

July 29, 2015

STATEMENT ON A NONPROPRIETARY NAME ADOPTED BY THE USAN COUNCIL

USAN (cd-14)	DEPATUXIZUMAB MAFODOTIN
PRONUNCIATION	de" pa tux iz' ue mab ma" foe doe' tin
THERAPEUTIC CLAIM	Treatment of cancer

CHEMICAL NAMES

1. Immunoglobulin G1, anti-(human epidermal growth factor receptor) (human-*Mus musculus* monoclonal ABT-806 heavy chain), disulfide with human-*Mus musculus* monoclonal ABT-806 light chain, dimer, tetrakis(thioether) with *N*-[6-(3-mercapto-2,5-dioxo-1-pyrrolidinyl)-1-oxohexyl]-*N*-methyl-L-valyl-L-valyl-(3*R*,4*S*,5*S*)-3-methoxy-5-methyl-4-(methylamino)heptanoyl-(α *R*, β *R*,2*S*)- β -methoxy- α -methyl-2-pyrrolidinepropanoyl-L-phenylalanine
2. Immunoglobulin G1-kappa, anti-(human EGFR protein (Epidermal growth factor receptor)); humanized mouse monoclonal antibody auristatin F conjugate: γ 1 heavy chain (1-446) [humanized VH (human IGHV4-30-4*01 (85%) –(IGHD)-IGHJ4*01 (93%)) [9.7.9] (1-116) –human IGHG1*01 (117-446)] (219-214')-disulfide with kappa light chain (1'-214') [humanized V-KAPPA (mouse IGKV14-100*01 (86%) –IGKJ1*01) [6.3.9] (1'-107') –human IGKC*01 (108'-214')] dimer (225-225":228-228")-bisdisulfide, an average of two disulfide bridges are reduced and the free thiols are S-substituted by (3*S*,6*R*,7*R*,8²*S*,11*R*,12*S*,15*S*,18*S*,26³*RS*)-12-[(2*S*)-butan-2-yl]-3-carboxy-7,11-dimethoxy-6,13,19-trimethyl-5,9,14,17,20,26^{2,5}-hepta-oxo-15,18-bis(propan-2-yl)-4,13,16,19-tetraaza-8(2,1),26(1)-dipyrrolidina-1(1)-benzenahexacosaphan-26³-yl radicals (mafodotin)

STRUCTURAL FORMULA

Heavy chain

QVQLQESGPG	LVKPSQTL ¹ SL	TCTVSGYSIS	SDFAWN ² WIRQ	PPGK ³ GLEW ⁴ MG	50
YISYSGNTRY	QPSL ⁵ KSRITI	SRDTSKNQFF	LKLNSV ⁶ TAAD	TATYYC ⁷ V ⁸ TAG	100
RGFPYWGQGT	LVTVSSASTK	GPSVFPLAPS	SKSTSGGTAA	LGCLV ⁹ KDYFP	150
EPVTVSWNSG	ALTS ¹⁰ GVHTFP	AVLQSSGLYS	LSSV ¹¹ TV ¹² PSS	SLGTQTYICN	200
VNHKPSNTKV	DKKVEPKSCD	KTHTCP ¹³ PCPA	PELLGGPSVF	LFPPK ¹⁴ PKDTL	250
MISRTPEVTC	VVVDVSHEDP	EVKFNWYVDG	VEVHNAKTKP	REEQY ¹⁵ NSTYR	300
VVSVLTVLHQ	DWLNGKEYKC	KVSNKALPAP	IEKTI ¹⁶ SKAG	QPREPQVYTL	350
PPSRDELTKN	QVSLTCLVKG	FYPSDIAVEW	ESNGQPENNY	K ¹⁷ TPPVLDSD	400
GSFFLYSKLT	VDKSRWQQGN	VFSCSV ¹⁸ MHEA	LHNHYTQKSL	SLSPGK	446

Light chain

DIQMTQSPSS	MSVSVGDRVT	ITCHSSQDIN	SNIGWLQQKP	GKSFKGLIYH	50'
GTNLDDGVPS	RFSGSGSGTD	YTLTIS ¹ SLQP	EDFATYYCVQ	YAQFPWTFGG	100'
GTKLEIKRTV	AAPSVFIFPP	SDEQLKSGTA	SVVCLLN ² FY	PREAKVQWKV	150'
DNALQSGNSQ	ESVTEQDSKD	STYLSSTLT	LSKADY ³ EKHK	VYACEVTHQG	200'
LSSPVTKSFN	RGE ⁴ C				214'

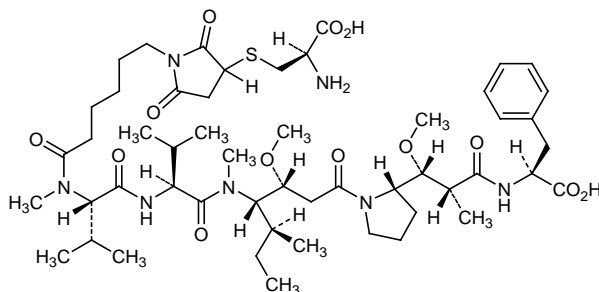
Disulfide bridges location

22-96	22"-96"	23'-88'	23'''-88'''	134'-194'	134'''-194'''	143-199	143"-199"
214'-219*	214'''-219'''*	225-225**	228-228**	260-320	260'''-320'''	366-424	366"-424"

* Partly reduced

Modified residues

C
An average of 2 disulfide bridges are reduced and the free cysteines are S-substituted as shown.

Glycosylation sites (N)

Asn-296 Asn-296''

MOLECULAR FORMULA

$$\text{C}_{6624}\text{H}_{10228}\text{N}_{1728}\text{O}_{2052}\text{S}_{42} \cdot (\text{C}_{49}\text{H}_{76}\text{N}_6\text{O}_{11})_n$$

(unglycosylated)

MOLECULAR WEIGHT

148.3 kDa (unglycosylated, $n = 4$)

TRADEMARK

None as yet

SPONSOR

AbbVie

CODE DESIGNATIONS

ABT-414

CAS REGISTRY NUMBER

1585973-65-4

UNII

F3R7A4P04N

WHO NUMBER

10263

gbk