

Evaluating patients' individual response regarding anti-inflammatory preparations using the TNF- α inhibitor test

TNF- α as a pro-inflammatory key cytokine

Tumour necrosis factor alpha (short: TNF- α) is a cytokine, which is involved in almost all inflammatory reactions. TNF- α is the first cytokine to be expressed in activated macrophages' signalling cascade. Its most important function is the activation of various immune cells. TNF- α has the ability to promote cell differentiation and the expression of other cytokines. Within the CNS, it causes fever, fatigue, gene-ral feeling of illness and affects the lipid meta-bolism, blood coagulation, insulin resistance, and the convalescence of patients with arteriosclerosis.



Fig.1 local and systemic effects of TNF- α

In rheumatology, TNF- α blockers are administered as longterm anti-rheumatic drugs. Remicade[®], Humira[®] and Enbrel[®] are antibody preparations that block TNF- α 's effect at its receptors on target cells. The preparations above do not inhibit the expression of TNF- α by macrophages. For this therapy approach only the typical anti-inflammatory preparations (e.g. prednisolone) as well as numerous phytopharmaceuticals and some dietary supplements with described anti-inflammatory properties are available so far.

However, regarding their clinical activity, patients show varying responses, which is due to highly differing and individual numbers of toll-like receptors on macrophages.

Therefore, running the TNF- α inhibitor test as a pretest is sensible, if the most effective preparation for an adjuvant anti-inflammatory therapy in an individual case needs to be pre-selected.

Principle of the test

The standardised expression of LPS-induced TNF- α (TNF- α base value) is used as a reference value. In replicate controls, the LPS-induced TNF- α secretion is analysed under the influence of added preparations.

Note: LPS (lipopolysaccharide) is a surface molecule of gram-negative bacteria, which induces a significant inflammatory response (TNF- α secretion) in monocytes/macrophages by binding to their CD14 molecules.

Is the test sensible only in cases with increased $\text{TNF-}\alpha$ levels in blood?

No, TNF- α is used only as a marker cytokine for the activation of macrophages. By detecting the release of TNF- α , the test assesses whether the respective preparation has an impact on the inflammatory cascade. Regarding the detection of LPS-induced inflammatory responses, TNF- α has proven a more sensitive and better test than IL-1 or IL-6. Therefore, the TNF inhibitor test is beneficial in context with any type of myelomonocytic inflammation, even if it has been proven only via increased CRP, IL-1, or IL-6. The TNF- α inhibitor test is a global inflammation inhibitor test.

Material

10 ml heparin blood

Sample receipt within 24 hrs has to be ensured. The sample should be stored and transported at room temperature. Within the Berlin city area, we offer a courier service (+49 (0)30 7701- 250). For collections beyond Berlin, please contact our complimentary courier service (+49 (0)30 77001- 450).

Costs

Costs for the test are a one-time payment of 17.31 \in for the basic TNF- α response and 18.47 \in per each desired preparation.

Labor Berlin	medical report			
Test	Result	Unit		
TNF-alpha inhibitor test The TNF-a base value is the reference value that is used to com- pare corresponding TNF- α levels of each preparation to. Values below the base value indicate an anti-inflammatory effect. Va- lues above the base value indicate a pro-inflammatory in vitro effect of the respective preparation.				
TNF-α base value (LPS-stimulated) TNF-α preparation 1 (1) prednisolone	1760 212	pg/ml pg/ml		
TNF-α preparation 2 (2) Boswellia	354	pg/ml		
TNF-α preparation 3 (3) milk thistle	1782	pg/ml		
TNF-α preparation 4 (4) curcumin	2544	pg/ml		
Next to prednisolone, especially Boswellia has a proven TNF-inhibi- ting and thus anti-inflammatory effect. Milk thistle does not show any effect, while curcumin even has an activating effect.)				

Fig. 2 sample medical report

Do you have questions? Our serviceteam will be happy to support you: +49 (0)30 770 01-220.

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The following preparations are established for the TNF-alpha inhibitor test and are available in the laboratory (alphabetic order):

Preparation (producer)	Medical compound	Preparation (producer)	Medical compound
Alpha Liponsäure Mono (Capsumed)	Alpha lipoic acid	MSM (Essential Food)	Methyl-sulfonyl-methane
Angocin Anti-Infekt N (Repha)	Nasturtium/horseradish	Omega 3 Max (Capsumed)	Omega 3 fish oil
Artemisia Annua pur (Capsumed)	Annual wormwood	Peony Immune (life Extensions)	White peony roots
Arthroflexan plus (Natures Own)	Glucosamine sulphate/Chondroi-	Phyto Curcuma (Capsumed)	Curcuma/turmeric
Banderol (TS Products)	tin sulphate Bark extract	Phyto Isoflavonoid (Capsumed)	Soy isoflavones and red clover
Bicorsan (Jabosan Health Care)		Phyto Weihrauch 400 (Capsumed)	Indian incense
	Boswellia, curcumin, linseed oil	Prednisolon (Mibe)	Glucocorticoid
Boscari (Olibanum)	African incense	Pro Agalostrum (Tisso)	Colostrum, vitamin D3, Agaricus
Boswellion Plus (Natures Own)	Weihrauch, Teufelskralle	Procain (Dr. Loges)	Neuraltherapeuticum
Bromelain (Vita World)	Cysteine protease	Pro Carnitin (Tisso)	L-Carnitine
Catechin-Loges (Dr. Loges)	Green tea extract + vitamin C	Pro Curmin Complete II (Tisso)	Curcuma
CBD – Cannabis Öl (HEALTH+)	Cannabidiol	Pro Dialvit (Tisso)	Combined preparation
Cilantris (Nestmann Pharma)	Coriander	Pro Immupec (Tisso)	Natural fermentation product with vitamin D3
Cistus Incanus (Hannes Pharma)	Cistus		
Coenzym Q 10 (Zein Pharma)	Ubiquinone	Pro Mucosa (Tisso)	L-Glutamine, quercetin, lecithin, broccoli, MSM
Colestyramin (Hexal)	Cholesterol inhibitor	Pro Sirtusan (Tisso)	Bioactive polyphenols
Colostrum (Allcura)	First milk preparation	ProEmsan (Tisso)	Probiotic micro-organism
Cordyceps Extrakt (Capsumed)	Vitality mushroom	Propolis (Langer Vital)	Bee propolis
Cranberry (11 A Nutritheke)	American Cranberry	Quercetin (Bios Medical)	Onion extract
Curcumin-loges® (Dr. Loges)	Curcumin/ vitamin D	reduziertes Glutathion (Apozen)	GSH
Derris Scandens (Siam Heilkräuter)	Flavonoids, isoflavones	Reishi (Capsumed)	Medicinal fungus
Eicosapen (Truw Arzneimittel)	Omega 3 fish oil	Resveratrol (Zein Pharma)	Polyphenol phytoalexin
Eleu Kokk (Boehringer)	Siberian ginseng (Eleutherococ- cus senticosus)	Rizol-Omega Borax-Lösung nach Dr. Steidl (Fürstenplatz Apotheke)	Rizol-Omega, Borax, Chloroquin
Enzym pur (Capsumed)	Bromelain/Papain/Rutin	SAMe 200 pur (Capsumed)	S-Adenosylmethionine
Felis (Hexal)	St. John's Wort	Silymarin (AL)	Milk thistle
Hagebutten Extrakt (Trendbalance)	Rosehip	Super-S-Plus (Dr. Hittich)	MSM-incense-ginger
Hericium-erinaceus	Lion's mane mushroom	Synerga (Laves Arzneimittel)	E.coli
(Mycovital Heilpilze) Hox Alpha (Strathmann)	Stinging nettle	Tigovit (EGCG)	Green tea extract
Inflam effekt (Life Light)	Micronutrients	TNF balance (Trendvital)	Boswellia carterii, curcumin
		TNF direkt (Viathen)	Incense resin
Karazym (Volopharm)	Enzyme preparation	Weihrauch 250 (Natures Own)	Incense
Kelt Vital (Trendvital)	Uridin monophosphat, vitamin B12, 5-Methyltetrahyd- rofol acid	Weihrauch serrata (Trendbalance)	Incense, willow bark, ginger
Koreanischer roter Ginseng (KGV-Korea-Ginseng Vertrieb)	Ginsenosides	Essential oils	Medical compound
Krillöl (Capsumed)	Krill oil	Lavender oil (various pharmacies)	Lavender
L-Carnitin (Diamant Natur)	Amino acid	Lemon oil (various pharmacies)	Lemon
L-Curcumin (Trendbalance)	Curcugreen	Lemongrass oil (various pharmacies)	Lemongrass
L-Curcumin (TrendVital med)	Curcuma longa extract	Thyme oil (various pharmacies)	Thyme
Leinöl (Berco Arzneimittel)	Linseed oil	Rosemary oil (various pharmacies)	Rosemary
Lidocain (Braun)	Neuraltherapeuticum	Manuka oil (various pharmacies)	Manuka
Lassain (Braan)	arattici apeaticulti		

In case you would like to test a preparation that is not listed above, we would kindly ask you to add it to your sample when sending it to us. Upon request, the preparation can be stored up to 3 months in the laboratory to assure substance stability for further tests (please indicate on the transfer form).

hibitor test is not useful, since the effect of these substances is blood or at the target cell.

 $\begin{array}{ll} \mbox{Regarding "TNF-alpha blockers" such as Enbrel^{\tiny (Etanercept), } \\ \mbox{Remicade}^{\tiny (Infliximab) or Humira}^{\tiny (Adalimumab), the TNF incorrect for the substances block TNF-$$$$ are free to the substances block TNF-$$$$$$$$$$$$$$$ are free to the substances block TNF-$$$$$$$$$$ are free to the substances block TNF-$$$$$$$$$$ are free to the substances block TNF-$$$$$$$$$ are free to the substances block the sub$