

STATEMENT ON A NONPROPRIETARY NAME ADOPTED BY THE USAN COUNCIL

USAN (LM-25)

NEMOLIZUMAB

PRONUNCIATION

ne" moe liz' ue mab

THERAPEUTIC CLAIM

Treatment of atopic dermatitis (AD), treatment of prurigo nodularis (PN) and chronic kidney disease associated pruritus (CKD-aP)

CHEMICAL NAMES

Immunoglobulin G2, anti-(interleukin 31 receptor A) (human-Mus musculus monoclonal CIM331 γ -chain), disulfide with human-Mus musculus monoclonal CIM331 κ -chain, dimer

immunoglobulin G2-kappa, anti-[*Homo sapiens* IL31RA (interleukin 31 receptor subunit alpha)], humanized monoclonal antibody gamma2 heavy chain (1-445) [humanized VH (*Homo sapiens* IGHV1-2*02 (83.70%) -(IGHD)-IGHJ5*01) [8.8.14] (1-121) -*Homo sapiens* IGHG2*01 (CH1 C10>S (135), R12>K (137), E16>G (141), S17>G (142) (122-219), hinge C4>S (223) (220-231), CH2 H30>Q (268) (232-340), CH3 R11>Q (355), Q98>E (419) (341-445)) (122-445)], (224-214')-disulfide with kappa light chain (1'-214') [humanized V-KAPPA (*Homo sapiens* IGKV1-39*01 (82.10%) -IGKJ4*01) [6.3.9] (1'-107') -*Homo sapiens* IGKC*01 (108'-214')]; dimer (227-227":230-230")-bisdisulfide

STRUCTURAL FORMULA

Heavy chain (CIM331 (anti-IL-31 RA); γ -chain); X, X")

QVQLVQSGAE	VKKPGASVKV	SC KAS GYTFT	GY IMNWVRQA	PGQGLEWMGL	50
IN PYNGG TD Y	N PQ FQDRVTI	TADKSTSTAY	MELSSLRSED	TAVYY CARDG	100
YDDG PY LET	WGQGLVTVS	SASTKGPSVF	PLAP SK STS	GG TAAL GCLV	150
KDYFPEPVTV	SWNSGALTSG	VHTFPAVLQS	SGLYSLSSVV	TVPSSNFGTQ	200
TYT C NVDHKP	SNTKVDK TVE	RK SC VE CPPC	PAPPVAGPSV	FLFPPKPKDT	250
LMISRTPEVT	C VVV DVS QED	PEVQFNWYVD	GVEVHNAKTK	PREEQF N STF	300
RVVSVLTVVH	QDWLN G KEYK	C KVSNKGLPA	PIEKTISKTK	GQPREPQVYT	350
LPPS Q EEMTK	NQVSLT CL VK	GFYPSDIAVE	WESNGQPENN	YKTT P PMLDS	400
DGSFFLYSKL	TVDKSR WQ EG	NVFS C SVMHE	ALHNHYTQKS	LSLSP	445

Light chains (CIM331 κ -chain); X', X")

DIQMTQSPSS	LSASVGDRVT	IT CQ AS EDIY	SF VAWYQQKP	GKAPKLLI YN	50'
AQ TEA Q GVPS	RFSGSGSGTD	F TLT ISS LQP	EDFATYY CQH	HYDS PL TFGG	100'
G TK VEIKRTV	AAPSVFIFPP	SDEQLKSGTA	SVV CL LNNFY	P RE AKVQWKV	150'
DNALQSGNSQ	ESVTEQDSKD	STYLSSTLT	LSKADYEKHK	VY ACE VTHQG	200'
LSSPVTKSFN	R GE C				214'

Disulfide bridges (denoted in red, bold text, interchain is shaded grey)

22-96	148-204	261-321	367-425
22"-96"	148"-204"	261"-321"	367"-425"
23'-88'	134'-194'	23'''-88'''	134'''-194'''
224-214'	224"-214'"	227-227"	230-230"

Glycosylation sites (most represented N-glycan is G0F)

297 297''

Substitutions into heavy chain variable regions to lower isoelectric point, increase thermal stability, increase neutralizing activity and reduce immunogenicity

S59D Q62P K63Q K65Q
G66D M108L D109E Y110T

Substitutions into light chain variable regions to lower isoelectric point, increase thermal stability, increase neutralizing activity and reduce immunogenicity

R24Q T25A N28D L33V
K52Q L54E K56Q E93D

The CDRs are displayed in bold, underlined (IMGT)

Heavy Chain Modifications (substitutions/deletions) based on human IG2; to improve properties

C131S R133K E137G S138G
C219S H268Q R355Q Q419E
G446[del] K447[del]

MOLECULAR FORMULA	C ₆₃₈₄ H ₉₈₁₄ N ₁₆₇₈ O ₂₀₃₄ S ₄₈ (non-glycosylated)
MOLECULAR WEIGHT	144.15 kDa (non-glycosylated)
TRADEMARK	None as yet
SPONSOR	GALDERMA Research and Development LLC.
CODE DESIGNATIONS	CD14152, CIM331
<u>CAS</u> REGISTRY NUMBER	1476039-58-3
UNII	GN465U8B72
WHO NUMBER	10064

gbk