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2 STUDY SYNOPSIS

Names of Company: GluckTpe"cpf "GluckNvf 0	RF&F WCN"UWVF ["VCDNG"	*Hqt" P c&qpcri' Cwj qtkv{ " Wug" Qpn{ "+"
Name of Finished Product: Ngpxcvpkl"G92: 2+"qten'cdngv	Tghgttkpi "q"Rctv"K" qh'yj g'F quukgt"	
Name of Active Ingredient: Ngpxcvpkl"G92: 2+	Volume:	Page:

Study Title:	Rj cug"4.'O wnegpvgt.'Qr gp/rdgn'Ulpi rg'Cto 'Vtkri'q'Gxcnvcg'yj g'Uchgv{ 'cpf "Ghlece{ "qh' Qten'G92: 2'lp'O gf wmt { 'cpf "Kf lpg/353'Tghtcevt { .'Wptgugevcdng'F khtgpkvcgf "Vj { tqkf " Ecpegtu.'Utcvklgf "d{ 'J kurnqi {
Investigators/ Sites:	O wnegpvgt<"52'ukgu'lp'yj g'WU.'Kcn{ .'Wpksf 'Mpi f qo .'Cwutcrk.'Rqmpf.'cpf "Hcpeg0"C" rku'qh'lxguki cvqtu'cpf 'ukgu'ku'r tqxf gf 'lp"Cr r gpf k'38060
Publications based on study:	<p>3+ Uj gto cp'U.'Lct cd'D.'Ecdeplmru'O . 'Nlektc'N.'RcepkH'O ctvpu'T. 'gv'crf0C'r j cug'K' vlcrl'qh'yj g'o wnkcti gvgf 'nlpug'lpj kdkqt'G92: 2'lp'cf xcpegf 'tcf kqkf lpg'*TCK/ tghtcevt { 'f khtgpkvcgf 'y { tqkf 'ecpegt '*F VE+'Jcdutcev_0Rtgugpvcvqp'cv'yj g'69y " Co gtkecp'Uqelgv 'qh'ErklecniQpeqmi { 'CpwwriO ggvpki =4233' Lxp'6/: =Ej leci q'*KN+0 L'Erkp'Qpeqr04233-4; *Uwr r n=77250'</p> <p>4+ Uj gto cp'U.'Lct cd'D.'Ecdeplmru'O . 'Nlektc'N.'RcepkH'O ctvpu'T. 'gv'crf0C'r j cug'K' vlcrl'qh'yj g'o wnkcti gvgf 'nlpug'lpj kdkqt'G92: 2'lp'cf xcpegf 'tcf kqkf lpg'*TCK/ tghtcevt { 'f khtgpkvcgf 'y { tqkf 'ecpegt '*F VE+'Jcdutcev_0Rtgugpvcvqp'cv'yj g' y " CppwriO ggvpki 'qh'Lcr cpug'Uqelgv 'qh'O gf leci'Qpeqmi { =4233' Lwn43/45= " [qnqj co c' "Lcr cp=0'</p> <p>5+ Uj gto cp'U.'Lct cd'D.'Ecdeplmru'O . 'Nlektc'N.'RcepkH'O ctvpu'T. 'gv'crf0G92: 2" *Ngpxcvpkl+lp'Cf xcpegf 'Tcf kqkf lpg'*TCK/Tghtcevt { 'f khtgpkvcgf 'y { tqkf 'ecpegt '*F VE+=Tguwru'qh'c'O wnkEgpvgt'Rj cug'K'vlcrl'qtcrn'rgugpvcvqp_0Rtgugpvcvqp'cv' yj g'57y 'CpwwriO ggvpki 'qh'yj g'Gwtqr gcp'Vj { tqkf 'Cuuqekvcqp=4233' Ugr '32/36= 'Mtcnqy "Rqmpf +04233-QR230' j wr <leqpvgtv'cti gt(qo Rtgf wmgF Dlr tqf wmgBur Av{ r ? r f h f qk?552787"</p> <p>6+ DcmfF . 'Uj gto cp'U.'Lct cd'D.'Ecdeplmru'O . 'Nlektc'N.'RcepkH'gv'crf0C'r j cug'K'vlcrl' qh'yj g'o wnkcti gvgf 'nlpug'lpj kdkqt'ngpxcvpkl"G92: 2+lp'cf xcpegf 'tcf kqkf lpg" *TCK/tghtcevt { 'f khtgpkvcgf 'y { tqkf 'ecpegt '*F VE+<eqttgrvcqp'qh'v'gco gpv' qweqo gu'y kj 'wo qt' i gpgvle'cpcn'uku.'ugtwo 'dlqo ctngtu'cpf 'r j cto ceqnlpgvleu" Jcdutcev_0Rtgugpvcvqp'cv'yj g': 3uv'CpwwriO ggvpki 'qh'yj g'Co gtkecp'Vj { tqkf " Cuuqekvcqp=4233'Qev'48/52=lpf lcp'Y gmu'EC-0Vj { tqkf 0'4233'Qev'43*3" Uwr r n=C8/90'</p> <p>7+ DcmfY . 'Uj gto cp'UK'Lct cd'D.'Ecdeplmru'O G.'O ctvpu'T. 'Uj cj 'O J . 'gv'crf0 Ngpxcvpkl'tgco gpv'qh'cf xcpegf 'TCKtghtcevt { 'f khtgpkvcgf 'y { tqkf 'ecpegt '*F VE+<e{ qmkg'cpf 'cpi kji gple'fcevq '*ECH+r tqhki 'lp'eqo dlpvcqp'y kj 'wo qt' i gpgvle'cpcn'uku'v'q'f gpvhl' 'o ctngtu'cuuqekvcgf 'y kj 'tgr qpug'Jcdutcev_0Rtgugpvcvqp' cv'yj g'4234' CUEQ' CppwriO ggvpki =4234' Lxp'3/7=Ej leci q'*KN+0L'Erkp'Qpeqr0' 4234-52*Uwr r n=773: 0"</p> <p>8+ Uej nwo dgti gt'O . 'Lct cd'D.'Ecdeplmru'O G.'Tqdlpuqp'D.'RcepkH'DcmfY . 'gv'crf0C" r j cug'K'vlcrl'qh'yj g'o wnkcti gvgf 'nlpug'lpj kdkqt'ngpxcvpkl"G92: 2+lp'cf xcpegf " o gf wmt { 'y { tqkf 'ecpegt '*O VE+'Jcdutcev_0Rtgugpvcvqp'cv'yj g'6: y 'Co gtkecp' Uqelgv 'qh'ErklecniQpeqmi { 'CpwwriO ggvpki =4234' Lxp'3/7=Ej leci q'*KN+0L'Erkp' Qpeqr04234-52*Uwr r n=77; 30'</p>

	<p>9+ Hwpcj cuj k[. 'Mcf qy cnk'V. 'Vqj { co c'Q. 'O wq'J . 'O cvl gxl e'O. 'Ucej f gx'R'gv'cnf' Ngpxcvlkd'tgcvo gpv'qh'f khtgtpvkn'yj { tqkf "ecpegt" *F VE+<Cpcn' uku'q'f gpv'kh' " dkqo ctngtu'cuuqekcvf'y kj 'tgr qpug'Jcdutcev_0'Rtgugpvcvqp'cv'yj g'93^u'Lcr cpug' Ecpegt'Cuqekcvqp'CpwcniO ggkpi . '4234'Ugr "3; /43=Ucr r qtq. 'Lcr cp0'</p> <p>Ugg"Cr r gpf lz"380830'</p>
Study Period:	28'P qx"422: "q"33'Cr t"4233"
Phase of Development:	Rj cug"4"
Objectives:	<p>Primary Objective</p> <p>Kp"uwlgeu'y kj "o gf wmt { "j { tqkf "ecpegt" *O VE+<qt'tcf kqkf kpg" *⁵³K/tghcevt { ltgukcpv f khtgtpvkvf'yj { tqkf "ecpegt" *F VE+< 'y g'r tlo ct { "qdlgev'xgu'y gtg'q<"</p> <ul style="list-style-type: none"> F gvgto kpg"y g"ghgev'qh'ngpxcvlkd"qp'yj g'qdlgev'xg'tgur qpug'tcvg" *QTT+<dcugf "qp" yj g'o qf khtg' "Tgur qpug" Gxcnvcvqp" Etkgtk'kp "Uqkf" Vwo qtu" *TGEKV+<d { "cp kpf gr gpf gpv'lo ci kpi 'tgxky " *KT+<" F gvgto kpg" yj g" r j cto ceqnlpgve" *RM+< r tqhkg" cpf" yj g" r j cto ceqnlpgve" r j cto ceqf { pco le" *RMIRF +<tgrvkvpuj kr u'qh'ngpxcvlkd0' <p>Secondary Objectives</p> <p>Kp" uwlgeu' y kj " O VE" qt'tcf kqkf kpg" *⁵³K/tghcevt { ltgukcpv' F VE." yj g'ugeqpf c t { " qdlgev'xgu'y gtg'q<"</p> <ul style="list-style-type: none"> F gvgto kpg"y g"ghgev'qh'ngpxcvlkd"qp'f wcvkp'qh'tgur qpug'd { "yj g'KT" O gcuwtg"y g"ghgev'qh'ngpxcvlkd"qp'yj g'f kugcug"eqpvtqitcvg" *F ET+<eqo r ngvg' tgur qpug"JET_ "r ctvkn'tgur qpug"JRT_ "qt'uvcdrg'f kugcug"JUF_+<cpf "enplecn'dgpgk" tcvg" *EDT+<ET. 'RT. 'cpf "f wcdrg'UF +<d { "KT" F gvgto kpg"y g"vko g'vq'tgur qpug'd { "KT" Gxcnvcv"y g"ghgev'qh'ngpxcvlkd"qp'r tqi tguukp/hgg'wtxkcn' *RHU+<d { "KT" cpf " qxgtcn'wtxkcn' *QU+<" Gxcnvcv"y g'uchgv' "cpf "vqrgcdkkr { "qh'ngpxcvlkd" Cuuguu"y g" kphwpeg" qh' F P C'ug s wpeg" xctkcpw" qp" o gxcdqrle" gpl { o gu" cpf " vcpur qtvgtu" r quukdn { " kpxqkrf" kp" xctkcdkkr { " qh' ngpxcvlkd" RM' r ctco gvgtu" d { " i gpqv' r kpi "uwlgeu'i gpqo le' F P C'wukpi "y g'Chh { o gvtlz' F O GVl " cttc { " F gvgto kpg"y g"dkqej go lecn'tgur qpug" wukpi "wo qt" o ctngtu" *gkj gt" yj { tqi rwdwkp" hqt" F VE" uwlgeu' qt'ecrekvlp" cpf "ect elpqgo dt { qple" cpki gp" JEGC_ "hqt" O VE" uwlgeu+<" Cuuguu"y g"ghgev'qh'u qo cve" F P C'ug s wpeg" xctkcpw" kp" DTCH"J / . "M" cpf "P / Tcu" cpf "TGVRVE3. "4. "cpf "5. "cpf "i gto rlp" F P C'ug s wpeg" xctkcpw" kp" TGV" cpf " pgct" HQZG3 " *tu; 87735+<cpf "P MZ4/3" *tu; 664: ; +<qp" uwlgev' tgur qpug" q" uwf { " vtgcvo gpv" Kpxgunki cvg"y g'r qvkvcn'eqtgrvkv"qh'yj g'hqmy kpi "dkqo ctngtu'y kj "ghlece { <" É Ugtwo "r tqvqo g"gzr tguukp" *e { vqnlpg" cpf "cpi kqi gple' hcevt { "JECH+<" É Ugtwo "dkqo ctngtu" qh' cr qr vuku" *ecur cug" 5 9" JECur " 5 9_ " e { vqej tqo g" e JE { vqE_ " cpf "O 52" pgqcpki gp" JO 52_+<"
Methodology:	<p>Vj ku"y cu'c'Rj cug"4. "o wnekgpvt. "qr gp/rcdgn" ukpi rg/cto "uwf { . "utcvkhgf" d { "j kvqni { " *F VE" qt' O VE+0"</p> <p>Vj ku"uwf { "eqpvkpgf "yj tgg'Rj cugu<" yj g'Rtgtgcvo gpv'Rj cug. "y g"Vtgcvo gpv'Rj cug. "cpf "y g" Gzvpukp'Rj cug0"</p>

	<p>Vj g"Pretreatment Phase"ruxgf "pq"npi gt "y cp"4: "f c { u0"Kphqto gf "eqpugpv'y cu'qdvckpgf" cpf "r tqvqeqn'gri kdkk" { "cpf "f kugcug"ej ctcevgtkueu'y gt g'gucdkuj gf "r tkt "vq"tgcvo gpv0"</p> <p>Vj g"Treatment Phase"eqpukugf "qh" c "Vtgcvo gpv"Rgtkf "cpf "c" Hq nny /wr "Rgtkf 0" Vj g' Vtgcvo gpv"Rgtkf "qh" yj g' Vtgcvo gpv"Rj cug" dgi cp" cv'yj g' vko g' yj cv'yj g' hktu' uwdlgev' dgi cp' uwwf { "f twi "cf o kpkutcvkqp" cpf "gp gf "cv'yj g' vko g' yj gp" cm' uwdlgeu' gptqmgf "eqo r ngvgf " gki j v'e { engu' qh' tgcvo gpv' qt "f kueqpvpwgf "uwwf { "tgcvo gpv' r tkt "vq" yj g' gki j yj "e { eng0" Cm' uwdlgeu' yj gp' gpvgf "y j g' Gz vgpukp" Rj cug0"</p> <p>Vj g"Extension Phase"eqpukugf "qh" c "Vtgcvo gpv"Rgtkf "cpf "c" Hq nny /wr "Rgtkf 0" Vj g' Gz vgpukp" Rj cug" dgi cp" cv'ko o gf kvgn' "chgt "y j g' Vtgcvo gpv"Rj cug" gp gf "cpf "kpenw gf "cm' uwdlgeu' yj cv'y gt g' gki gt "uknitgegkpi "tgcvo gpv' qt "kp' hq nny /wr 0"</p> <p>Kp' gki gt "y j g' Vtgcvo gpv"Rj cug" qt "Gz vgpukp" Rj cug. "y j g' uwdlgev' f kueqpvpwgf "uwwf { "f twi "cf o kpkutcvkqp" y j gp' qpg' qh' yj g' hq nny kpi "qewttgf <" f kugcug' r tqi tguukp. "y j g' f gxgr o gpv' qh' wpccegr vdrng' vqzlek' . "y j g' uwdlgeu' yj k f tcy cniq' h' eqpugpv' hqt' r ctvek' cvkqp' kp' yj g' uwwf { . "qt "y j g' uwdlgeu' ej qleg' vq' uqr "uwwf { "tgcvo gpv0" Vj g' f kueqpvpwgf "uwdlgev' cf "c" Hkpcn' Xkukv qt "Vgto kpcvqp" Xkukv' 52' f c { u' hq nny kpi "hpcn' uwwf { "f twi "cf o kpkutcvkqp. "cpf "y cu' hq nny gf "kp' yj g' Hq nny /wr "Rgtkf "qh" yj g' Vtgcvo gpv"Rj cug" qt "Gz vgpukp" Rj cug" f gr gp f kpi "qp" y j cv' r j cug' yj g' uwdlgev' y cu' kp' y j gp' f kueqpvpwgf "qewttgf 0" Vj g' uwdlgev' y j q' f kueqpvpwgf "uwwf { "tgcvo gpv' dghqt g' f kugcug' r tqi tguukp" cpf "f k' "pqv' y k f tcy "eqpugpv' eqpvpwgf "vq' wpf gti q" wo qt "cuuguo gpv' gxgt { "5" o qpj u' uctvki "5" o qpj u' htqo "y j g' rux' wo qt "cuuguo gpv' cpf "eqpvpwgf "wpki' f qewo gpv' cvkqp' qh' f kugcug' r tqi tguukp0" Hqt' uwdlgeu' y j q' f kueqpvpwgf "y k f r tqi tguukp" f kugcug' uwtxk' cni' hq nny /wr "qewttgf "gxgt { "5" o qpj u' hqt' yj g' hktu' 4" { gctu' qh' uwwf { "tgcvo gpv' yj gp' gxgt { "8" o qpj u' f wtkpi "I gctu' 5" cpf "6. c pf " { gctn' " yj gt gchgt "wpki' uwdlgev' gcj "qewttgf 0"</p> <p>Vj g' vko g' qh' f cv' ewqhi' hqt' yj g' r tko ct { "uwwf { "cpcn' uku' qewttgf "y j gp' cm' uwdlgeu' gptqmgf "kp' yj g' uwwf { "eqo r ngvgf "gki j v'e { engu' qh' tgcvo gpv' qt "f kueqpvpwgf "uwwf { "tgcvo gpv' r tkt "vq" yj g' gki j yj "e { eng0"</p>
Number of Subjects (Planned and Enrolled):	<p>Rrppgf <" r r tqzko cvgn' "326" uwdlgeu"</p> <p>Gptqmgf "Uetggpgf <" 384" uwdlgeu"</p> <p>Vtgcvgf <" 339" uwdlgeu" 7: "F VE" uwdlgeu" cpf "7; "O VE" uwdlgeu" +</p>
Diagnosis and Main Criteria for Inclusion and Exclusion:	<p>Uwdlgeu" ≥ "3: " { gctu' qh' ci g' y k f "j kuxqni kecm' / "qt' e { vqni kecm' / eqphkto gf "f kci pquku' qh' gki gt "F VE" qt "O VE" wptgugevdrng' f kugcug' yj cv'y cu' pqv' co gpcedng' vq' uwti gt { "y k f "gxkf gpeg' qh' f kugcug' r tqi tguukp" dcugf "qp" o qf kkgf "TGEKUV" y k f kp" 34" o qpj u" 3" o qpj "vq' cmjy " hqt' xctkpegu" kp' uwdlgev' uecpplpi "kpvgtxcni" r tkt "vq" uwwf { " gpv' { "y gt g' gki kdrng' vq" dg' gptqmgf "kp' yj ku' uwwf { 0"</p> <p>Inclusion criteria:</p> <p>30 Y k f "j kuxqni kecm' / "qt' e { vqni kecm' / eqphkto gf "f kci pquku' qh' qpg' qh' yj g' hq nny kpi <" c0 F VE "kpenw kpi "cp { "qh' yj g' hq nny kpi "uwdv { r gu<"</p> <ul style="list-style-type: none"> Rcr knct { "y { tqkf "ecpegt "RVE" + <ul style="list-style-type: none"> Hqnlewrct "xctkcpv" Xctkcpw" "kpenw kpi " dw' "pqv' rko ksf " vq' cm' egm' eq nwo pct" egm' etkdt kqto / o qt wrct. "uqrk' . "qz { r j kn" Y ctvj kpa' r kng. " vtdgedwrct. " wo qt " y k f "pqf wrct" hucelku' r kng" utqo c. " J A' vj ng' egm' x ctkcpv' qh' r cr knct { " ectekp qo c. " r qqt n' "f kkgtpv' cvgf +" Hqnlewrct "y { tqkf "ecpegt "HVE" + <ul style="list-style-type: none"> J A' vj ng' egm' Engct "egm' Kpuwrct " <p>d0 O VE"</p> <p>40 Y k f "o gcuwtdrng' f kugcug' o ggkpi "y j g' hq nny kpi "etkgtkqp <" c0 Cv' ngcu' qp g' ngukp " ≥ 30" eo "kp' npi guv' f kco gygt "JNF " hqt' pqpni o r j "pqf gu" cpf " ≥ 40" eo "kp' NF " hqt' h' o r j "pqf gu" y j kej "y cu' ugtkcm' " cpf "cee wtcvgn' " o gcuwtdrng' ceeqtf kpi " vq' o qf k kkgf " TGEKUV" wukpi "gki gt" e qo r wgtk gf "</p>

Exclusion criteria:

	<p>o gvcucuku+ "cpf "y j g' uwdlgev'y cu'npqy p "vq" j cxg' dpgp' o gvcucuku. 'y j g' rlxgt/ ur gekhe' cmerkp' r j qur j cvug' o wuv' j cxg' dggp' ugr ctevgf "htqo "y j g' vqwn' cpf " y j g' rlxgt' ur gekhe' cmerkp' r j qur j cvug' cnpgp' uij qwrf " j cxg' dggp' wugf "vq' cuuguu' rlxgt' hmpvklp0'</p> <p>f0 etgcwklpg' ergetcpeg' > "82" o Nlo kp' r gt' y j g' Egenetqhn' cpf "I cwn' hqto wrt" "</p> <p>50' Uki phtecpv' ectf kqxcuewrt "lo r ckt o gpv' j kxqt { "qh' eqpi gukxg' j gctv' hckwtg' @P gy " [qtni' J gctv' Cuqekvklp"] P [J C_ " Ercuu' KK" wpuvcdng' cpi kpc" qt" o { qctf' kcn' kphctevklp' y kj kp' 8" o qpj u' qh' uwwf { "uvtv' qt' ugt' kqwu' ectf kce' cttj { y j o kc+ "</p> <p>60' Cevxg' j go qr v' uku' *dtki j v' tgf "dmqf' qh' cv' hgcuv' " vgcuv' qqp+ "lp' y j g' 4: "f c{ u' r tkqt " vq' uwwf { "gpv' { "</p> <p>70' Dnggf kpi "qt' y j tqo dqvle' f kuqt' ftu' qt' wug' qh' cp' v' eqci wcpvu. "uwej "cu' y cthctkp. "y kj " c' y j gter gwle' kp' vgt' pcvklp' cn' pqt o cki gf "tevlq' *KP T+ "</p> <p>80' Rqukkxg' j kxqt { " qh' J KK. "cev' x g' j gr' cvklku' D' q t" cev' xg' j gr' cvklku' E' q t' ugxgt' ghpeqptqngf' "lp' vgt' ewt' gpv' kmpguu' qt' "lphgevlq' "</p> <p>90' Qti cp' cmqi tchm' tgs vkt kpi "lo o wpuwr r tguukxg' t' gcv o gpv' "</p> <p>: 0' Rtkqt' o cki pcpe { " qj gt' y j cp' pqp o gncp qo c' "unkp' ecepgt' qt' egt' xlecn' ectekp qo c' "lp' ukw' wprguu' y j g' r tkqt' o cki pcpe { "y cu' f kci pqugf' cpf "f gh' pklxgn' "tgcvgf' > "7" { gctu' r t' gxlqwu' y kj "pq' uwdugs wgpv' gxf' gpeg' qh' t' g' ewt' gpeg' "</p> <p>; 0' Dtlcp' qt' rgr' vqo gplpi gcn' *egpvt' cn' pgt' xqwu' u' { ugo "]EP U_+ " o gvcucugu0' " Uwdlgevu' y kj " ucdng' qt' r t' gxlqwu' " ktcf' k' cvgf' " dtclp' o gvcucugu' y j g' g' cnq' gzenmf gf 0' "</p> <p>320' O ctngf' " dcugr' kpg' r t' q' npi cvklp' qh' S V S Ve" "lp' vgt' xcn' " S Ve" "lp' vgt' xcn' > "722" o uge+ " wukpi "y j g' Hkf' g' t' k' c' o g' j qf " S Ve" ? " S V I T ²⁰⁵ + " hqt' S Ve' cpcn' uku' "</p> <p>330' I tgcvgt' y j cp' 3- " r t' q' v' k' p' w' t' c' " qp' w' k' p' g' f' k' r' u' k' m' v' g' u' k' p' i " qt' @52" o i f N0' " Uwdlgevu' y kj " r t' q' v' k' p' w' t' c' @3- " qp' w' k' p' g' f' k' r' u' k' m' v' g' u' k' p' i " y qwrf " j cxg' w' p' f' g' t' i p' p' g' 46/ j qwt' w' t' k' p' g' e' q' n' g' e' v' k' l' p' h' q' t' s' w' c' p' v' k' c' v' x' g' c' u' g' u' o' g' p' v' q' h' r' t' q' v' k' p' w' t' c' 0' " Uwdlgevu' y kj " 46/ j qwt' r t' q' v' k' p' > "3" i 46' j qwtu' y j g' g' t' g' l' p' g' r' i k' d' ng' 0' "</p> <p>340' J kxqt { "qh' i' c' u' t' q' l' p' v' g' u' k' p' c' n' o' c' m' d' u' q' t' r' v' k' l' p' " qt' j' c' x' k' p' i " w' p' f' g' t' i p' p' g' " u' w' t' i g' t' { " t' g' s' v' k' t' k' p' i " i' c' u' t' q' l' p' v' g' u' k' p' c' n' o' c' p' c' u' q' o' q' u' g' u' y j k' p' 6" y j g' g' m' i " qh' u' v' t' v' k' p' i " v' j g' t' c' r { " qt' y j q' u' g' " y j q' j cxg' p' q' v' t' g' e' q' x' g' t' g' f' " h' t' q' o " o' c' l' q' t' u' w' t' i g' t' { " y j k' p' 6" y j g' g' m' i " qh' u' v' t' v' k' p' i " y j g' t' c' r { " Y qo gp' y j q' y j g' t' g' r' t' g' i p' c' p' v' q' t' " dtgcuv' h' g' f' k' p' i = y qo gp' qh' e' j k' f' d' g' c' t' k' p' i " r' q' v' g' p' v' k' c' n' y kj " c' r' q' u' k' x' g' r' t' g' i p' c' p' e { " v' g' u' v' c' v' r' t' g' t' g' c' v' o' g' p' v' q' t' " p' q' r' t' g' i p' c' p' e { " v' g' u' 0' " Y qo gp' qh' e' j k' f' d' g' c' t' k' p' i " r' q' v' g' p' v' k' c' n' w' p' r' g' u' i " 3+ " u' w' t' i k' e' c' m' i " u' n' g' t' k' g' " qt' " 4+ " w' u' k' p' i " c' f' g' s' w' c' v' g' o' g' c' u' w' t' g' u' qh' e' q' p' t' c' e' g' r' v' k' l' p' " * k' p' e' n' f' k' p' i " v' y q' h' q' t' o' u' qh' e' q' p' t' c' e' g' r' v' k' l' p' . " q' p' g' qh' y j k' e' j " o' w' u' v' d' g' c' " d' c' t' t' l' g' t' o' g' v' j qf + " lp' y j g' q' r' k' p' k' p' qh' y j g' l' p' x' g' u' k' i c' v' q' t' 0' " R' g' t' k' o' g' p' q' r' c' w' u' c' n' y qo gp' o' w' u' v' j cxg' dggp' co gp' q' t' t' j g' l' e' " h' q' t' " c' v' r' g' c' u' v' 34" o' q' p' j u' " v' q' d' g' e' q' p' u' k' f' g' t' g' f' " qh' p' q' p' e' j k' f' d' g' c' t' k' p' i " r' q' v' g' p' v' k' c' n' 0' " H' g' t' v' k' g' " o' c' r' g' u' y kj " h' g' o' c' r' g' r' c' t' v' p' g' t' u' y j q' y j g' t' g' p' q' v' y k' r' k' p' i " v' q' w' u' g' e' q' p' t' c' e' g' r' v' k' l' p' " qt' y j q' u' g' h' g' o' c' r' g' r' c' t' v' p' g' t' u' y j g' t' g' p' q' v' w' u' k' p' i " c' f' g' s' w' c' v' g' " e' q' p' t' c' e' g' r' v' k' x' g' r' t' q' v' g' e' v' k' l' p' y j g' t' g' z' e' n' f' g' f' 0' "</p> <p>360' Qj gt' uki phtecpv' f kugcug' qt' f kuqt' ftg' y j cv' "lp' y j g' l' p' x' g' u' k' i c' v' q' t' a' u' q' r' k' p' k' p' . " y qwrf " gzenmf g' y j g' uwdlgev' h' t' qo " y j g' uwwf { " "</p> <p>370' Rtgxlqwu' h' g' p' x' c' v' k' l' d' y j g' t' c' r { " "</p> <p>380' Rtgxlqwu' t' g' c' v' o' g' p' v' y kj " cp' " l' p' x' g' u' k' i c' v' k' p' c' n' i' f' t' w' i . " y kj " y j g' g' z' e' g' r' v' k' l' p' qh' y j q' u' g' k' f' g' p' v' h' g' f' " lp' " k' p' e' n' u' k' l' p' " E' t' k' g' t' k' l' p' " %0. " y kj kp' y j g' 52" f c { u' r' t' k' q' t' " v' q' y j g' u' v' t' v' q' h' r' g' p' x' c' v' k' l' d' c' f' o' k' p' k' u' t' c' v' k' l' p' "</p> <p>390' J kxqt { "qh' c' n' e' q' i q' r' k' u' o . " f' t' w' i " c' f' f' k' e' v' k' l' p' . " r' u' { e' j k' c' v' t' k' e' " qt' r' u' { e' j q' r' i q' i k' e' c' n' e' q' p' f' k' k' l' p' . " qt' u' q' e' k' c' n' u' k' w' c' v' k' l' p' " y j k' e' j . " lp' y j g' q' r' k' p' k' p' qh' y j g' l' p' x' g' u' k' i c' v' q' t' . " y qwrf " lo r ckt' uwwf { " eqo r r' k' c' p' e' g' "</p> <p>3: 0' Ngi cnlpecr cekf 0</p>
Test Treatment, Dose, Mode of Administration, and Batch Numbers:	<p>Ngpxcvklp' y cu' cf o l' p' k' u' n' g' t' g' f' "lp' y q' f' q' u' c' i g' u' f' w' t' k' p' i " y j k' u' u' w' f { 0 " C' e' e' q' t' f' k' p' i " v' q' y j g' q' t' k' i k' p' c' n' r' t' q' v' e' q' n' " y j g' f' q' u' c' i g' y cu' q' p' g' 32" o i " v' c' d' n' g' y j k' e' g' f' c' k' n' " * D K F + 0 " V y q' u' w' d' l' g' e' v' u' y j g' t' g' v' t' g' c' v' g' f' " y kj " y j k' u' f' q' u' c' i g' 0 " V j g' f' q' u' c' i g' y cu' e' j c' p' i g' f' " v' q' 46" o i " q' p' e' g' f' c' k' n' " * S F + y j k' e' j " e' q' p' u' k' u' g' f' " qh' y j q' 32" o i " v' c' d' n' g' u' c' p' f' " q' p' g' 6" o i " v' c' d' n' g' u' * C o g' p' f' o' g' p' v' 23+ 0 " C " v' q' w' c' n' qh' 37" u' w' d' l' g' e' v' u' y j g' t' g' v' t' g' c' v' g' f' " y kj " y j k' u' f' q' u' c' i g' 0 " Ngpxcvklp' y cu' u' g' r' h' c' f' o' l' p' k' u' n' g' t' g' f' " q' t' c' m' i " d { " u' w' d' l' g' e' v' u' 0 " U' g' g' " V' c' d' n' g' 4 " h' q' t' d' c' v' e' j " p' w' o d' g' t' u' 0</p>

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Reference Therapy, Dose, Mode of Administration, and Batch Numbers:"	P qv'cr r decdng
Duration of Treatment:	Uwdlgeu'y gtg"tgcvgf"lp"v j g"Vtgcwo gpv'Rgtkqf"qh"vj g"Vtgcwo gpv'Rj cug"cpf" Gzvgpukqp Rj cug0"Vj g"uwdlgev'eqpvpwgf"uwwf { "tgcwo gpv'wpvki'f kugcug"r tqi tguukqp."f gxgtr o gpv'qh' wpceegr vcdng"vqzlek{ .f gcvj . "uwdlgeu"y kj f tcy ch'qh'e qpugpv"ltqo "r ctvlekr cvkqp"kp"vj g" uwwf { . "qt"uwdlgeu'ej qleg"q"uqr"uwwf { "tgcwo gpv0
Criteria for Evaluation:	<p>Efficacy:</p> <p>Tumor Assessments for Response Assessment:</p> <p>Tumor Assessments for Subjects with DTC:</p> <p>Rtgtgcwo gpv'wo qt"cuuguuo gpw"vulpi "EV"qh"vj g"pgemlej guvcdqo gp'lr grklu"cpf" qvj gt" ctgcu"qh'npqy p" f kugcug"qt"pgy n' "uwwr gev'f" f kugcug"y gtg"r gthqto gf"y kj kp"6"y ggmu'r tkqt" vq"vj g"htuv'f qug"qh'ngp xcvpkd"cf o kpkntcvkqp0"Uecpu"qh"vj g"pgem"cdf qo gp."r grklu"cpf" qvj gt"ctgcu"qh"vj g"dqf { "o c { "j cxg"dggp" f qpg"y kj "O TKlpugcf"qh"EV."dw"gxcmcvkqp"qh" vj g"ej guv'y gtg"vq"j cxg"dggp" f qpg"y kj "EV0"EV"uecpu"y gtg"r gthqto gf"y kj "qten"cpf" kqf kpcvgf"lk0'eqpvcuv"cpf"O TKuecpu"y kj "lk0'i cf qrkpkwo "ej grvg"wpugui"vj gtg"y cu"o o gf lecn'eqpvcupf lecvkp"vq"eqpvcu0"</p> <p>Hqmqy /vr" wo qt"cuuguuo gpw"qh"vj g"pgemlej guvcdqo gp"cpf" qvj gt" ctgcu"qh' npqy p" f kugcug"cvRtgtgcwo gpv'qt"pgy n' "uwwr gev'f" f kugcug"y gtg"vq"j cxg"dggp"r gt hqto gf"qpeg" gxgt { "qvj gt"e { eng"dgw ggp"Fc { u'43"cpf"4: "qt"uqpgt"ht"vj gtg'y cu'gxkf gpeg"qh'r tqi tguukxg" f kugcug" cpf" uq qwf"j cxg" wvkl gf" vj g" uco g" o gy qf qmri { "EV"qt" O TK" cpf" uecp" ces wvklqp" vgej pls wgu" kpenf lpi "wug"qt" pqpwwg"qh"lk0'eqpvcuv"cu"y gtg"v ugf" hqt"vj g" Rtgtgcwo gpv'cuuguuo gpw0"C"ej guv'z/tc { "qt"ungrvzn/z/tc { "vj cv'ergetn" f go qpustcvf" c'pgy " o gvcucvle"ngukqp"o c { "j cxg"dggp" wugf"vq" f qewo gpv'r tqi tguukqp"kp"lkw"qh"vj g"EV"qt"O TK uecpu0"</p> <p>Vwo qt"cuuguuo gpv'cv'vj g"HpcriXkuk'y cu'qpn { "p geguuct { "ht'pqv'f qpg"y kj kp"6"y ggmu"qh'rcuv" wo qt"cuuguuo gpv0</p> <p>EV"cpf"O TKces wvklqp"i wlf grkgu"ht"uwdlgeu'y kj "FVE"y gtg" f gvckgf"lp"vj g"Vgej plecn' Ukg"O cpwcn'r tqxkf gf"vq" gcej "ukgd { "vj g"ko ci lpi "ETQ" f guki pcvgf" d { "vj g"ur qpugt"vq" r gthqto "vj g"lpf gr gpf gpvtgxky "qh"vj g"wo qt"cuuguuo gpw0</p> <p>Tumor Assessments for Subjects with MTC:</p> <p>Y kj kp"6"y ggmu'r tkqt"vq"vj g"htuv'f qug"qh'ngpvcvpkd"cf o kpkntcvkqp."rtgtgcwo gpv'wo qt" cuuguuo gpw"vulpi "EV"qh"vj g"pgemlej guv'lr grklu"cpf"qvj gt"ctgcu"qh'npqy p" f kugcug"qt"pgy n' "uwwr gev'f" f kugcug."cpf" gkj gt"O TK"qh"vj g"cdf qo gp"y kj "cpf"y kj qw'eqpvcuv"qt"EV"qh' cdf qo gp"y kj "vkr ng"r j cug"ko ci lpi "qh"vj g"rkxgt."y gtg"vq"j cxg"dggp"r gthqto gf0"Vj g" uecpplpi "ugs wpeg"ht"EV"qh"vj g'cdf qo gp"y kj "vkr ng"r j cug"ko ci lpi "qh"vj g"rkxgt"y cu"vq" j cxg"dggp"pqpeqpvcuv"EV"qh"vj g'rkxgt" hqmqy gf" d { "KX"eqpvcuv"lp'gevkqp"cpf" ctv'lecn'r j cug" ko ci lpi "qh"vj g'rkxgt."hqmqy gf"ko o gf levgn" d { "EV"qh"vj g"pgem"ej guv'cdf qo gp"cpf"r grklu0 Kp"cf f kqkp"vq"uecpu"qh"vj g'cdf qo gp."uecpu"qh"vj g"pgem"r grklu"cpf" qvj gt"ctgcu"qh'f kugcug" o c { "j cxg"dggp" f qpg"y kj "O TKlpugcf"qh"EV."dw"vj g"ej guv'gxcmcvkqp"y cu"vq"j cxg"dggp" f qpg"y kj "EV0"EV"uecpu"y gtg"vq"j cxg"dggp"r gthqto gf"y kj "qten"cpf" kqf kpcvgf"lk0' eqpvcuv"cpf"O TKuecpu"y kj "lk0'i cf qrkpkwo "ej grvg"eqpvcuv"wpugui"vj gtg"y cu"o o gf lecn' eqpvcupf lecvkp"vq"eqpvcu0"</p> <p>Hqmqy /vr" wo qt"cuuguuo gpw"qh"vj g"pgemlej guvcdqo gp"cpf" qvj gt" ctgcu"qh' npqy p" f kugcug"cvRtgtgcwo gpv'qt"pgy n' "uwwr gev'f" f kugcug"y gtg"vq"j cxg"dggp"r gt hqto gf"qpeg" gxgt { "qvj gt"e { eng"dgw ggp"Fc { u'43"cpf"4: "qt"uqpgt"ht"vj gtg'y cu'gxkf gpeg"qh'r tqi tguukxg" f kugcug" cpf" uq qwf"j cxg" wugf"vq" g" uco g" o gy qf qmri { "EV"qt"O TK"cpf" uecp"ces wvklqp" vgej pls wgu" kpenf lpi "wug"qt" pqpwwg"qh"KX"eqpvcuv"cu"y gtg"v ugf" hqt"vj g"r tgtgcwo gpv' cuuguuo gpw0"Cu"cvRtgtgcwo gpv"y g'cdf qo gp"o wu"j cxg"dggp" gxcmcvgf" vulpi "gkj gt" O TKy kj "cpf"y kj qw'eqpvcuv"qt"EV"qh"vj g"cdf qo gp"y kj "vkr ng"r j cu g"ko ci lpi "qh"vj g"</p>

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<p>rkxgt0"C"ej guv/z/tc{ "qt'u ngrgcn/z/tc{ "vj cv' ergetn{ "f go qpuntcvf "c"pgy "o gxcvckle"ngukqp o c{ 'j' cxg'dggp'wugf "q"i qewo gpvr tqi tguukqp'lp'hw'qh'vj g'EV'qt'O T Kuecpu0"</p> <p>Vvo qt"cuuguuo gpv'cv'vj g'Hkpcn'Xkuk'y cu'qpn{ 'p geguuct{ "h'pqv'f qpg'y kj lp'6'y ggmu'qh'vj g rcuv'wo qt'cuuguuo gpv0'</p> <p>EV'cpf "O T Kces wukukqp"i wlf grkpgu'hqt'uwdlgeu'y kj "O VE'y gtg"f gvcckgf "lp'vj g'Vej plecn' Ukg'O cpvcr'r tqxkf gf "q" gcej "ukg'd { "vj g"lo ci lpi "ETQ"i guki pcvgf "d { "vj g"ur qpuqt"vq" r gthqto "vj g'lpf gr gpf gpvt'gxlgy "qh'vj g'wo qt'cuuguuo gpv0'</p> <p>For All Subjects:</p> <p>Vj g'r tghgtgf "v' r g'qh'tcf kqmi lecn'uecp'y cu'c'f kci pquke's wcrk{ "ur kcn'qt"o wnkf gvevqt'EV" uecp'y kj "lk0'cpf "qtcn'eqpvtcu0"Kk0'eqpvtcu'y cu'eqpvtclpf lecvgf."vj g'ej guv'gxcnckvqp" y cu'r gthqto gf "y kj "pqeqpvtcu'EV'cpf "vj g'cdf qo gp lr grkku'gxcnckvqp" f qpg'y kj "O TK' Dtclp'uecpu."y j gtg'tgs vktgf."y gtg'r gthqto gf "y kj "lk0'eqpvtcu'wulpi "gkj gt"O TK'y kj " i cf qnplkw "ej grv'g'eqpvtcu'wrguu'vj gtg'y cu'c'eqpvtclpf lecvgf"vq'vj ku'v' r g'qh'eqpvtcu'qt" EV0'Nqy "f qug'EV'uecpu" hqo "c'eqo dlpckvqp"RGV/EV'uecppgt"y gtg'pqv'ceeg r wdrng' Wmtcuwpgf "cpf "hwqtkp/3: /4/hwqtk/4/f gqz { /F/i meug/r quktqp" go kukqp" vqo qi tcr j { " *HF I /RGV+ "y gtg'pqv'vq"j cxg" dggp'wugf "hqt"tcf kqi tcr j le"wo qt'cuuguuo gpv0'K'y cu" tgeqo o gpf gf "vj cv' ur kcnlo wnkf gvevqt"EV"dg" r gthqto gf "y kj "c"≤"7"o o "eqpvi wqwu" tgeqpuv'ekqp"cn qtkj o 0'K'dqf { "O T Kuecpu'y gtg'r gthqto gf."vj g' { "y gtg'tgeqo o gpf gf "vq" dg" f qpg"y kj "eqpvi wqwu"urlegu'q h"≤"7"o o 0'K'uwde wcpqgwu"o cuugu"qt"pqf gu"y gtg" r cr cdrng" *g0 0'dwnf "+cpf "y gtg"cuuguucdrng"d { "dqj "erplecn'cpf "tcf kqi tcr j le"vej pls wgu." vj g'tcf kqi tcr j le"EV"qt'O TK'vej pls wg'y cu"vq"j cxg" dggp'wugf "hqt"vj g'cuuguuo gpv'qh' vcti gv'cpf "pqpvti gv'ngukpu0'Cuuguuo gpv'y cu'r gthqto gf "cv'vj g'ukg"d { "cr r tqr tlcvg" s wcrk'gf "r gtuqppgn"tcf kqmi ku'lp'eqplwepvqp'y kj "erplecn'lp'xgukv cxtq'+cpf "tguwmu'y gtg" tgeqtf gf "qp'vj g'cr r tqr tlcvg'ETH'ci g*u0'</p> <p>"</p> <p>Target and Nontarget Lesions:</p> <p>Measurable Disease:</p> <p>Ceeqtf lpi "vq'vj g" qtki lpcn' TGEKUV'cpf "o qf kkgf "TGEKUV."o gcuwcdrg" f kugcug'y cu " f ghkpgf "d { "vj g'r tgupeg'qh'cv'ngcu'qpg'o gcuwcdrg'ngukqp0"C"o gcuwcdrg'ngukqp'ku'qpg'y cv' eqwf "dg'ceewtcvgn"o gcuwgf "lp'cv'ngcu'qpg'f kco gvg' "cv'ngcu'32"o o "lp'NF" d { "ur kcn'EV" uecp"qt"cv'ngcu'42"o o "d { "ucpf ctf "EV"qt'O TK'vej pls wgu"qt"d { "erplecn" o gcuwtgo gpv0' Hqt"vj ku'tlcn'cm'uwdlgeu'y gtg'tgs vktgf "vq"j cxg'cv'ngcu'q pg'ngukqp"≥"30"eo "lp'NF" hqt" pqpni o r j "pqf gu'cpf "≥"40"eo "lp'NF" hqt"n{ o r j "pqf gu"vj cv'y gtg'ugtkcmf "cpf "ceewtcvgn" o gcuwcdrg'ceeqtf lpi "vq"o qf kkgf "TGEKUV" wulpi "gkj gt"EV"qt'O TK'Ngukpu"vj cv'j cf " tgegkxgf "GDTV"o wuv"j cxg"uj qy p" gxf gpeg'qh'r tqi tguukg" f kugcug'dcugf "qp"o qf kkgf " TGEKUV"vq'dg'f ggo gf "c'vcti gv'ngukqp0'K'c'ngukqp'y cu'cuuguucdrng"d { "dqj "tcf kqmi lecn'cpf " erplecn'vej pls wgu."tcf kqmi lecn'vej pls wgu"y gtg"vq"j cxg" dggp'wugf 0'K'vj g'o gcuwcdrg' ngukqp'y cu'c'h{ o r j "pqf g."k'o wuv"j cxg'o gcuwgf "cv'ngcu'42"o o "lp'NF 0'K'c'ukpi ng'ngukqp" y cu'kf gpv'kkgf "cu'vj g"vcti gv'ngukqp."c'e { "vqmi lecn'qt"j kuqmi lecn'eqphko cvkqp"qh'vj { tqkf " ectekpqo c'y cu'tgs vktgf 0'</p> <p>Vcti gv'ngukpu"y gtg'ugrgev'f "q p"vj g"dcuku'qh'vj gkt"uk g'*NF "+cpf "vj gkt"uwkcdkxk{ "hqt" ceewtcv'gtr gcv'f "o gcuwtgo gpv0'Cm'lo gcuwcdrg'ngukpu"wr "vq"o czlo wo "qh'7"ngukpu" r gt"qti cp'ukg"cpf "32"ngukpu"lp"vqcn'vj cv'y gtg'tgr tgupeg'vckg'qh'cm'lp'xqkxgf "qti cp'ukkgu" y gtg'kf gpv'kkgf."o gcuwgf."cpf "tgeqtf gf "cu'vcti gv'ngukpu"qp'vj g'cr r tqr tlcvg'dcugrkpg'ETH' Vj g'uco g'ngukpu'y gtg'o gcuwgf "cpf "tgeqtf gf "cv'cn'hmjy /wr "ko g'r qkpwo"</p> <p>Cm'qj gt"ngukpukkgu"qh'f kugcug"y gtg'kf gpv'kkgf "cu"pqpvti gv'f kugcug."tgeqtf gf "qp"vj g" dcugrkpg" ETH'cpf "cuuguuf" cv'cn'hmjy /wr "ko g" r qkpwo" cu" opq" ej cpi glucdrng.ö" ödcugrvlf kucr r gctgf.ö"öwpgs wlxqecni'y qtuqplpi.ö"qt"öqj gt"ur gekh{ +0""Cp{ "pgy "ngukpu" f gvev'f "tcf kqi tcr j lecnf "y gtg'tgeqtf gf "qp'vj g'cr r tqr tlcvg'pqpvti gv'ngukqp'ETH'ci g0'</p> <p>Clinically Evaluable Disease:</p> <p>Enklecni "gxcnckdrng'ngukpukkgu"qh'f kugcug"*uej "cu'r ngwtcn'gh'wukqp."cukgu."ewcpqgwu"</p>
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	<p>n{ o r j cpi kku+ y gtg" k f gpv hkf " cpf " tgeqtf gf " cu" pqpvti gv" ngukpu" qp" y j g" cr r tqr tkvg" dcugr kpg"ETHO"O gcuwtgo gpv"qh'y gug'ngukpu'y gtg"pqv'tgs vkt gf "dw'y j g{ "y gtg"hmny gf " cpf "tgeqtf gf "cu"öpq'ej c pi g luvdr g.ö"öcdugpvlf kucr r gctgf.ö"öwpgs wkgced'y qtugplpi .ö"qt öqy j t"ur gekh{ :ö"kp'y j g'r quwtgcvo gpv'ETHuO"C f f kkpccm{ . "y j g"cr r gctcpeg"qh'cp{ "pgy" erplecm{ "gxcnwdng'ngukpu'y cu'tgeqtf gf "qp'y j g'r quwtgcvo gpv'ETHuO"</p> <p>Bone Scan: C"r tgv gcv gpv'dqpg"uecp"y cu'r gthqto gf "y kj kp"8"y ggmu'r tkqt"vq"cf o kpkutcvkp"qh'y j g" hktuv'f qug"qh'ngpxcvkpkd0"Vj g'dqpg"uecp"y cu'tgr gcvgf "gxgt { "6'e{ ergu'dgwy ggp"Fc{ u'43"cpf " 4: "cpf "cv'y j g"Hkpcn'Xkuk'wprguu'c'uecp"y cu'qdvkpgf "nguu'y cp"38"y ggmu'r tkqt"vq'y j g"Hkpcn' Xkuk+0"C"dqpg"uecp"y cu'tgs vkt gf "cv'y j g"eqphko cvqt { "ko g'r qkp'lp"cmiwdlgeu'y kj "ET" cpf "RT0"HF1 "RGV" y cu'pqv'cp'ceeg r vdr g"cngtpevkg"vq'y j g'uecpf ctf " ; o "vgej pgkwo " r qn{ r j qur j qpvcg"dqpg"uecp0" Y j qrg"dqf { "O TKo c { "j cxg"dgpp"wgf "cu'cp"cngtpevkg"vq" dqpg"uecpu"cv'r tgv gcv gpv'cpf "hmny /wr "h"wgf "eqpukngpv" cetquu'cnr'tgs vkt gf "ko g' r qkp'lp0"Dqpg"uecpu'y gtg"wgf "vq" f gvev'pgy "dqp { "ngukpu0"Kic"pgy "ngukp"cr r gctgf "qp" c " hmny /wr "dqpg"uecp. "k'y cu'tgeqo o gpf gf "y j cv'z/tc{ . "EV. "qt "O TKdg"wgf "vq"eqphko "y j g" o cki pcv'pcwtg"qh'y j g'pgy "tcegt"wr vng0"</p> <p>Evaluation of Tumor Response: Vwo qt" tgr qpug" y cu'gx cnwvgf " cv'y j g" uksg'd { " s wcn'hkf " r gtuqppgi" *c" tcf kqni kuv' kp" eqplwpevqp"y kj "y j g"erplecni'kpxguki cvqt+ wulpi "o qf hkf "TGEKUV0" Cp" KKT"qh"wo qt" tgr qpugu'y cu'cnq'r gthqto gf "d { "Eqtgrcd"Rctvpgtu0" Tgr qpugu'y gtg"cuuguuf "ceeqt f kpi "vq" o qf hkf "TGEKUV. "f gckrgf "kp"cp"Kpf gr gpf gpv'ko ci kpi "Tgxky "Ej ctvgt. "y j lej "y cu'kuwgf " r tkqt "vq"y j g'uctv'qh'y j g'lpf gr gpf gpv'tgxky 0"Kpxguki cvqtu'y gtg'tgs vkt gf "vq"r tqxkf g"eqr kgu" *r tghgtcdn{ "kp" f ki kcn'hqto cv"qh'ko ci gu'hqt"wo qt"cuuguu gpv"EV. "O TK'dqpg"uecpu"vq" c " egptcn'fcekks{ 0"C"o cpwcn'f gckkpi "tgeqo o gpf gf "uecpkpi "r etco gvgu. "ko ci g"j cpf r kpi . " cpf "uj k r r kpi "y cu'r tqxkf gf "vq" gcej "ukg0"Vj ku'f qewo gpv'ku'hqecvgf "kp"y j g'v'kcn'o cuvt "hkg0" Vj g"KKT" gxcnwcqp"y cu'wgf "hqt"y j g'r tko ct { "cpf "ugeqpf ct { "ghlece { "cuuguu gpv"qh'y j g" uwf { 0"</p> <p>Kp"qt f gt "hqt"y j g"UF "f guki pcvqp"vq"dg"i kxgp"ht"dguv'qxtcm'tgr qpug" *DQT+ "cv'ngcu'pqp" r quwtgcvo gpv'o gcuwtgo gpv'o wuv'j cxg'o gv'y j g"UF "etkgtlc" c'o kpo wo "qh'9"y ggmu'chgt " hktuv'uwf { "f twi "cf o kpkutcvkp0"C"ET"qt "RT"cuuguuf "cv'c"o kpo wo "qh'9"y ggmu'chgt "uctv' qh'tgcvo gpv'y kj qww'c"eqphko cvkp"qh'RT"qt "ET"cv'ngcu'6"y ggmu'hvgt "d { "hmny /wr "uecpu. " dw"j cxkpi "c"uudugs wgpv'r tqi tguukxg" f kugcug"cuuguu gpv."y cu'eqpukf gtgf "UF "hqt"y j g"dguv' tgr qpug0"J qy gxgt. "ET"qt "RT"cuuguuf "nguu'y cp"9"y ggmu'chgt "uctv'qh'tgcvo gpv'y kj "c' uudugs wgpv'r tqi tguukxg" f kugcug"y cu'eqpukf gtgf "r tqi tguukxg" f kugcug"ht"y j g"dguv'tgr qpug0"</p> <p>Eqphko gf "tgr qpugu" *ET"qt "RT+ "y gtg" f gvgto kpgf "cv'c'tgr gcv'wo qt" gxcnwcqp"cv'ngcu'6" y ggmu'chgt "dgkpi "hktuv'qdugtxgf "d { "gkij gt "y j g" kpxguki cvqt "qt "KKT"y j q"o cf g"y j g" kpkkn' cuuguu gpv0"Vj g"pgz'v'gxcnwcqp"ht"r qvcpv'cn'eqphko cvkp"qh'y j g" hktuv'RT"qt "ET"eqwv " j cxg"y ckxgf "wpv'v'v' g"pgz'v'tgi wctn{ "uej gf wrgf "wo qt"cuuguu gpv0"Vwo qt"cuuguu gpv'cv' y j g"Hkpcn'Xkuk'y cu'qp n{ "pgeguuct { "h'p'qv'f qpq"y kj kp"6"y ggmu'qh'y j g'ncu'v'tgcvo gpv'e{ eng0" C"ej guv'z/tc{ "qt"ungv'cn'z/tc{ "y j cv'engctn{ "f go qpuxcvgf "c'pgy "o gxcu'cve'ngukp"o c { "j cxg" dgpp"wgf "vq" f qewo gpv'r tqi tguukp. "kp" hkw'qh'y j g"EV"qt "O TKuecpu0"</p> <p>Tumor Assessments if Subjects Discontinued from Study Treatment Without Progressive Disease: Ki' uwdlgeu' f kueqpvkpgf "htqo "uwf { "tgcvo gpv'y kj qww'r tqi tguukxg" f kugcug" cpf "f kf" pqv' y kj f tcy "eqpugpv"wo qt"cuuguu gpv'y gtg'r gthqto gf "cv'y j g"Hkpcn'Xkuk"y kj kp"52" f c { u'qh' y j g'ncu'uwf { "f twi "cf o kpkutcvkp+ h'p'qv'r gthqto gf "y kj kp"y j g'r tgxkqwu"6"y ggmu. "cpf "y j g" gxgt { "5"o qpj u'uctv'kpi "htqo "y j g'ncu'wo qt"cuuguu gpv'wpv'v' tqi tguukxg" f kugcug. "f gcj . "qt" y j g'uctv'qh'cpqj gt "cpv'ecpeg"y j g'tcr { "qeewtgtf 0"</p> <p>Pharmacokinetics: Ngpxcvkpkd" r ncuo c" eqpegpvcvqp" f cvc"y gtg "eqmgev" c pf "dgecwug" qh" y j g" ur ctug" RM" uco r kpi "ko gu" y cu'r qq r g" y kj "y j g" kpxgukxg" RM" f cvc "htqo "y j tgg'r tgxkqwu" Rj cug"3" uwf kgu"kp" uwdlgeu'y kj "uqkf"wo qtu" *G92: 2/G266/323. "G92: 2/C223/324. "cpf "G92: 2/</p>
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	<p>I2: 3/325+!hqt"RM"o qf grlf gxmqr o gpv0"</p> <p>Hqt"j g"y q"uwlgeu"y j q"y gtg"tgcvgf"y kj "rgpxcvkpld"32"o i "DIF ."c"vqcn'qh"33"RM'dmqf" uco r ngu'y gtg"qdvckpgf"r gt"uwlgeu"Uco r ngu'y gtg"qdvckpgf"qp"E{erg"3"Fc{u"3"cpf"4"cpf" qp"E{erg"4"Fc{"30"Hqt"j g"tgo ckpki "337"uwlgeu"y j q"tgegkxgf"rgpxcvkpld"46"o i "S F ."c" vqcn'qh"plpg"RM'd mqf"uco r ngu'y gtg"qdvckpgf"r gt"uwlgeu"Uco r ngu'y gtg"qdvckpgf"qp" E{erg"3"Fc{u"3"cpf". .E{erg"4"Fc{"3."cpf"E{erg"5"Fc{"30"</p> <p>Pharmacokinetic/Pharmacodynamic (PK/PD):</p> <p>Cpcn'ugu"y gtg"eqpf wevgf"d{"o qf grkpi "vq"gzr nqtg"gzr quwtgótgur qpug"tgrvckpuj kr u"htq" uchgv{cpf"ghhece{0"</p> <p>Pharmacodynamics:</p> <p>Vj g"r nppgf"RF"cuuguu gpw"lpenwf gf"j g" hmqy kpi "ugtuo "dkqo ctngtu<"3+"j {tqlf" hmpvckp"vguu"lpenwf kpi "hgg"V6"cpf"VUJ =4+vj {tqi mqdwkp"FE"uwlgeu"qpn{=5+ ecrekqplp"cpf"EGC"O VE"uwlgeu"qpn{=6+dkqo ctngtu"qh'cr qr vaku"Ecur "59."E{vqE." cpf"O 52+cpf"7+r tqvgqo g"ECH"dkqo ctngtu"qh"cpk qj gple li tqy j "hcevtu."lpenwf kpi ej go qnkp."gpf qj grkwo "hmpvckp."cpf"kvgtngwnk lko o wpg"tgr qpug=Rj cto ceqf {pco le" cpf"wo qt'tgur qpug'tgrvckpuj kr u'y gtg"gzr nqtg 0"</p> <p>Pharmacogenomics:</p> <p>Vj g"r nppgf"r j cto ceqi gpqo le"RI +cuuguu gpw"lpenwf gf"j g" hmqy kpi <"3+"wo qt" o wcvkpu"uqo cve"FP C"ugs wpeg"xctkcpw"lp"BRAF, H-, K-"cpf"N-Ras,"cpf"RET/PTC1, 2,"cpf"3"cuuguu"lp"htgo ctkp/hkzgf"r ctchkp/go dgf f gf"HHRG+"wo qt"vuuv"cp f"4+" i gto rkp"FP C"ugs wpeg"xctkcpw"lp"i gpgu"lpxqngf"lp"CF O G"cu'y gni'cu"RET"cpf"FP C" xctkcpw"pgct"FOXEl"rs965513+"cpf"NKX2-1"rs944289+"cuuguu"ltqo "uwlgeu'dmqf" uco r ngu"Rj cto ceqi gpqo le"cpf"v wo qt" tgr qpug"tgrvckpuj kr u"y gtg"r nppgf"vq" dg" gzr nqtg 0"</p> <p>Safety:</p> <p>Uchgv"cuuguu gpw"eqpukvgf"qh"o qpkqtkpi "cpf"tgeqtf kpi "cmf'cf xgtug"gxgpw"CGu+"cpf" ugtkqu"cf xgtug" gxgpw"UCGu+eqpego kcpv"o gf lecvkpu."tgi wct"o qpkqtkpi "qh" j go cvqri {."dmqf"ej go knt {."cpf"wtkp"xcnwg"r gtlkf le"o gcuwgo gpv"qh'xkcn'uki pu." GEQI "r gthqto cpeg"ucwu."P[J C"cuuguu gpw."grgevtqectf kqi tco u"GEI u+gej qectf kqi tco u=cpf"r gthqto cpeg"qhr j {ulecn'gzco kpcvku0"</p>
Bioanalytical Methods:	<p>Rruu c"eqpepvtcvkpu"qh'r ctgpf"ftwi "y gtg"s wcpvklgf"d{"rks wlf"ej tqo cvqi tcr j {lo cuu" ur gevto gt {lo cuu"ur gevto gt {"NE IO UIO U+"o gj qf qri { "wukpi "c"r tgxkquw{"xcnkf cvg" cuu{0"</p>
Statistical Methods:	<p>F cv"qdvckpgf"vr "vq"j g"fcv"ewqhh"fcvg"qh"33"Cr t"4233"ctg"lpenwf gf"lp"j ku"tgr qtv0"kp" cf f kkp."f cv"ht"cuuguu gpvlgxgpw"y kj "c'uctvf" cvg"r tktq"vq"j g"ewqhh"fcvg"j cv" eqpvkpgf"chgt"j g"ewqhh"fcvg"ctg'cnq "lpenwf gf" g0 0"CG.e qpego kcpv"o gf lecvkpp." r tqegf wtg."cpf"uwf {"tgcvo gpv0"Hwtj gto qtg."uwxkcn'fcv"eqngevgf"chgt"j g"ewqhh"fcvg" ctg'cnq"lpenwf gf"lp"qtf gt"vq"qdvckp"uwlgeu"uwxkcn'ucwu"cv'j g"ewqhh"fcvg0"</p> <p>Vj g"Kp'gpv"vq"VtgcV"KV+Rqr wcvkpp"lpenwf gf"cm'uwlgeu"y j q"tgegkxgf"cv'rgcu'ppg"t qug" qh"j g"uwf {"f twi "cpf"y cu'j g"r tko ct {"cpcn'uku'ugv'wugf"htq"ghhece{ "cpcn'ugu0"Vj g" Ghhece{"Gxcnwdrg"Rqr wcvkpp"lpenwf gf"cm'uwlgeu"y j q"tgegkxgf"cv'rgcu'ppg"t qug"qh"j g" uwf {"tgcvo gpv"j cf "c"dcugnkp"cpf"cv'rgcu'ppg"r quwtgcvo gpv"wo qt"tgr qpug"gxcnvckp." cpf"hwtkngf"lpenwkp"etkgtk"3."4."5."cpf"60"Vj g"Uchgv"Rqr wcvkpp"lpenwf gf"cm'uwlgeu" y j q"tgegkxgf"cv'rgcu'ppg"t qug" qh"uwf {"f twi "cpf"j cf "cv'rgcu'ppg"r quwtgcvo gpv"uchgv" cuuguu gpv'cpf"y cu'j g"r tko ct {"cpcn'uku'ugv'htq"cm'uchgv"fcv0"Vj g"uchgv"cpn'ugu"y gtg" eqpf wevgf"d{"utcw"FE"cpf"O VE+ugr ctevg"cp f"y kj "j g"y q"utcv"eqo dlpf 0"kp" cf f kkp."j g"y q" F VE"uwlgeu"gpqmgf"wpf gt"j g"qtki kpcn' r tqveqn"y kj "lplcn'DIF" f qukpi "y gtg"cpn{ gf"ugr ctevg"cu'c'i tqwr."cn uq"vqi gj gt"y kj "cm'qj gt"uwlgeu"cu"j g" qxgtcm'uo o ct {"ht"j g"uwf {0"</p>

Eisai

"

	<p>cugugf "hqt"j { r gtvukqp"cpf "r tqvkwk0"Qvj gt"uchgv"r ctco gvtu"y gtg"cnq"i tcr j lecnf" cugugf "hqt"j g'tgrvukpuij k'y kj "ngpxcvkpld"gzr quwtg'y j lej "lpenwf gf "y g'CGu"qh'f kettj gc." hcvk wg."f getgcugf "crr gvkqg."pcwugc."cpf "y gki j v'f getgcug."cpf "rdqtcvqt { "r ctco gvtu"y hqt" j go cvqmi { "<"j go cvqetk."j go qi mdkp."y j kg'drqf "egm."pgwtqr j kn"cpf "r rvgv'eqwpu=" hqt" dkqej go kwt { "<" cmekpg"r j qur j cvug." dkkwdkp." cmekpg" co kqvtcpuhgtcug." cur ctvvg co kqvtcpuhgtcug."ugt wo "cndwo kp."ugt wo "etgcvkpg."cpf "qvcn'r tqvklp-0"</p> <p>Tgrvukpuij k'u'y gtg"gzr mgtgf "dgw ggp"ngpxcvkpld"gzr quwtg"cpf "y g'hmqy kpi "dkqo ctngtu" dkqej go lecn'o ctngtu"y { tqi mdkp"hqt"y g'F VE"eqj qtv"ecrekqpkp"cpf "EGC"hqt"y g'O VE" eqj qtv" cpf " y { tqkf " hapevqp" vgu'tguw nu" *Hgg"V6 " cpf " Hgg"VUJ " hqt" cni'wd lgeu=" dkqo ctngtu" qh' cr qr vuku" *Ecur " 59." E { vqE." cpf "O 52+" cpf "ugt wo " r tqvgo g' ECH" dkqo ctngtu." lpenwf kpi " cpi kqi gple li tqy yj " hrevqt." ej go qnkpg." gpf qvj gkwo " hapevqp." kpvtrgwukp l'ko o wpg'tgur qpug'hrevqtu."cpf "qvj gt'hrevqtu0"</p> <p>Uchgv" f cvc."lpenwf kpi "CG"cpf "rdqtcvqt { "xcnwg."r tlqt"cpf "eqpeqo kcpv'o gf lecvkqp."cpf " uwf { "f twi "gzr quwtg."GEI "cpf "gej qectf kqi tco "hkp kpi u"y gtg"cpn { j gf "wukpi "f guetk vxg" uc'vukku" *p."o gcp."o gf kcp."ucpf ctf " f gxcvqp." tci g+" hqt" eqp'p'wqu" o gcuwgu." cpf " lpekf gpeg"eqwpu" *pwo dgt "cpf "r gtegpv" hqt"ecvgi qtlecn'xctkcdrgu0"</p>
Results:	<p>Analysis Sets</p> <p>Vj g'KVVRqr wrcvqp"cpf "y g'Uchgv" Rqr wrcvqp"y gtg'yj g'uco g'lp"y ku'uwf { "<"c'vqcn'qh'339" uwdlgeu"tgcvgf =7: "uwdlgeu"lp"y g'F VE"eqj qtv"y q" f qugf "32"o i "DKF"cpf "78" f qugf "46" o i "S F "+"cpf "7: "uwdlgeu"lp"y g'O VE"eqj qtv" f qugf "46"o i "S F 0"Vj g'Ghlee { "Gxcnwdrg" Rqr wrcvqp"lpenwf gf "77"uwdlgeu"lp"y g'F VE"eqj qtv"cpf "72"uwdlgeu"lp"y g'O VE"r qr wrcvqp *, 7" "cpf": 7" "qh'yj g'KVVRqr wrcvqp."tgr gevkgu { +0</p> <p>Subject Disposition"</p> <ul style="list-style-type: none"> Qh'yj g'7: "uwdlgeu"lp"y g'F VE"eqj qtv"y g'y q"uwdlgeu"y j q"y gtg'f qugf "y kj " ngpxcvkpld" 32"o i " DKF" f kueqpvpwgf " yj g' Vtgcvo gpv' Rj cug" f wg" vq" f kugcug" r tqi tguukqp"cpf "f kf "pqv'eqpvpwg"tgcvo gpv'lpv"y g'Gzvgpukp"Rj cug0"Qh'yj g'78" uwdlgeu" y kj " F VE" f qugf "y kj " ngpxcvkpld" 46"o i " S F." 45" *63' "+" eqpvpwgf " tgcvo gpv'lp"y g'Gzvgpukp"Rj cug"cp f "55" *7; ' "+" y gtg'f kueqpvpwgf "Htqo " yj g' tgcvo gpv'lp"y g'Vtgcvo gpv' Rj cug0"Ukvggp"uwdlgeu" *4; ' "+"lp"y g' S F "f qulpi " i tqvr "f kueqpvpwgf "f wg"vq" f kugcug"r tqi tguukp0"C"vqcn'qh'37"uwdlgeu" *49' "+"lp" yj g' S F "f qulpi " i tqvr "f kueqpvpwgf "f wg"vq" CGu="36" *47' "+" y kj "CGu"cu"vj g' r tko ct { " tgcup" hqt" f kueqpvpwcvqp" cpf "3" uwdlgev" y kj " uwdlgev" ej qleg" cu"vj g' r tko ct { " tgcup" hqt" f kueqpvpwcvqp" cpf "y kj " cp" CG"cu"y g'ugeqpf ct { " tgcup0 Qpg"cf f kkpnci"uwdlgev" f kueqpvpwgf "f wg"vq"uwdlgev" ej qleg" cpf " cpqy gt" uwdlgev y kj f tgy "eqpugp'cu"r tko ct { " tgcup" hqt" f kueqpvpwcvqp0" Qh'yj g'7: "uwdlgeu"y kj "O VE."4; " *6; ' "+"eqpvpwgf "tgcvo gpv'lp"y g'Gzvgpukp" Rj cug" cpf "52" *73' "+" y gtg'f kueqpvpwgf "Htqo " yj g' Vtgcvo gpv' Rj cug0" Hknggp" uwdlgeu" *47' "+" f kueqpvpwgf "f wg"vq" f kugcug"r tqi tguukp0"C"vqcn'qh'36"uwdlgeu" *46' "+" f kueqpvpwgf "f wg"vq" CGu="35" *44' "+" cf "CGu"cu"y g' r tko ct { " tgcup" hqt" f kueqpvpwcvqp" *lpenwf kpi "3"uwdlgev" y kj "dqy "y g' r tko ct { "cpf "dqy gtö"tgcup"lp" cf f kkp"vq"y g' r tko ct { " tgcup_ "hqt" f kueqpvpwcvqp"pqvgf "cu"dgkpi "f wg"vq" CGu" cpf "3"uwdlgev" y kj "uwdlgev" ej qleg"cu"y g' r tko ct { " tgcup" hqt" f kueqpvpwcvqp" cpf " y kj " cp" CG"cu"y g'ugeqpf ct { " tgcup0"Qpg"cf f kkpnci"uwdlgev" f kueqpvpwgf "f wg"vq" uwdlgev" ej qleg"cu"y g' r tko ct { " tgcup" hqt" f kueqpvpwcvqp0 Cv'yj g'vko g'qh'f cvc"ewqhi" hqt"y ku'EUT" *33 "Cr t"423 3+."74"uwdlgeu" *45"lp"y g' F VE"eqj qtv"cpf "4; "lp"y g' O VE"eqj qtv"y gtg"uwnitgegkxkpi "tgcvo gpv'lp"y g' Gzvgpukp"Rj cug0 <p>Exposure</p> <ul style="list-style-type: none"> Hqt"y g'F VE"eqj qtv"y g'o gf kcp" f wcvqp"y cu"5; 50" f c { u" *tci g<"42"vq"75: " f c { u"0"Vj g'o gcp"pwo dgt "qh'e { engu"y cu"330."y g'o gf kcp"pwo dgt "qh'e { engu"y cu" 350." cpf " yj g' o kpo wo " cpf " o czko wo " pwo dgt " qh' e { engu" y cu" 3" cpf " 3; .

	<p>tgur gevkgñ(0"</p> <ul style="list-style-type: none"> Hqt"vj g"O VE"eqj qtv"vj g"o gf kcp"f wcvkqp"y cu"48602"f c{u"ticipi g<"3 5"vq"769" f c{u"0"Hqt"vj g"O VE"eqj qtv"vj g"o gcp"pwo dgt"qh'e{ ergu"y cu"; 09."vj g"o gf kcp" pwo dgt"qh'e{ergu"y cu"320."cpf"vj g"o kpo wo "cpf"o czko wo "pwo dgt"qh'e{ ergu"y cu"3"cpf"42."tgur gevkgñ(0" <p>Efficacy</p> <ul style="list-style-type: none"> Ghhece{ "gpfr qkpw"y gtg"cuugugf "kp"vj g"K/V"Rqr wcvkqp"wpiguu"qvj gty lug"pqvgf 0" Vj g"r tko ct{ "gpfr qkpv"qh"vj g"uwf { "y cu"vj g"QTT"dcugf "qp"vj g"cuuguo gpw'd{ "vj g" KKT"0"Vj g"QTT"y cu"72" "kp"vj g"F VE"eqj qtv"cpf"58" "kp"vj g"O VE"eqj qtv"0" Ugeqpf ct{ "ghhece{ "tguwu"htq"vj g"O VE"eqj q t v'y gtg"cu"hmmy u0"Cv'c'o kpo wo " hmmy /w"r gtlkf "qh"36"o qpj u."vj g"o gf kcp"guvko cvg"qh"RHU"dcugf "qp"KKT"y cu" 340"o qpj u0"Vj g"8/o qpj "RHU"tcvg"y cu"9: " "cpf"vj g"34/o qpj "RHU"tcvg"y cu" 77" 0"Vj g"o gf kcp"QU"dcugf "qp"Mcrrp/O gkt"cpn{uku"eqwnf "pqv"dg"tgrkdn{ " guvko cvgf 0"Vj g"qxtcm"uwtxkcn"tcvg"y cu": 8" "cv"34"o qpj u"cpf"y cu"9: " "cv'dqvj 3: "cpf"46"o qpj u0"Vj g"o gf kcp"hmmy /w"vko g"y cu"380"o qpj u0"Dcugf "qp" cuuguo gpw'd{ "vj g"KKT."vj g"DET"y cu"; 5" "cpf"vj g"EDT"y cu"9: " 0"Vj g"o gf kcp" f wcvkqp"qh"tgur qpug"htq"uwlgeu"y kj "c"DQT"qh"ET"qt"RT"*p?4; +y cu"340" o qpj u0"Vj g"o gf kcp"vko g"vq"tgur qpug"htq"uwlgeu"y kj "c"DQT"qh"ET"qt"RT"kp" vj g"Ghhece{ "Gxcwcdng"Rqr wcvkqp"*p?4: +y cu"50"o qpj u0" Ugeqpf ct{ "ghhece{ "tguwu"htq"vj g"O VE"e qj qtv'y gtg"cu"hmmy u0"Cv'c'o kpo wo " hmmy /w"r gtlkf "qh": "o qpj u."vj g"o gf kcp"guvko cvg"qh"RHU"y cu"; 0"o qpj u0"Vj g" 8/o qpj "RHU"tcvg"y cu"89" "cpf"vj g"34/o qpj "RHU"tcvg"y cu"68 " 0"Vj g"o gf kcp" QU"dcugf "qp"Mcrrp/O gkt"cpn{uku"eqwnf "pqv"dg"tgrkdn{ "guvko cvgf 0"Vj g"qxtcm" uwtxkcn"tcvg"y cu"98" "cv"34"o qpj u"cpf"y cu"pqv"guvko cdng"cv3: "cpf"46"o qpj u0" Vj g"o gf kcp"hmmy /w"vko g"y cu"330"o qpj u0"Dcugf "qp"cuuguo gpw'd{ "vj g"KKT." vj g"DET"y cu": 2" "cpf"vj g"EDT"y cu"86" 0"Vj g"o gf kcp"f wcvkqp"qh"tgur qpug"htq" uwlgeu"y kj "c"DQT"qh"ET"qt"RT"*p?43+y cu"pqv"tgcj gf 0"Vj g"o gf kcp"vko g"vq" tgur qpug"htq"uwlgeu"y kj "eqphko gf "c"DQT"qh"ET" qt"RT"kp"v j g"Ghhece{ " Gxcwcdng"Rqr wcvkqp"*p?3; +y cu"50"o qpj u0" Hqt"vj g"F VE"eqj qtv"vj g"QTT."dcugf "qp"cuuguo gpw'd{ "vj g"KKT."y cu "7; " "kp" uwlgeu"y kj "r tkt"XGI H"vcti gvgf "tgcvo gpv"*p?39+cpf"y cu"68" "kp"uwlgeu" y kj qw'r tkt"XGI H"vj gter { "p?63+0"Hqt"vj g"O VE"eqj qtv"vj g"QTT"y cu"vko kkt" kp"uwlgeu"y kj "r tkt"XGI H"vj gter { "p?48+eqo r ctgf "y kj "vj qug"y kj qw'r tkt" XGI H"tgcvo gpv"*p?55+*57" "cpf"58" .tgur gevkgñ(0" Ugeqpf ct{ "ghhece{ "gpfr qkpw"y gtg"cuugf "qp"vj g"kpvguvi cvqtu0"cuuguo gpw'd{ Vj g"ugeqpf ct{ "ghhece{ "gpfr qkpw"vj qy gf "pq"o clqt"t hgtgpegu"dcugf "qp"vj g"vy q" uqwtgu"qh"cuuguo gpw'd{ <p>Pharmacokinetics/Pharmacodynamics</p> <ul style="list-style-type: none"> Vj g"RM"qh"ngpxcvkpld"kp"vj g"eqo dlpgf "F VE"cpf"O VE"eqj qtv"y gtg"vko kkt"vq" vj cv"qdugtvgf "kp"vj g"Rj cug"3"uvwf lgu0" "P q" f kgev"tgrcvkpuj kr" qh" ngpxcvkpld gzr quwtg"*uvgcf { "uvcg"WE+eqwf"dg"qdugtvgf"y kj "QTT"cpf"RHU" <p>Safety</p> <ul style="list-style-type: none"> Cm'uwlgeu"kp"dqj "eqj qtv"gzr gtlkepgf "cv'hcuv"qpq"v tgcvo gpv/go gti gpv"CG" *VCGG+" cpf" cv' ngcu"qpq"VCGG"tg r qtv"cu"tgcvo gpv'tgrcvf 0"Vj g" o quv htgs wgpw{ "tgr qtv"VCGG"*62" "qh"uwlgeu"kp"gkj gt"eqj qtv"y gtg"tctj gc." j {r gtvgpukp."rtqvgpwtlc."h'vki wg."y gli j v'f getgcugf."f getgcugf"cr r gvkg."pcwugc." j gcfcej g."eqwi j ."cpf" f {ur j qpk0" Vqzlekkgu"y gtg"o cpci gf" d{ "uwf { "f twi " kpwttrv kqp" cpf" uwdugs wgpv" f qug" tgf wcvkqp"vq"tgf weg"vj g"pgeguks{ "htq"uwf { "f twi "y kj f tcy cn0"VCGG"u'hc f lpi "vq" uwf { "f twi "y kj f tcy cn'qeewtg f "kp"37"uwlgeu"*48" +kp"vj g"F VE"eqj qtv"cpf"kp" 36"uwlgeu"*46" +kp"vj g"O VE"eqj qtv"0"
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	<ul style="list-style-type: none"> • VGCu"qh'ur gekn'kpvgu'lpemf gf "j { r gtvgpukqp"cpf "rtqvgkwk."y j lej "ctg"CGu npqy p"q"dg"cuuqekvgf "y kj "tgcvo gpv'y kj "ngpxcvkpld"cpf "qvj gt"XGI H'vcti gvgf "ci gpv0"J { r gtvgpukqp"y cu"q"dg"tgcvgf "y kj "cpvj { r gtvgpukg"tgcvo gpv'cpf "y kj uwf { "f twi " lpvgttwr vkp" cpf " uwdugs wgpv" f qug" tgf vevkp0 "Vj ku" o cpci go gpv uej go g"cr r gctgf "q"dg"ghgevkxg0"Vj g"o clqtkv "qh'gxgpw'y gtg"l tcf g"3"qt"l tcf g"40"O quv'qh"y j g"gxgpw'qh"j { r gtvgpukqp"cpf "rtqvgkwk."y gtg"o cpci gf "y kj qw uwf { "f twi "y kj f tcy cn'y kj f tcy cn'ltqo "y j g"uwf { "tgcvo gpv'f wg"vq"j { r gtvgpukqp qeewttgf "lp"qpg"uwdlgev'gcej "lp"y j g"VE" cpf "O VE"eqj qtv0" Rtqvgkwk"tcf "q" uwf { "f twi "y kj f tcy cn'lp"y j tgg"uwdlgeu'lp"y j g"VE"eqj qtv'cpf "pq"uwdlgeu'lp"y j g"O VE"eqj qtv0"P q"l tcf g"6"qt"l tcf g"7"j { r gtvgpukqp"qt"rtqvgkwk"gxgpw'y gtg tgr qtv'gf 0"J { r gtvgpukqp."cu'cp"UCG."qeewttgf "lp"y q"uwdlgeu'lp"y j g"VE"eqj qtv' cpf "lp"qpg"uwdlgev'lp"y j g"O VE"eqj qtv0"Rtqvgkwk."cu'cp"UCG."qeewttgf "lp"qpg' uwdlgev'lp"y j g"VE"eqj qtv'cpf "pqv'lp"cp { "uwdlgev'lp"y j g"O VE"eqj qtv0" • Kp"y j g"VE"eqj qtv."y j g"o quv'ltgs wgpw' "tgr qtv'gf "l tcf g"5"VGCu"qeewttkpi "lp" cr r tqzko cvgn "32" "qh'uwdlgeu"y gtg'y gki j v'f getgcugf .f kcttj gc."j { r gtvgpukqp." cpf "rtqvgkwk0"Uz"l tcf g"6"VGCu"qeewttgf <"j { r qecre go lc."j { r gtnre go lc." cdcukc."cpf "cewg"o { qectf kn'lphtevkp"qpg"tgr qtv'gcej + "cpf "w y q"tgr qtv'qh' r wv qpct { "go dqruo 0"Kp"y j g"O VE"eqj qtv"y j g"o quv'ltgs wgpw' "tgr qtv'gf "l tcf g"5" VGCu"qeewttkpi "lp"cr r tqzko cvgn "32" "qh'uwdlgeu'qh'uwdlgeu"y cu'f kcttj gc0 Hlxg"l tcf g"6"VGCu"qeewttgf <"co { rug"lpetgcugf . "nr cug"lpetgcugf . "rpgwo qplc" cur kcvkp. "gzhqkcvkxg"tcuj . "cpf "ceekf gpv'lxg'f qug"qpg"tgr qtv'gcej +0" • UCGu"dqy "hvcn'cpf "pqphvcn"qeewttgf "lp"4: "uwdlgeu"6: " +lp"y j g"VE"eqj qtv' cpf "lp"52"uwdlgeu"73" +lp"y j g"O VE"eqj qtv0"UCGu'y j lej "qeewttgf "lp"cv'ngcu' y q" uwdlgeu" y gtg" cu' hmqy u< "k p" dqy " eqj qtv." r wv qpct { " go dqruo " cpf " f gj { f tcvkp= qpn { " lp" y j g" F VE" eqj qtv." ny gt" cdf qo kpcl' r clp." j { r gtvgpukqp." j { r gtvgpukqp."cpf "ectf kce"lckwtg=cpf "qpn { "lp"y j g"O VE"eqj qtv."cdf qo kpcl' r clp." npi "lp'hgkvq."f getgcugf "cr r gvkg."cpf "r tgo cwtg'o gpqr cwug0" • F gcj .f wtkpi "tgcvo gpv'qt'y kj lp"52"fc { u'qh'y j g'rcuv'f qug'qh'uwf { "f twi . "qeewttgf "lp"cv'qh'qh'ugxgp"uwdlgeu'lp"y ku'uwf { <"y j tgg"lp"y j g"VE"eqj qtv'cpf "hwt"lp"y j g" O VE"eqj qtv0"Vy q" f gcj u."qpg"lp"y j g"VE"eqj qtv'cpf "qpg"lp"y j g"O VE"eqj qtv." y gtg"tgr qtv'gf "f wg"q"enkplecn'r tqi tguukqp"qh'f kugcug0"Hlxg" f gcj u'y gtg"tgr qtv'gf " cu'cp"qweqo g"qh'cp"UCG<"y q "lp"y j g"VE"eqj qtv'cpf "y j tgg"lp"y j g"O VE"eqj qtv0 Y kj "y j g"gxncvkvq"qh'y j g'kphqto cvkp"cxckndrg"htq"y j g"ug'hxg" f gcj u"cuuqekvgf " y kj "UCGu"qp g"qh'y j g"y q" f gcj u"lp"y j g"VE"eqj qtv"y cu'f vgto kpgf "q"dg" cuuqekvgf "y kj "r tqi tguukqp"qh'f kugcug."cpf "y j g"qvj gt" f gcj "y cu'f wg"q"ectqkf" ctvt { "j go qttj ci g'y j lej "y cu'tgrvgf "q"necn'lphtco o cvkp"qh'y j g"xguun'ltqo "c" r tgxkqwu"tcej gquqo { 0"Kp"v j g"O VE"eqj qtv."y j g"v j tgg" f gcj u"cuuqekvgf "y kj "UCGu"y gtg" f vgto kpgf "q"dg"cuuqekvgf "y kj "r tqi tguukqp"qh'f kugcug0"Vj gtghqtg" qh'y j g'ugxgp" f gcj u"qeewttkpi "f wtkpi "tgcvo gpv'qt'y kj lp"52"fc { u'qh'y j g'rcuv'f qug' qh'uwf { "f twi . "ulz"y gtg"cuuqekvgf "y kj "r tqi tguukqp"qh'f kugcug."cpf "qpg"y cu'y j g" qweqo g"qh'cp"UCG0" • P q"enkplecn { "ko r qtvcpv'ej cpi gu"lp"o gcp"j go cvqni { "cpf "dkqej go kn { "xcnvgu" ltqo "dcugrkpg"q"y j g"gpf "qh'y j g"xctkqwu"e { engu'y gtg"qdugtxgf 0"t guwmu'htq'o gcp" r nvrgv'cpf "pgwtqr j kn'eqwpu'uj qy "c'u rki j v'f getgcukpi "vtgpf . "dw'y gtg'y kj lp" pqto cn'tepi g'ht"gcej " r ctco gvg0"Vj gug"tguwmu'y gtg"uggp"lp"dqy "F VE"cpf " O VE"eqj qtv0"Vj g'uj kn'cpcn'uku'ht"EVECG'i tcf gu'tgxgcrgf "pq"uj kn'qh'enkplecn' eqpegt'ht"j go cvqni { "qt'enkplecn'ej go kn { "r ctco gvgu'ltqo "dcugrkpg"q"y j g"gpf " qh'gcej "e { eng0"lpetgcugu"lp"rtqvgk"lp"y j g"vtkpg'y gtg"qdugtxgf "qxtg"ko g'y j lej "ku' eqpukngpv'y kj "y j g"qeewttgpeg"qh'r tqvgkwk"tgr qtv'gf "cu"VGCu0" • Enkplecn { "ko r qtvcpv'ej cpi gu"lp"o gcp"xkn'uki pu'ltqo "dcugrkpg"q"y j g"gpf r qkvu' cv'xctkqwu"xluku"y gtg"qdugtxgf 0"Drqf "r tguwtg"ej cpi gu"qeewttgf "cpf "y gtg" tgr qtv'gf "cu" VGCu" kn' f ggo gf " enkplecn { " ko r qtvcpv' d { " y j g" lpxgunk cvqt0 Ngpxcvkpld"tgcvo gpv'y cu'eqttgrvgf "y kj "cp"lpetgcug"lp"drqf "r tguwtg0"O quv'qh' y j g" lpetgcugu"lp" drqf "r tguwtg"qeewttgf "f wtkpi "y j g" htu'v' e { eng0" Chgt" y j g"
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	<p>lpetgcug."f qy py ctf "tgpfu"lp"dqj "UDR"cpf "F DR"y gtg"qdugtxgf "r tko ctkf "f wg"qj tgcvo gpv" y kj " cpvj { r gtvgpukxg" o gf kecvkpu" cpf lqt " f qug" lpgttwr vkp" qt tgf wevkp0'</p> <ul style="list-style-type: none"> P q"enklecni "tgrxcpv"lpf lpi u"lqt"ej cpi gu"lp"GEI u"qt"gej qectf kqi tco u"y gtg qdugtxgf 0"
Conclusions:	<p>Kp" yj ku" Rj cug" 4" uwf { " lp" uwdlgeu" y kj " gkj gt" O VE"qt " F VE" Kf kpg/353" tghtcevt { . wptgugewdng" f kugcug" y kj " gxf gpeg"qh" f kugcug" r tqi tguukp. "ngpxcvkpd" y cu" cf o lpkngt gf "cv" c" uctv kpi " f qug" qh" 46" o i " S F . " gzeqr v" hqt " y q " uwdlgeu" y j q " y gtg " tgcvgf " cv" y j g " f qug" qh" 32" o i " D K F . " cu" r nppgf " lp" y j g " qtki lpcn" r tqv qeqr0" Uwdlgeu" eqpvkpwgf " tgcvo gpv" wvkv" f kugcug" r tqi tguukp. " f gxmqr o gpv" qh" wpccegr vdrng " qzlek { . " f gcj . " qt" y kj f tcy cni" qh" eqpugp0" Vj g" o gf lcp" f wcvkqp" qh" tgcvo gpv" y cu" 5; 50" f c { u" t cpi g < " 42" q" 75: " f c { u" hqt " y j g " F VE " eqj qtv" cpf " 4860" f c { u" t cpi g < " 35" q" 769" f c { u" hqt " y j g " O VE " eqj qtv0" C vj g " vko g " qh" f c v " ewqhl hqt" y j ku" EUT " * 33 " Crt " 4233 + " 74 " uwdlgeu" * 45 " lp" y j g " F VE " eqj qtv" cpf " 4; " lp" y j g " O VE " eqj qtv" y gtg " uvmi tgegk kpi " tgcvo gpv" lp" y j g " Gz vgpukp " Rj cug0'</p> <p>Vj g " RM " qh" ngpxcvkpd " lp" y j g " F VE " cpf " O VE " eqj qtv" y gtg " uko krt " vj " y j cv" qdugtxgf " lp" y j g " Rj cug" 3 " uwf lgu0" P q " f kgev" tgrvkpuj k " qh" ngpxcvkpd " gzt quw" g " ugc { " uvcg " C WE + " eqwf " dg" qdugtxgf " y kj " QTT " cpf " RHU0</p> <p>Ngpxcvkpd " u j qy gf " o gcpkpi hmi" cpvko qt " cevkv { " lp" dqj " y j g " F VE " cpf " O VE " j kxqni lecn" eqj qtv0" Vj g " r tko ct { " gpf r qkp v" qh" y j g " uwf { . " QTT " dcugf " qp" y j g " cuuguu gpw" d { " y j g " KKT . " y cu" 72' " lp" y j g " F VE " eqj qtv" cpf " 58' " kp" y j g " O VE " eqj qtv0" Uge qpf ct { " ghlece { " tguwmu" uj qy gf " y j g " QTT . " dcugf " qp" cuuguu gpw" d { " y j g " lpxgunki cvqtu " y cu" 75' " lp" y j g " F VE " eqj qtv" cpf " 6; ' " lp" y j g " O VE " eqj qtv0" F kht gpegu" dgw ggp " y j g " tguwmu" qdvkpgf " d { " y j g " KKT " cpf " y j g " lpxgunki cvqtu " y gtg " gzt gev" f . " cpf " qxgtcm " y j g " tguwmu" ctg " lp" eqpe qtf cpeg0" Qj gt " ugeqpf ct { " ghlece { " gpf r qkp v. " lpenv lpi " F E T . " E D T . " cpf " RHU . " cnuq " u j qy g f " cpvko qt " cevkv { " qh" ngpxcvkpd0" Hqt " y j g " F VE " eqj qtv" y kj " c " hmqy / w r " vko g " qh" 36" o qpj u . " y j g " o gf lcp " guko cvg" qh" RHU " y cu" 3408" o qpj u . " cpf " hqt " y j g " O VE " eqj qtv" y kj " c " hmqy / w r " vko g " qh" : 0 " o qpj u . " y j g " o gf lcp " guko cvg " qh" RHU " y cu" , 0 " o qpj u0" Hqt " y j g " F VE " eqj qtv" y kj " c " o gf lcp " hmqy / w r " vko g " qh" 3808" o qpj u . " y j g " qxgtcm " uwxkcn " tcv " y cu" : 8' " cv" 34" o qpj u " cpf " y cu" 9: ' " cv" dqj " 3: " cpf " 46" o qpj u0" Hqt " y j g " O VE " eqj qtv" y kj " c " o gf lcp " hmqy / w r " vko g " qh" 3308" o qpj u . " y j g " qxgtcm " uwxkcn " tcv " y cu" 98' " cv" 34" o qpj u " cpf " y cu" p qv " guko cdng " cv" 3: " cpf " 46" o qpj u0" O gcp lpi hmi" cpvko qt " cevkv { " y cu" cnuq " qdugtxgf " lp" uwdlgeu" y j q " j cf " r t g x l q w u n " dggp " tgcvgf " y k j " qj gt " r t k q t " X G I H " v c t i g v g f " c i g p u 0 " V j g " g h l e c e { " f c v " qdugtxgf " lp" y j ku" uwf { " eqo r c t g " h x q t c d n { " y kj " r w d i k u j g f " f c v " h q t " q j g t " c p v " X G I H " c i g p u 0 " Kp" y j ku" uwf { . " ngpxcvkpd " j cf " cp " ceegr vdrng " uchgv { " r tqhkg " hqt " u w d l g e u " y kj " t g h t c e v t { " y j { t q k f " e c p e g t 0 " P q " p g y " u c h g v { " e q p e g t p u " y g t g " q d u g t x g f 0 " " C u " r t g x l q w u n " u g g p " k p " q j g t " uwf lgu. " j { r g t v g p u k p " c p f " r t q v k p w t k c " y g t g " u c h g v { " e q p e g t p u " y kj " ngpxcvkpd . " cu" y kj " qj g t " f t w i u " y j c v " c h g e v " y j g " X G I H " u k i p c r k p i " r c v j y c { 0 " D q j " j { r g t v g p u k p " c p f " r t q v k p w t k c " f g x g n r g f " g c t n { " lp" y j g " eqwtug " qh" uwf { " f t w i " c f o l p k n t c v k p 0 " Kp" i g p g t c n { " j { r g t v g p u k p " y c u " y g m i e q p v t q m g f " y kj " cpvj { r gtvgpukxg" o gf kecvkpu" cpf lqt " f qug" lpgttwr vkp" qt " tgf wevkp" qh" ngpxcvkpd0"</p> <p>Vj ku" uwf { " uwi i guw" y j cv" r tqo kulpi " cpvko qt " cevkv { " cpf " cp " ceegr vdrng " uchgv { " r tqhkg " ecp " dg" cej lxxgf " wulpi " y j g " f qulpi " r ncp . " y j k e j " l p k l c v g u " y g t c r { " c v j g " o c z k o w o " v a n g t c v g f " f q u g " * O V F + " v " l p f w e g " y j g " d g u v " q x g t c m " w o q t " t g u r q p u g " c p f " y j g p " l p f k k f w e n k " g u " y j g " f q u g " d c u g f " qp " q d u g t x g f " u c h g v { " r c t c o g v g t u . " w u l p i " r n p p g f " f q u g " t g f w e v k p u " q h " n g p x c v k p d 0 " V j k u " f q u l p i " r n p " y k n i d g " g x c n c v g f " h m t j g t " l p " u w d u g s w g p v " c p f " r c t i g t " u w f l g u 0 " V j g " g h l e c e { " c p f " u c h g v { " q h " ngpxcvkpd " ku" ewt g p w { " d g l p i " v g u g f " l p " c " R j c u g " 5 " t k n l p " u w d l g e u " y kj " F V E 0 "</p>
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