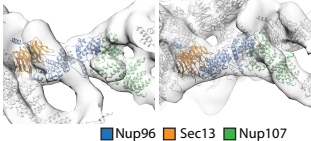

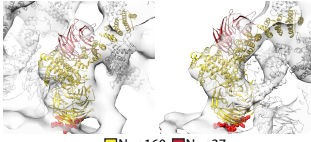

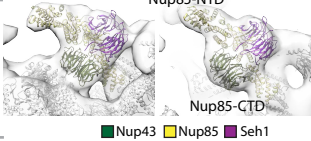

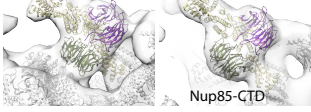
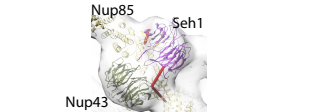

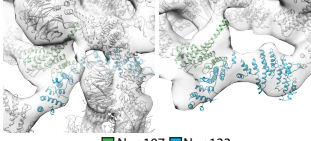

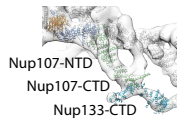
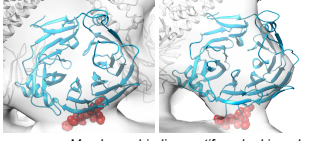

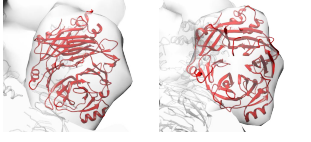

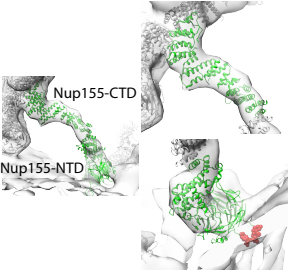
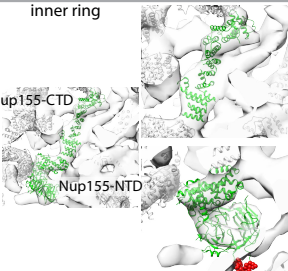
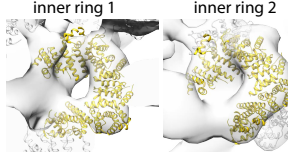
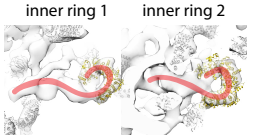
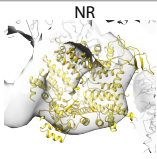
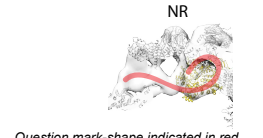


Component	Fit into the tomographic map	Localization	Orientation
Nup107-NTD/ Nup96/ Sec13	inner stem base NR outer stem base NR  ■ Nup96 ■ Sec13 ■ Nup107	Location: unambiguous <ul style="list-style-type: none"> restrained by known interactors within the Y-complex^{1,2,3,4} consistent with C2 symmetric localization to CR and NR^{5,6,7} consistent with measured stoichiometry⁸ 	Orientation: unambiguous <ul style="list-style-type: none"> consistent with known orientation within larger vertex structure^{1,9} identified by systematic fitting with very high confidence (Ext. Data Fig. 7) pronounced intrinsic asymmetry of template structure (Ext. Data Tab. 1)
Nup160/ Nup37	inner large arm CR outer large arm CR  ■ Nup160 ■ Nup37 <i>Predicted membrane binding motif marked in red</i>	Location: unambiguous <ul style="list-style-type: none"> restrained by known interactors within the Y-complex^{1,2,3,4} consistent with C2 symmetric localization to CR and NR^{6,7} consistent with measured stoichiometry⁸ 	Orientation: unambiguous <ul style="list-style-type: none"> consistent with known orientation within larger vertex structure^{1,9} identified by systematic fitting with very high confidence pronounced intrinsic asymmetry of template structure (Ext. Data Tab. 1) consistent with predicted membrane binding motif¹⁰ (Ext. Data Fig. 6b)
Nup85-NTD/ Seh1	inner small arm NR outer small arm NR Nup85-NTD  ■ Nup43 ■ Nup85 ■ Seh1	Location: unambiguous <ul style="list-style-type: none"> restrained by known interactors within the Y-complex^{1,2,3,4} consistent with C2 symmetric localization to CR and NR^{6,7} consistent with measured stoichiometry⁸ 	Orientation: unambiguous <ul style="list-style-type: none"> consistent with known orientation within larger vertex structure^{1,9} identified by systematic fitting with very high confidence pronounced intrinsic asymmetry of template structure (Ext. Data Tab. 1)
Nup85-CTD	 Nup85-CTD	Location: unambiguous <ul style="list-style-type: none"> restrained by known interactors within the Y-complex^{1,2,3,4} consistent with C2 symmetric localization to CR and NR^{6,7} consistent with measured stoichiometry⁸ 	Orientation: partially restrained <ul style="list-style-type: none"> consistent with known orientation within larger vertex structure^{1,9} but may rotate around long axis of the fold Homology model extending Nup85-NTD. The model has low confidence in exact helix positioning
Nup43	 Nup43 Nup85 Seh1	Location: unambiguous <ul style="list-style-type: none"> restrained by known interactors within the Y-complex^{1,11} consistent with C2 symmetric localization to CR and NR^{6,7} consistent with measured stoichiometry⁸ 	Orientation: unambiguous <ul style="list-style-type: none"> restrained by cross-linking MS data¹ consistent with local refinement using Haddock without restraints (see Methods)
Nup107-CTD/ Nup133-CTD	inner stem tip NR outer stem tip NR  ■ Nup107 ■ Nup133	Location: unambiguous <ul style="list-style-type: none"> restrained by known interactors within the Y-complex^{1,2,3,4} and head to tail contact to Nup160^{12,13} consistent with C2 symmetric localization to CR and NR^{5,6,7} consistent with measured stoichiometry⁸ 	Orientation: unambiguous <ul style="list-style-type: none"> restrained known connectivity to vertex and Nup133 structure^{1,9} consistent with systematic fitting pronounced intrinsic asymmetry of template structure (Ext. Data Tab. 1) 
Nup133-NTD	inner stem tip NR outer stem tip NR  <i>Membrane binding motif marked in red</i>	Location: unambiguous <ul style="list-style-type: none"> restrained by known interactors within the Y-complex^{1,2,3,4} and head to tail contact to Nup160^{12,13} consistent with C2 symmetric localization to CR and NR^{6,7} consistent with measured stoichiometry⁸ 	Orientation: partially restrained <ul style="list-style-type: none"> although restrained by its known membrane binding motif¹⁰ (Ext. Data Fig. 6b), the beta propeller might flip by 180 degree around an axis parallel to the nucleocytoplasmic transport axis. The orientation shown would be consistent with its connectivity to the Nup133-CTD (Ext. Data Fig. 6b) but the domain linker is flexible.
Elys-NTD	inner outer 	Location: unambiguous <ul style="list-style-type: none"> restrained by known interactors within the Y-complex^{1,3,14} consistent with asymmetric localization to NR^{6,15} and additional density observed in the tomographic map consistent with measured stoichiometry⁸ 	Orientation: partially restrained <ul style="list-style-type: none"> restrained only by the local asymmetry of the tomographic density that is highly consistent with the shape of the high resolution structure. A 180 degree flip of the structure around an axis parallel to the NE might be possible.

Component	Fit into the tomographic map connector between IR and CR/NR	Localization	Orientation
Nup155-NTD/ Nup155-CTD		<p>Location: unambiguous</p> <ul style="list-style-type: none"> • consistent with proximity to membrane, possibly transmembrane domain ¹⁶ (Ext. Data Fig. 1a) and Nup155 CTD • consistent with C2 symmetric localization to IR ⁶ • NTD identified by systematic fitting with very high confidence (Ext. Data Fig. 7); extension with CTD is highly consistent with tomographic density • consistent with observed weak interactions to CR and/or NR members (e.g. Nup214 and Sec13, Ext. Data Fig. 8) 	<p>Orientation: unambiguous</p> <ul style="list-style-type: none"> • NTD identified by systematic fitting with very high confidence (Ext. Data Fig. 7); extension with CTD is highly consistent with tomographic density • pronounced intrinsic asymmetry of template structure (Ext. Data Tab. 1) • consistent with experimentally determined membrane binding motif (Fig. 3b, d)
Nup155-NTD/ Nup155-CTD	<p>inner ring</p> 	<p>Location: suggestive</p> <ul style="list-style-type: none"> • suggestive assignment because of limited structural and biochemical information available for the IR members • consistent with proximity to membrane, and possibly transmembrane domain ¹⁶ (Ext. Data Fig. 1a) and Nup155 CTD • consistent with C2 symmetric localization to IR ⁶ • NTD identified by systematic fitting with very high confidence (Ext. Data Fig. 7); extension with CTD highly consistent with tomographic density 	<p>Orientation: unambiguous</p> <ul style="list-style-type: none"> • if suggestive position is correct, the tomographic density would not permit an alternative orientation because of the pronounced intrinsic asymmetry of template structure (Ext. Data Tab. 1) • NTD identified by systematic fitting with very high confidence (Ext. Data Fig. 7); extension with CTD highly consistent with tomographic density • consistent with experimentally determined membrane binding motif (Fig. 3b, d)
Nup205-NTD	<p>inner ring 1 inner ring 2</p> 	<p>Location: suggestive</p> <ul style="list-style-type: none"> • suggestive assignment because of limited structural and biochemical information available for the IR members • structurally homologous to Nup188 ¹⁸ that might substitute this position • consistent with C2 symmetric localization ^{5, 6} • NTD identified by systematic fitting with very high confidence (Ext. Data Fig. 7); extension with CTD consistent with question mark-shape observed in isolation ¹⁷ • consistent with measured stoichiometry ⁸ 	<p>Orientation: partially restrained</p> <ul style="list-style-type: none"> • might be flipped around an axis parallel to the NE plane • identified by systematic fitting with very high confidence (Ext. Data Fig. 7) <p>inner ring 1 inner ring 2</p>  <p><i>Question mark-shape indicated in red</i></p>
Nup188-NTD	<p>NR</p> 	<p>Location: suggestive</p> <ul style="list-style-type: none"> • suggestive assignment because of limited structural and biochemical information available for the IR members • structurally homologous to Nup205 ¹⁸ that might substitute this position • consistent with biochemical data (Fig. 3, Ext. Data Fig. 8), cross-linking MS data ¹, FRET data ¹⁹ • NTD identified by systematic fitting with very high confidence (Ext. Data Fig. 7); extension with CTD consistent with question mark-shape observed in isolation ¹⁷ • consistent with measured stoichiometry ⁸ 	<p>Orientation: partially restrained</p> <ul style="list-style-type: none"> • might be flipped around an axis parallel to the NE plane • identified by systematic fitting with very high confidence (Ext. Data Fig. 7) <p>NR</p>  <p><i>Question mark-shape indicated in red</i></p>

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