDOIC ACID REDUCES CHOLESTEROL SYNTHESIS IN THE LIVER, LEADING TO UPREGULATION OF LDL RECEPTORS AND ENHANCED CLEARANCE OF LDL CHOLESTEROL FROM THE BLOODSTREAM. THIS NOVEL APPROACH COMPLEMENTS STATINS, WHICH INHIBIT HMG-CoA REDUCTASE DIRECTLY.

PRODRUG ACTIVATION AND TISSUE SPECIFICITY

BEMPEDOIC ACID IS A PRODRUG THAT REQUIRES ACTIVATION BY VERY-LONG-CHAIN ACYL-CoA SYNTHETASE 1 (ACSVL1), AN ENZYME EXPRESSED PREDOMINANTLY IN THE LIVER BUT ABSENT IN SKELETAL MUSCLE. THIS SELECTIVE ACTIVATION LIMITS THE POTENTIAL FOR MUSCLE-RELATED SIDE EFFECTS COMMONLY ASSOCIATED WITH STATINS, MAKING BEMPEDOIC ACID A FAVORABLE OPTION FOR PATIENTS EXPERIENCING STATIN INTOLERANCE.

PHARMACOKINETICS AND METABOLISM

THE PHARMACOKINETIC PROFILE OF BEMPEDOIC ACID FEATURES ORAL ADMINISTRATION WITH ONCE-DAILY DOSING, A HALF-LIFE CONDUCTIVE TO STEADY LIPID-LOWERING EFFECTS, AND HEPATIC METABOLISM. ITS ELIMINATION PATHWAYS AND MINIMAL DRUG-DRUG INTERACTIONS CONTRIBUTE TO ITS SUITABILITY IN COMBINATION REGIMENS WITH STATINS OR EZETIMIBE.

NEJM CLINICAL TRIALS AND EVIDENCE

THE NEW ENGLAND JOURNAL OF MEDICINE HAS PUBLISHED SEVERAL LANDMARK STUDIES EVALUATING THE EFFICACY AND SAFETY OF BEMPEDOIC ACID. THESE TRIALS PROVIDE HIGH-QUALITY EVIDENCE SUPPORTING ITS CLINICAL USE.

CLEAR HARMONY AND CLEAR WISDOM TRIALS

TWO PIVOTAL PHASE 3 TRIALS, CLEAR HARMONY AND CLEAR WISDOM, ASSESSED THE LIPID-LOWERING EFFECTS OF BEMPEDOIC ACID IN PATIENTS WITH HYPERCHOLESTEROLEMIA ALREADY RECEIVING MAXIMALLY TOLERATED STATIN THERAPY. BOTH TRIALS DEMONSTRATED STATISTICALLY SIGNIFICANT REDUCTIONS IN LDL CHOLESTEROL COMPARED TO PLACEBO, CONFIRMING THE DRUG'S EFFICACY AS AN ADD-ON TREATMENT.

CARDIOVASCULAR OUTCOMES AND SAFETY DATA

SUBSEQUENT ANALYSES, INCLUDING CARDIOVASCULAR OUTCOMES STUDIES, EVALUATED THE IMPACT OF BEMPEDOIC ACID ON MAJOR ADVERSE CARDIOVASCULAR EVENTS (MACE). WHILE LIPID REDUCTION WAS ROBUST, ONGOING STUDIES AIM TO CLARIFY LONG-TERM CARDIOVASCULAR BENEFITS. SAFETY DATA FROM NEJM PUBLICATIONS HIGHLIGHT A FAVORABLE PROFILE WITH LOW INCIDENCE OF MUSCLE-RELATED ADVERSE EVENTS AND MANAGEABLE SIDE EFFECTS.

IMPLICATIONS FOR CLINICAL PRACTICE

THE NEJM EVIDENCE SUPPORTS THE INTEGRATION OF BEMPEDOIC ACID INTO LIPID MANAGEMENT PROTOCOLS, ESPECIALLY FOR PATIENTS WHO REQUIRE ADDITIONAL LDL LOWERING BEYOND STATINS OR WHO CANNOT TOLERATE STATINS. THESE STUDIES HAVE INFLUENCED GUIDELINES AND THERAPEUTIC DECISION-MAKING WORLDWIDE.

THERAPEUTIC USES AND PATIENT POPULATIONS

BEMPEDOIC ACID IS INDICATED PRIMARILY FOR ADULTS WITH HYPERCHOLESTEROLEMIA OR MIXED DYSLIPIDEMIA REQUIRING FURTHER LDL CHOLESTEROL REDUCTION. ITS SPECIFIC APPLICATIONS REFLECT ITS PHARMACOLOGICAL PROPERTIES AND CLINICAL TRIAL FINDINGS.

PATIENTS WITH STATIN INTOLERANCE

STATIN INTOLERANCE, CHARACTERIZED BY MUSCLE SYMPTOMS OR OTHER ADVERSE EFFECTS, LIMITS EFFECTIVE LDL CHOLESTEROL MANAGEMENT IN MANY PATIENTS. BEMPEDOIC ACID OFFERS A VALUABLE ALTERNATIVE DUE TO ITS LIVER-SPECIFIC ACTIVATION, WHICH MINIMIZES MUSCLE EXPOSURE AND ASSOCIATED SIDE EFFECTS. THIS EXPANDS TREATMENT OPTIONS AND IMPROVES PATIENT ADHERENCE.

COMBINATION THERAPY WITH STATINS AND EZETIMIBE

FOR PATIENTS NOT ACHIEVING LDL CHOLESTEROL TARGETS ON STATINS ALONE, BEMPEDOIC ACID CAN BE COMBINED WITH STATINS OR EZETIMIBE TO ACHIEVE ADDITIVE LDL LOWERING. THIS COMBINATION APPROACH IS SUPPORTED BY CLINICAL TRIAL DATA AND ALIGNS WITH TREATMENT GUIDELINES AIMING FOR INTENSIVE LIPID CONTROL IN HIGH-RISK CARDIOVASCULAR PATIENTS.

POTENTIAL ROLE IN FAMILIAL HYPERCHOLESTEROLEMIA

PATIENTS WITH HETEROZYGOUS FAMILIAL HYPERCHOLESTEROLEMIA (HeFH) OFTEN REQUIRE MULTI-DRUG REGIMENS TO MANAGE ELEVATED LDL CHOLESTEROL. BEMPEDOIC ACID REPRESENTS AN EMERGING OPTION IN THIS GROUP, OFFERING INCREMENTAL LDL REDUCTION WHEN STANDARD THERAPIES ARE INSUFFICIENT.

SAFETY PROFILE AND SIDE EFFECTS

A THOROUGH UNDERSTANDING OF THE SAFETY PROFILE OF BEMPEDOIC ACID IS ESSENTIAL FOR ITS CLINICAL USE. NEJM REPORTS AND POST-MARKETING DATA PROVIDE INSIGHTS INTO ITS TOLERABILITY AND ADVERSE EVENT SPECTRUM.

COMMON ADVERSE EFFECTS

THE MOST FREQUENTLY REPORTED SIDE EFFECTS INCLUDE:

- UPPER RESPIRATORY TRACT INFECTIONS
- MUSCLE SPASMS
- BACK PAIN
- GOUT FLARE-UPS
- ELEVATIONS IN URIC ACID LEVELS

THESE ADVERSE EFFECTS ARE GENERALLY MILD TO MODERATE AND MANAGEABLE WITH APPROPRIATE CLINICAL MONITORING.

MUSCLE-RELATED SAFETY

BEMPEDOIC ACID'S LACK OF ACTIVATION IN SKELETAL MUSCLE REDUCES THE RISK OF MYOPATHY AND RHABDOMYOLYSIS, SIDE EFFECTS COMMONLY ASSOCIATED WITH STATINS. THIS FEATURE IS PARTICULARLY ADVANTAGEOUS FOR PATIENTS WITH PREVIOUS STATIN-ASSOCIATED MUSCLE SYMPTOMS.

CONTRAINDICATIONS AND PRECAUTIONS

BEMPEDOIC ACID SHOULD BE USED CAUTIOUSLY IN PATIENTS WITH A HISTORY OF GOUT OR HYPERURICEMIA DUE TO POTENTIAL INCREASES IN SERUM URIC ACID. ADDITIONALLY, RENAL FUNCTION SHOULD BE MONITORED TO DETECT ANY ADVERSE CHANGES DURING THERAPY.

COMPARATIVE EFFECTIVENESS WITH OTHER LIPID-LOWERING AGENTS

UNDERSTANDING HOW BEMPEDOIC ACID COMPARES TO EXISTING LIPID-LOWERING THERAPIES AIDS CLINICIANS IN SELECTING APPROPRIATE TREATMENT STRATEGIES FOR INDIVIDUAL PATIENTS.

COMPARISON WITH STATINS

WHILE STATINS REMAIN THE CORNERSTONE OF LDL CHOLESTEROL MANAGEMENT, BEMPEDOIC ACID PROVIDES A COMPLEMENTARY MECHANISM AND IS PARTICULARLY USEFUL IN PATIENTS WHO CANNOT TOLERATE STATINS OR REQUIRE ADDITIONAL LDL REDUCTION. ITS LDL-LOWERING EFFECT IS GENERALLY MORE MODEST THAN HIGH-INTENSITY STATINS BUT ADDITIVE WHEN COMBINED.

COMPARISON WITH EZETIMIBE

EZETIMIBE, AN INHIBITOR OF INTESTINAL CHOLESTEROL ABSORPTION, IS ANOTHER COMMONLY USED ADJUNCT TO STATINS. BEMPEDOIC ACID OFFERS A DIFFERENT MECHANISM BY TARGETING HEPATIC CHOLESTEROL SYNTHESIS. COMBINATION THERAPY WITH BOTH AGENTS CAN PRODUCE GREATER LDL CHOLESTEROL REDUCTION THAN EITHER ALONE.

EMERGING THERAPIES AND FUTURE DIRECTIONS

NEWER LIPID-LOWERING AGENTS SUCH AS PCSK9 INHIBITORS PROVIDE POTENT LDL CHOLESTEROL REDUCTION BUT ARE OFTEN LIMITED BY COST AND ADMINISTRATION ROUTE. BEMPEDOIC ACID'S ORAL ADMINISTRATION AND TOLERABILITY POSITION IT AS AN IMPORTANT OPTION AMONG EMERGING THERAPIES FOR DYSLIPIDEMIA MANAGEMENT.

FREQUENTLY ASKED QUESTIONS

WHAT IS BEMPEDOIC ACID AND HOW DOES IT WORK?

BEMPEDOIC ACID IS A LIPID-LOWERING MEDICATION THAT INHIBITS ATP CITRATE LYASE, AN ENZYME INVOLVED IN CHOLESTEROL SYNTHESIS IN THE LIVER, THEREBY REDUCING LDL CHOLESTEROL LEVELS.

WHAT ARE THE KEY FINDINGS ABOUT BEMPEDOIC ACID IN RECENT NEJM STUDIES?

RECENT NEJM STUDIES HAVE SHOWN THAT BEMPEDOIC ACID EFFECTIVELY LOWERS LDL CHOLESTEROL IN PATIENTS WITH HYPERCHOLESTEROLEMIA, INCLUDING THOSE WHO ARE STATIN-INTOLERANT, AND MAY REDUCE CARDIOVASCULAR EVENTS WHEN ADDED TO STANDARD THERAPY.

IS BEMPEDOIC ACID SAFE FOR PATIENTS WHO CANNOT TOLERATE STATINS?

YES, NEJM RESEARCH INDICATES THAT BEMPEDOIC ACID IS GENERALLY WELL TOLERATED AND OFFERS AN ALTERNATIVE LIPID-LOWERING THERAPY FOR PATIENTS WHO EXPERIENCE STATIN-ASSOCIATED MUSCLE SYMPTOMS OR INTOLERANCE.

HOW DOES BEMPEDOIC ACID COMPARE TO STATINS IN LOWERING LDL CHOLESTEROL?

WHILE STATINS REMAIN THE FIRST-LINE TREATMENT, BEMPEDOIC ACID PROVIDES A MODERATE LDL CHOLESTEROL REDUCTION AND CAN BE USED AS AN ADJUNCT OR ALTERNATIVE, PARTICULARLY IN PATIENTS WHO CANNOT TOLERATE HIGH-INTENSITY STATINS.

WHAT ARE THE COMMON SIDE EFFECTS OF BEMPEDOIC ACID REPORTED IN NEJM ARTICLES?

COMMON SIDE EFFECTS INCLUDE INCREASED URIC ACID LEVELS, GOUT, MUSCLE SPASMS, AND ELEVATED LIVER ENZYMES; HOWEVER, THESE ARE GENERALLY INFREQUENT AND MANAGEABLE.

CAN BEMPEDOIC ACID BE COMBINED WITH OTHER LIPID-LOWERING THERAPIES?

YES, BEMPEDOIC ACID HAS BEEN STUDIED IN COMBINATION WITH EZETIMIBE AND STATINS, SHOWING ADDITIVE LDL CHOLESTEROL LOWERING EFFECTS WITHOUT SIGNIFICANT DRUG INTERACTIONS.

WHAT IS THE IMPACT OF BEMPEDOIC ACID ON CARDIOVASCULAR OUTCOMES ACCORDING TO NEJM RESEARCH?

NEJM CLINICAL TRIALS SUGGEST THAT BEMPEDOIC ACID MAY REDUCE THE RISK OF MAJOR ADVERSE CARDIOVASCULAR EVENTS IN HIGH-RISK PATIENTS, ALTHOUGH ONGOING STUDIES CONTINUE TO EVALUATE ITS LONG-TERM BENEFITS.

ADDITIONAL RESOURCES

1. *BEMPEDOIC ACID AND CARDIOVASCULAR HEALTH: EMERGING THERAPIES IN LIPID MANAGEMENT*

THIS BOOK EXPLORES THE ROLE OF BEMPEDOIC ACID AS A NOVEL LIPID-LOWERING AGENT, FOCUSING ON ITS MECHANISM OF ACTION AND CLINICAL EFFICACY. IT PROVIDES A COMPREHENSIVE REVIEW OF CARDIOVASCULAR DISEASE MANAGEMENT, PARTICULARLY FOR PATIENTS WHO ARE STATIN-INTOLERANT OR REQUIRE ADDITIONAL LDL CHOLESTEROL REDUCTION. CASE STUDIES AND RECENT CLINICAL TRIAL DATA ARE INCLUDED TO ILLUSTRATE PRACTICAL APPLICATIONS.

2. *ADVANCES IN LIPID-LOWERING THERAPIES: THE PROMISE OF BEMPEDOIC ACID*

DETAILING THE LATEST ADVANCEMENTS IN LIPID-LOWERING TREATMENTS, THIS BOOK HIGHLIGHTS BEMPEDOIC ACID'S PLACE AMONG NEW THERAPEUTIC OPTIONS. IT COVERS PHARMACOLOGY, SAFETY PROFILES, AND COMPARATIVE EFFECTIVENESS AGAINST TRADITIONAL STATINS AND PCSK9 INHIBITORS. THE TEXT IS AIMED AT CLINICIANS SEEKING TO UPDATE THEIR KNOWLEDGE ON CUTTING-EDGE CARDIOVASCULAR MEDICATIONS.

3. *CLINICAL APPLICATIONS OF BEMPEDOIC ACID IN HYPERCHOLESTEROLEMIA*

FOCUSING ON THE CLINICAL USE OF BEMPEDOIC ACID, THIS VOLUME OFFERS PRACTICAL GUIDANCE FOR HEALTHCARE PROVIDERS MANAGING PATIENTS WITH HIGH CHOLESTEROL. IT DISCUSSES DOSING STRATEGIES, PATIENT SELECTION, AND MONITORING REQUIREMENTS. THE BOOK ALSO ADDRESSES CHALLENGES IN ADHERENCE AND LONG-TERM OUTCOMES.

4. *MECHANISMS OF ACTION AND THERAPEUTIC POTENTIAL OF BEMPEDOIC ACID*

THIS BOOK DIVES INTO THE BIOCHEMICAL PATHWAYS TARGETED BY BEMPEDOIC ACID, EXPLAINING HOW IT INHIBITS ATP CITRATE LYASE TO REDUCE CHOLESTEROL SYNTHESIS. IT EXAMINES PRECLINICAL STUDIES AND THE DRUG'S IMPACT ON LIPID METABOLISM. RESEARCHERS AND PHARMACOLOGISTS WILL FIND THE DETAILED MECHANISTIC INSIGHTS PARTICULARLY VALUABLE.

5. *STATIN INTOLERANCE AND ALTERNATIVE LIPID-LOWERING AGENTS: SPOTLIGHT ON BEMPEDOIC ACID*

ADDRESSING THE CLINICAL PROBLEM OF STATIN INTOLERANCE, THIS TEXT REVIEWS ALTERNATIVE TREATMENTS WITH AN EMPHASIS ON BEMPEDOIC ACID. IT PROVIDES EVIDENCE-BASED RECOMMENDATIONS FOR MANAGING PATIENTS WHO CANNOT TOLERATE STATINS AND DISCUSSES COMBINATION THERAPIES TO OPTIMIZE LIPID CONTROL.

6. *BEMPEDOIC ACID IN CONTEMPORARY CARDIOVASCULAR MEDICINE*

THIS COMPREHENSIVE RESOURCE SITUATES BEMPEDOIC ACID WITHIN THE BROADER CONTEXT OF CARDIOVASCULAR DISEASE PREVENTION AND TREATMENT. IT COVERS EPIDEMIOLOGY, THERAPEUTIC GUIDELINES, AND INTEGRATION OF NEW AGENTS INTO PRACTICE. THE BOOK IS DESIGNED FOR CARDIOLOGISTS, INTERNISTS, AND CLINICAL PHARMACISTS.

7. *PHARMACOECONOMICS OF BEMPEDOIC ACID: COST-EFFECTIVENESS IN LIPID MANAGEMENT*

ANALYZING THE ECONOMIC IMPACT OF INTRODUCING BEMPEDOIC ACID INTO TREATMENT REGIMENS, THIS BOOK EVALUATES COST-EFFECTIVENESS COMPARED TO OTHER LIPID-LOWERING DRUGS. IT INCLUDES HEALTHCARE SYSTEM PERSPECTIVES, BUDGET IMPACT ANALYSES, AND PATIENT ACCESS CONSIDERATIONS. POLICYMAKERS AND HEALTHCARE ADMINISTRATORS WILL BENEFIT FROM ITS INSIGHTS.

8. *INTEGRATIVE APPROACHES TO DYSLIPIDEMIA: COMBINING LIFESTYLE AND BEMPEDOIC ACID THERAPY*

THIS TITLE EMPHASIZES A HOLISTIC APPROACH TO MANAGING DYSLIPIDEMIA, COMBINING PHARMACOLOGICAL TREATMENTS LIKE BEMPEDOIC ACID WITH DIET, EXERCISE, AND BEHAVIORAL CHANGES. IT DISCUSSES PATIENT EDUCATION STRATEGIES AND MULTIDISCIPLINARY CARE MODELS TO ENHANCE TREATMENT OUTCOMES.

9. *FUTURE DIRECTIONS IN LIPID-LOWERING AGENTS: BEYOND STATINS WITH BEMPEDOIC ACID*

LOOKING AHEAD, THIS BOOK SURVEYS EMERGING LIPID-LOWERING THERAPIES WITH A SPECIAL FOCUS ON BEMPEDOIC ACID'S DEVELOPMENT AND POTENTIAL NEW INDICATIONS. IT EXPLORES ONGOING CLINICAL TRIALS AND FUTURE RESEARCH PRIORITIES, PROVIDING A FORWARD-LOOKING PERSPECTIVE FOR CLINICIANS AND SCIENTISTS.

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