

SYMDEKO™ (tezacaftor/ivacaftor; ivacaftor) Tablets

Table 4 lists responsive *CFTR* mutations based on (1) a clinical FEV₁ response and/or (2) *in vitro* data in FRT cells, indicating that tezacaftor/ivacaftor increases chloride transport to at least 10% of untreated normal over baseline. *CFTR* gene mutations that are not responsive to ivacaftor alone are not expected to respond to SYMDEKO except for F508del homozygotes.

Table 4: List of <i>CFTR</i> Gene Mutations that Produce <i>CFTR</i> Protein and are Responsive to SYMDEKO					
<i>E56K</i>	<i>R117C</i>	<i>A455E</i>	<i>S945L</i>	<i>R1070W</i>	<i>3272-26A→G</i>
<i>P67L</i>	<i>E193K</i>	<i>F508del*</i>	<i>S977F</i>	<i>F1074L</i>	<i>3849+10kbC→T</i>
<i>R74W</i>	<i>L206W</i>	<i>D579G</i>	<i>F1052V</i>	<i>D1152H</i>	
<i>D110E</i>	<i>R347H</i>	<i>711+3A→G</i>	<i>K1060T</i>	<i>D1270N</i>	
<i>D110H</i>	<i>R352Q</i>	<i>E831X</i>	<i>A1067T</i>	<i>2789+5G→A</i>	
*A patient must have two copies of the <i>F508del</i> mutation or at least one copy of a responsive mutation presented in Table 4 to be indicated.					