

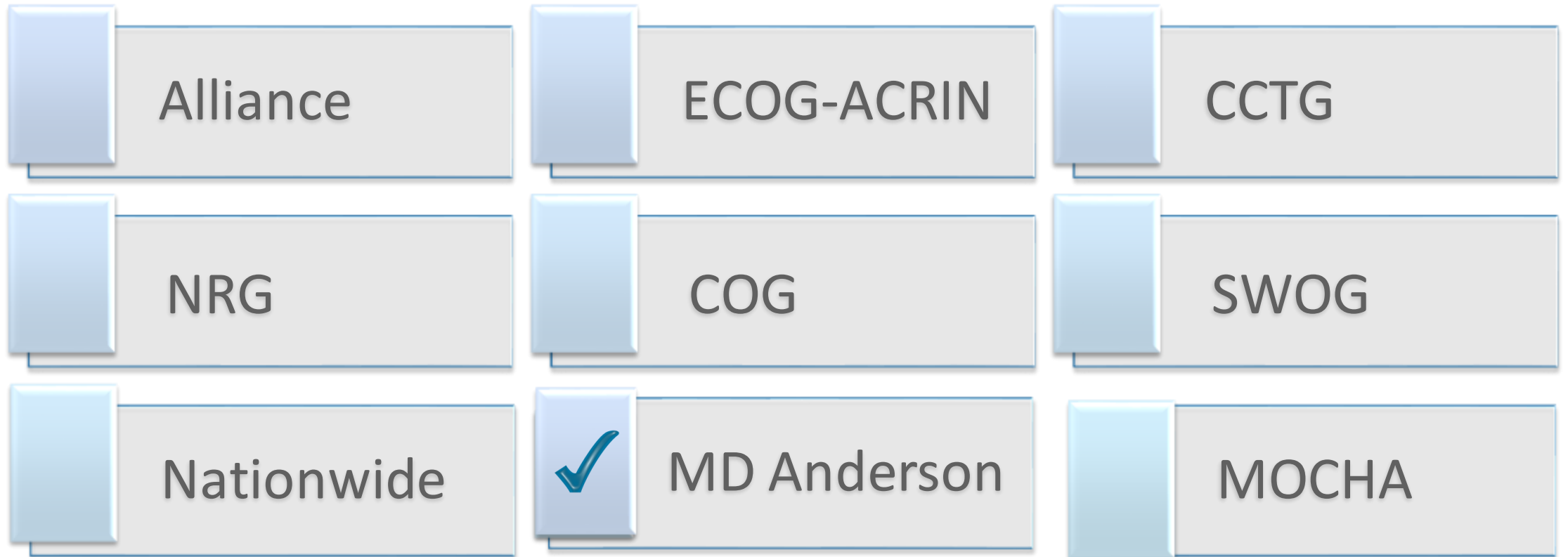
Precision Medicine Initiative (PMI) Committee Meeting

March 20, 2024

Agenda









- Role Call
- Project Updates
- Open Discussion

Stakeholder Representation



Project Updates

Project Updates – PMI Deliverables

	Deliverable	Target Release Date	Target Release Vehicle
	ComboMATCH changes (Disease fields, histology, behavior field code)	PROD Released: July 17, 2023	Screening Protocol EC Template v2.0
	Designated Labs for Combo	PROD Released: Sept 6, 2023	Screening Protocol EC Template v3.0 (UAT Release w/EA – 07/27/2023)
	Re-Screening ComboMATCH	PROD Released: Sept 27, 2023	MSRP Re-Screening EC Template v1.0 (UAT Release w/EA – 07/27/2023)
	Cohort Migration	PROD OPEN Released: Sept 28, 2023	Treatment Protocol Cohort Migration EC Template v1.0
		PROD EC Temp v1.0 Released: Oct 27, 2023	
	MyeloMATCH Tx Protocol Crossover (for S01, CTG01 and EA02)	n/a	Part of the existing Treatment Protocol workflow; no PRODUCTION OPEN Release required; the Westat team is prepared to support this workflow as Groups begin using it.
	Re-Screening MM	PROD RS EC Temp v2.0 Released: Nov 30, 2023	No PRODUCTION OPEN Release is required for MM Re-Screening activities; the Westat team is prepared to support this workflow as Groups begin using it. MSRP Re-Screening EC Template v2.0 (UAT Release w/SWOG – 10/6/2023; 2 nd UAT Release w/SWOG – 10/24/2023; 3 rd UAT Release to SWOG – 11/07/2023; PROD RS EC Template v2.0 target date 11/30/2023)
	MyeloMATCH Stratification	<i>PROD OPEN Release: February 12, 2023</i>	Part of existing Treatment Protocol Workflow.
	BETA PMI Screening Protocol ALS v2.0 New Forms: STMF, CLIA, PATH GROUP	Released: BETA Release: January 12, 2024	Target date for BETA Release for Groups to initiate UAT activities. (accompanied by BETA Screening Protocol Central Study ALS v2.0)
	PROD PMI Screening Protocol ALS v2.0 New Forms: STMF, CLIA, PATH GROUP	<i>PROD Release: April 22, 2024</i>	Target date for PRODUCTION Release for Groups to initiate implementation activities. (accompanied by PROD Screening Protocol Central Study ALS v2.0)

Project Updates - EC Template Release Schedule

	Template	Version	UAT Release Date	PROD Release Date	Details
✓	PMI Screening Protocol EC Template	V1.0	Released: 12/16/2022	Released: 02/15/2023	
✓	PMI Screening Protocol EC Template	V2.0	Released: 05/01/2023	Released: 07/17/2023	Includes CM changes (Disease fields, histology, behavior field code)
✓	PMI Screening Protocol EC Template	V3.0	Released: 07/27/2023	Released: 09/06/2023	Includes updates for Designated Labs (DLAP fields)
✓	<i>PMI Screening Protocol EC Template Fact Sheet</i>	v3.1	n/a	Released: 10/6/2023	Administrative version/release of the Fact Sheet to capture MyeloMATCH Field Help/Instructional Text for the 'Prior therapy name' field; to be used in conjunction with the PMI Screening Protocol EC Template v3.0.
✓	PMI Treatment Protocol EC Template	V1.0	Released: 12/16/2022	Released: 02/15/2023	
✓	<i>PMI Treatment Protocol EC Template Fact Sheet</i>	V1.1	n/a	Released: 11/16/2023	Administrative version/release of the Fact Sheet to provide a stand-alone TX Protocol-specific document allows Groups to easily access the Fact Sheet information; to be used in conjunction with the PMI Treatment Protocol EC Template v1.0.
✓	PMI MSRP Re-Screening EC Template	V1.0	Released: 07/27/2023	Released: 09/27/2023	<i>Supports ComboMATCH MSRP.</i>
✓	PMI MSRP Re-Screening EC Template	V2.0	Re-released: 11/07/2023 Re-released: 10/24/2023 Released: 10/06/2023	Released: 11/30/2023	<i>Supports MyeloMATCH MSRP; PROD release date confirmed.</i>
✓	PMI Treatment Protocol Cohort Migration EC Template	V1.0	Released: 09/18/2023	Released: 10/27/2023	Supports cohort migration activities for ComboMATCH.
✓	<i>PMI Treatment Protocol Cohort Migration EC Template Fact Sheet</i>	v1.1	Re-Release: 10/4/2023	Released: 10/27/2023	Administrative version/re-release of the UAT Fact Sheet to capture updated Field Help/Instructional Text for the 'Cohort migration assignment' field; to be used in conjunction with the UAT PMI Treatment Protocol Cohort Migration EC Template v1.0.

Group UAT Status : Screening Protocol ALS v2.0 (BETA)

- Beta ALSv2.0 Released on 1/15/24.
- Group UAT is underway
- Target date to complete Group UAT Testing (BETA) is **3/28/24**

ComboMATCH Priority 1 List

#	Protocol #	Protocol Title	Group	Activated	Current Status
1	EAY191	Molecular Analysis for Combination Therapy Choice (ComboMATCH)	ECOG-ACRIN	Yes	2/21: No Updates 3/20: No updates
2	EAY191-N4	Molecular Analysis for Combination Therapy Choice (ComboMATCH) EAY191-N4: A Randomized Trial of Selumetinib and Olaparib or Selumetinib Alone in Patients with Recurrent or Persistent RAS Pathway Mutant Ovarian and Endometrial Cancers A ComboMATCH Treatment Trial	NRG	Yes	2/21: No Updates 3/20: No updates
3	EAY191-E4	A ComboMATCH Treatment Trial ComboMATCH Treatment Trial E4: Nilotinib and Paclitaxel in Patients with Prior Taxane-Treated Solid Tumors	ECOG-ACRIN	Yes	2/21: No Updates 3/20: No updates
4	EAY191-N2	A ComboMATCH Treatment Trial EAY191-N2: Phase 2 Trial of Fulvestrant and Binimetinib in Patients with Hormone Receptor-Positive Metastatic Breast Cancer with a Frameshift or Nonsense Mutation or Genomic Deletion in NF1	NRG	Yes	2/7: Got comments from Amendment 1, need to update EC template & need updated protocol 2/21: Protocol Submitted – working on updates 3/20: No updates
5	EAY191-S3	Phase 2 Study of Paclitaxel + Ipatasertib in Taxane-Refractory Participants with AKT-Altered Advanced Non-Breast Solid Tumors	SWOG	Yes	2/7: Removed the SSN and full name and released 2/21: No updates 3/20: No Updates

ComboMATCH Priority 2 List

#	Protocol #	Protocol Title	Group	Activated	Current Status
6	EAY191-A6	A ComboMATCH Treatment Trial: FOLFOX in Combination with Binimetinib as 2nd Line Therapy for Patients with Advanced Biliary Tract Cancers with MAPK Pathway Alterations	Alliance	Yes	2/21: No Updates 3/20: No Updates
7	EAY191-A3	Palbociclib and Binimetinib in RAS-Mutant Cancers	Alliance	Yes	2/21: No Updates 3/20: Suspend Cohort 3.

ComboMATCH Priority 3 List

#	Protocol #	Protocol Title	Group	Current Status
8	EAY191-A2	Olaparib Plus Low-Dose Alpelisib for Breast Cancer	Alliance	DROPPED
9	EAY191-C1	Phase 2 Subprotocol of the Combination of a MEK Inhibitor and a Pan-RAF Inhibitor in Patients with Relapsed/Refractory Tumors Harboring Activating MAPK Pathway Mutations	COG	DROPPED
10	EAY191-E5	ComboMATCH Treatment Trial E5: A Randomized Phase II Study of AMG 510 (Sotorasib) with or Without Panitumumab in Advanced Solid Tumors	ECOG-ACRIN	<p>12/13: Almost complete with internal OPEN testing; contacting CTSU and Engineering for FFP Cohort Migration.</p> <p>1/10: No Updates</p> <p>2/7: Active</p> <p>2/21: No Updates</p> <p>3/20: No updates</p>
11	EAY191-N5	Molecular Analysis for Combination Therapy Choice (ComboMATCH) EAY191-N5: A Randomized Trial of Neratinib, A Pan-ERBB Inhibitor, Alone or in Combination with Palbociclib, a CDK4/6 Inhibitor, in Patients with HER2+ Gynecologic Cancers and Other Solid Tumors A ComboMATCH Treatment Trial	NRG	<p>12/13: Updated protocol submitted to CTEP on 12/4. Waiting to get test patients and working on the build.</p> <p>1/10: No Updates</p> <p>2/7: Released to production on 2/12</p> <p>2/21: No updates</p> <p>3/20: No updates</p>

MyeloMATCH Priority 1 List

#	Protocol #	Protocol Title	Group	Current Status
1	MYELOMATCH	Master Screening and Reassessment Protocol (MSRP) for the NCI MyeloMATCH Clinical Trials	SWOG	2/21: Completed, working through step to prep for activation – Westat is reviewing the bug found in OPEN...Westat will provide an update EOD 3/20: Pending is FFP; having trouble getting a volunteer for testing. If we do not identify someone soon, we will need to use an internal person.
2	MM1YA-S01	A Randomized Phase II Study Comparing Cytarabine + Daunorubicin (7 + 3) to (Daunorubicin and Cytarabine) Liposome, Cytarabine + Daunorubicin + Venetoclax, and Azacitidine + Venetoclax in Patients Aged 59 or Younger with High-Risk (Adverse) Acute Myeloid Leukemia; A MYELOMATCH Clinical Trial	SWOG	2/7: Ready for MATCHbox integration testing on Jan 29. 2/21: - Pending patient registration – until Westat bug is updated 3/20: Halfway through integration testing, have a few bugs they are working through.
3	MM1YA-CTG01	A Measurable Residual Disease (MRD) Driven, Phase II Study of Venetoclax Plus Chemotherapy for Newly Diagnosed Younger Patients with Intermediate Risk Acute Myeloid Leukemia: A Tier 1 MYELOMATCH Clinical Trial	CCTG	2/21: Waiting on new testing patients to complete testing – have a request in 3/20: Working on integration testing, received extra test patients. Put in a request with CTSU for a fix. Identified sites for FFP testing.
6	MM10A-EA02	A Randomized Phase II Study of HMA-Based Therapies for the Treatment of Older Adults with Newly Diagnosed FLT3-Mutated Acute Myeloid Leukemia: A myeloMATCH Treatment Trial	ECOG-ACRIN	2/7: Cohort/Stratum Assignment – Redoing the checklist – won't be ready for FFP by this week but working to meet that date. 2/21: Working with randostats team to get files in place to begin testing 3/20: Working on integration testing, hoping to complete in the next few days. Reached out to FFP testing, waiting on the sites to get back to us.

MyeloMATCH Priority 2 List

#	Protocol #	Protocol Title	Group	Current Status
4	MM2YA-EA01	Eradicating Measurable Residual Disease in Patients with Acute Myeloid Leukemia (AML) Prior to Stem Cell Transplantation (ERASE): A MyeloMATCH Treatment Trial	ECOG-ACRIN	2/21: No Updates 3/20: No Updates
5	MM10A-S02	A Randomized Phase II Study of Targeted Agents with the Oral DNMT1 Inhibitor Cedazuridine-Decitabine (ASTX727) for the Treatment of Newly Diagnosed TP53-Mutated Acute Myeloid Leukemia in Older Patients: A myeloMATCH Treatment Trial	SWOG	2/21: No Updates 3/20: No Updates
7	MM1MDS-EA03 (Concept)	ERADICATE MDS with TP53 Study: Efficacy of targeted Agents Combinations with Oral DNMT1 Inhibitor C-DEC (ASTX727) for Treatment of Higher Risk Myelodysplastic Syndromes with TP53 Mutations: A Non-Comparative, Parallel, Multi-Arm Phase 2 Study	ECOG-ACRIN	2/21: Withdrawn 3/20:

Open Discussion

- **Updates on EC Workflow:** Hoping to have conversations with ECOG to roll it out on their UAT soon. Third case requested for after the slot expires...they would require an amendment to protocol.
- Demo to the PMI Committee on the EC Workflow. Want to know how TAP works but how it's different between CM and MM.

Next Steps

Communication



Contact the PMI Mailbox for any PMI related questions & comments

pmistandards@nih.gov

The project team will respond within 48 hours with a response or a follow up



PMI Wiki

<https://wiki.nci.nih.gov/display/CDISC/Precision+Medicine+Initiative>

All presentations, recordings, minutes, project documents and releases will be posted on this wiki



**NATIONAL
CANCER
INSTITUTE**

www.cancer.gov

www.cancer.gov/espanol

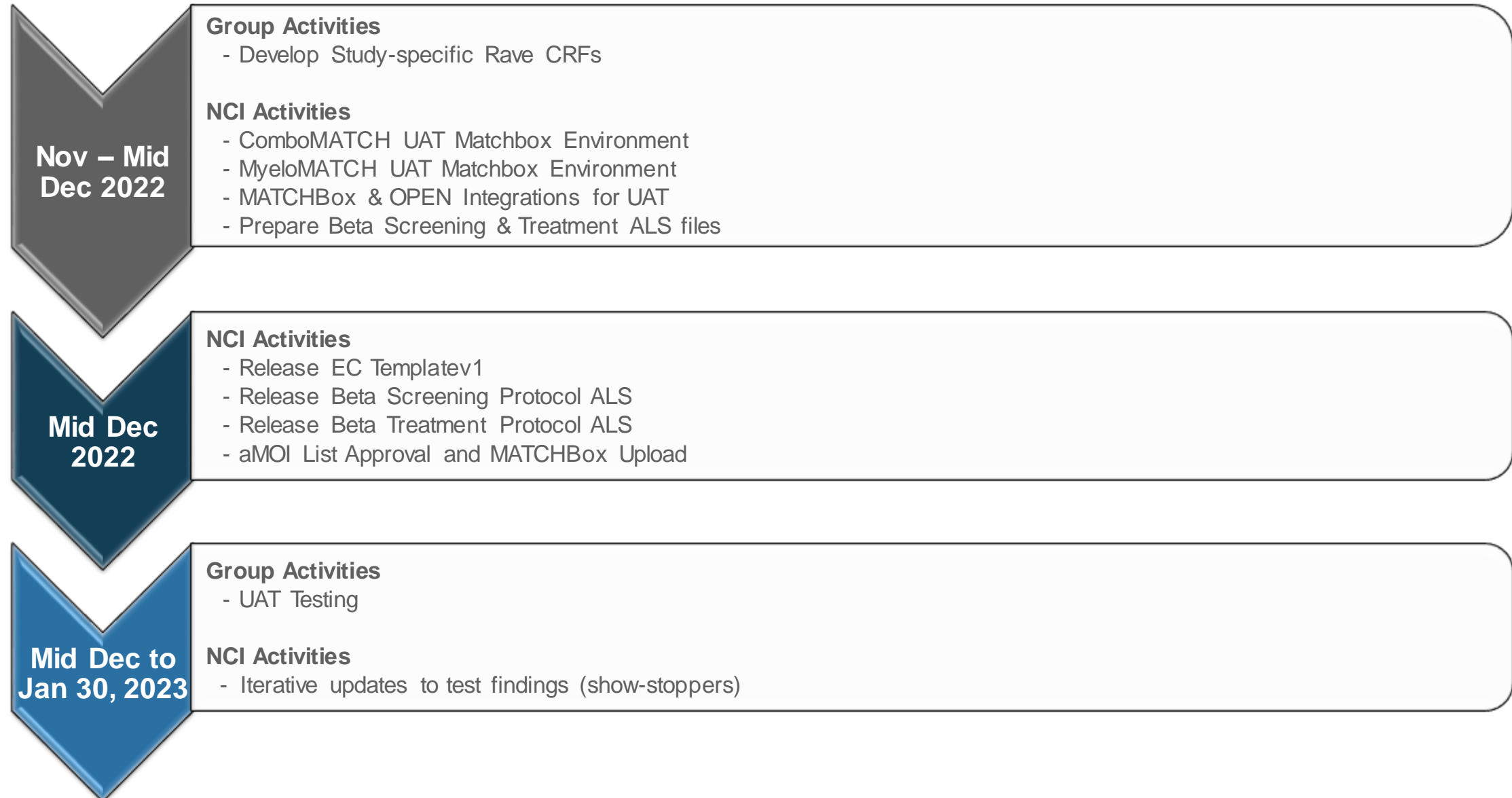
Appendix

CTEP CDISC Policy Governance Review

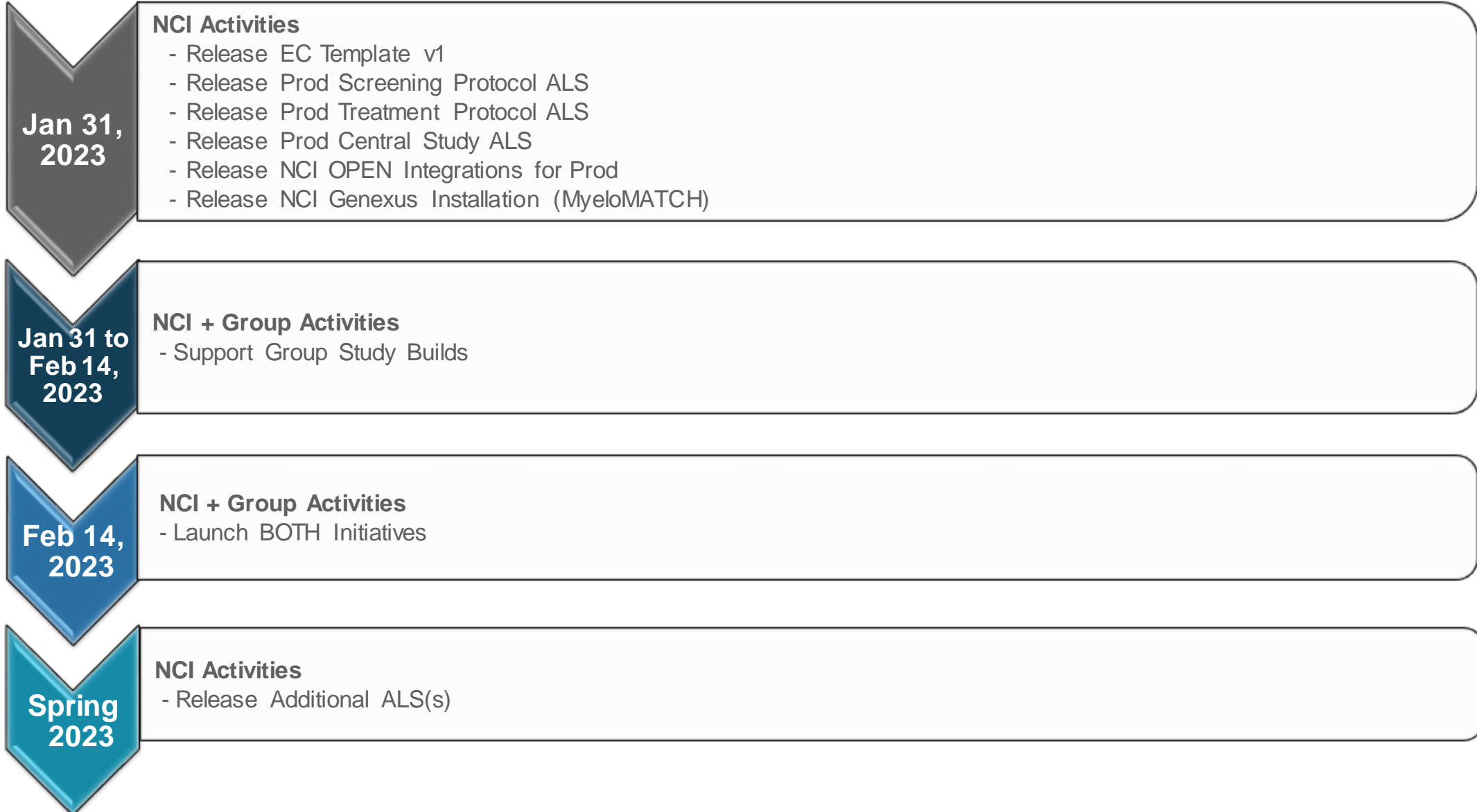
Compliance for Use of ALS Versions

- Standard forms built in the LPO's Global Library within **60 days** of production ALS release to reflect additions/updates included in the ALS
 - Example: Release ALSv7.X on November 1, LPOs would be expected to incorporate the ALSv7.X into their Global Library by January 1
 - ALS versions for new study builds
 - LPOs will be required to use the most current version of the ALS available at the time of each study build

Target Timeline



Target Timeline



ComboMATCH Priority 1 List

Study #	Document Number	Document Title	Group	Current Status	Next Steps	Primary Agent Name (P) Other Agent Name (O)
1	EAY191	Molecular Analysis for Combination Therapy Choice (ComboMATCH)	ECOG-ACRIN	8/15/2022 Approval on Hold	Revision 3 back from CIRB with approval. On hold for DTL, funding sheet and IND filing.	
2	EAY191-N4	Molecular Analysis for Combination Therapy Choice (ComboMATCH) EAY191-N4: A Randomized Trial of Selumetinib and Olaparib or Selumetinib Alone in Patients with Recurrent or Persistent RAS Pathway Mutant Ovarian and Endometrial Cancers A ComboMATCH Treatment Trial	NRG	8/15/2022 Approval on Hold	Revision 4 back from CIRB with approval. On hold for DTL, funding sheet and IND filing.	(P) Olaparib (AZD2281) (747856) (O) Selumetinib (AZD6244 hydrogen sulfate) (748727)
3	EAY191-E4	A ComboMATCH Treatment Trial ComboMATCH Treatment Trial E4: Nilotinib and Paclitaxel in Patients with Prior Taxane-Treated Solid Tumors	ECOG-ACRIN	8/26/2022 Approval on Hold	Revision 3 back from CIRB with approval. On hold for DTL, funding sheet and IND filing.	(P) Nilotinib (747599) (O) Paclitaxel (Taxol) (673089)
4	EAY191-N2	A ComboMATCH Treatment Trial EAY191-N2: Phase 2 Trial of Fulvestrant and Binimetinib in Patients with Hormone Receptor-Positive Metastatic Breast Cancer with a Frameshift or Nonsense Mutation or Genomic Deletion in NF1	NRG	8/26/2022 Approval on Hold	Revision 3 back from CIRB with approval. On hold for DTL, funding sheet and IND filing.	(P) Binimetinib (788187) (O) Fulvestrant (Faslodex) (719276)
5	EAY191-S3	Phase 2 Study of Paclitaxel + Ipatasertib in Taxane-Refractory Participants with AKT-Altered Advanced Non-Breast Solid Tumors	SWOG	8/29/2022 Approval on Hold	Revision 4 back from CIRB with approval. On hold for DTL, funding sheet and IND filing.	(P) Ipatasertib (781451) (O) Paclitaxel (Taxol) (673089)

ComboMATCH Priority 2 List

Study #	Document Number	Document Title	Group	Current Status	Next Steps	Primary Agent Name (P) Other Agent Name (O)
6	EAY191-A6	A ComboMATCH Treatment Trial: FOLFOX in Combination with Binimetinib as 2nd Line Therapy for Patients with Advanced Biliary Tract Cancers with MAPK Pathway Alterations	Alliance	Approval on Hold 9/14/2022	Rev 4 approved by CIRB on 10/24. CTEP currently reviewing stips.	(P) Binimetinib (788187) (O) OXALIplatin (Eloxatin) (266046) (O) 5-Fluorouracil (5-FU) (19893) (O)Leucovorin calcium (3590)
7	EAY191-A3	Palbociclib and Binimetinib in RAS-Mutant Cancers	Alliance	Approval on Hold 8/26/2022	Rev 2 sent to CIRB . Reviewed on 10/6, study team responding to stips	(P) Palbociclib (PD-0332991) (772256) (O) Binimetinib (788187)

ComboMATCH Priority 3 List

Study #	Document Number	Document Title	Group	Current Status	Next Steps	Primary Agent Name (P) Other Agent Name (O)
8	EAY191-A2	Olaparib Plus Low-Dose Alpelisib for Breast Cancer	Alliance	In Review 4/4/2022	Rev 3 recv;d 10/19, currently with LR. Looks like it'll be another disapproval. Not yet sent to CIRB for initial review.	(P) Olaparib (AZD2281) (747856) (O) Alpelisib (BYL719) (801658)
9	EAY191-C1	Phase 2 Subprotocol of the Combination of a MEK Inhibitor and a Pan-RAF Inhibitor in Patients with Relapsed/Refractory Tumors Harboring Activating MAPK Pathway Mutations	COG	In Review 9/29/2022	Protocol reviewed at PRC 10/20; CR is being put together	(P) Selumetinib (AZD6244 hydrogen sulfate) (748727) (O) DAY101 (TAK-580, MLN-2480) (800798)
10	EAY191-E5	ComboMATCH Treatment Trial E5: A Randomized Phase II Study of AMG 510 (Sotorasib) with or Without Panitumumab in Advanced Solid Tumors	ECOG-ACRIN	In Review 2/22/2022	Waiting for revision 2 since 9/28; reminder was sent to EA 10/24	(P) AMG 510 (Sotorasib) (825510) (O) Panitumumab (742319)
11	EAY191-N5	Molecular Analysis for Combination Therapy Choice (ComboMATCH) EAY191-N5: A Randomized Trial of Neratinib, A Pan-ERBB Inhibitor, Alone or in Combination with Palbociclib, a CDK4/6 Inhibitor, in Patients with HER2+ Gynecologic Cancers and Other Solid Tumors A ComboMATCH Treatment Trial	NRG	In Review 11/22/2021	Rev 3 disapproval sent 10/15; waiting for Rev 4	(P) Neratinib (783782) (O) Palbociclib (PD-0332991) (772256)

MyeloMATCH Priority 1 List

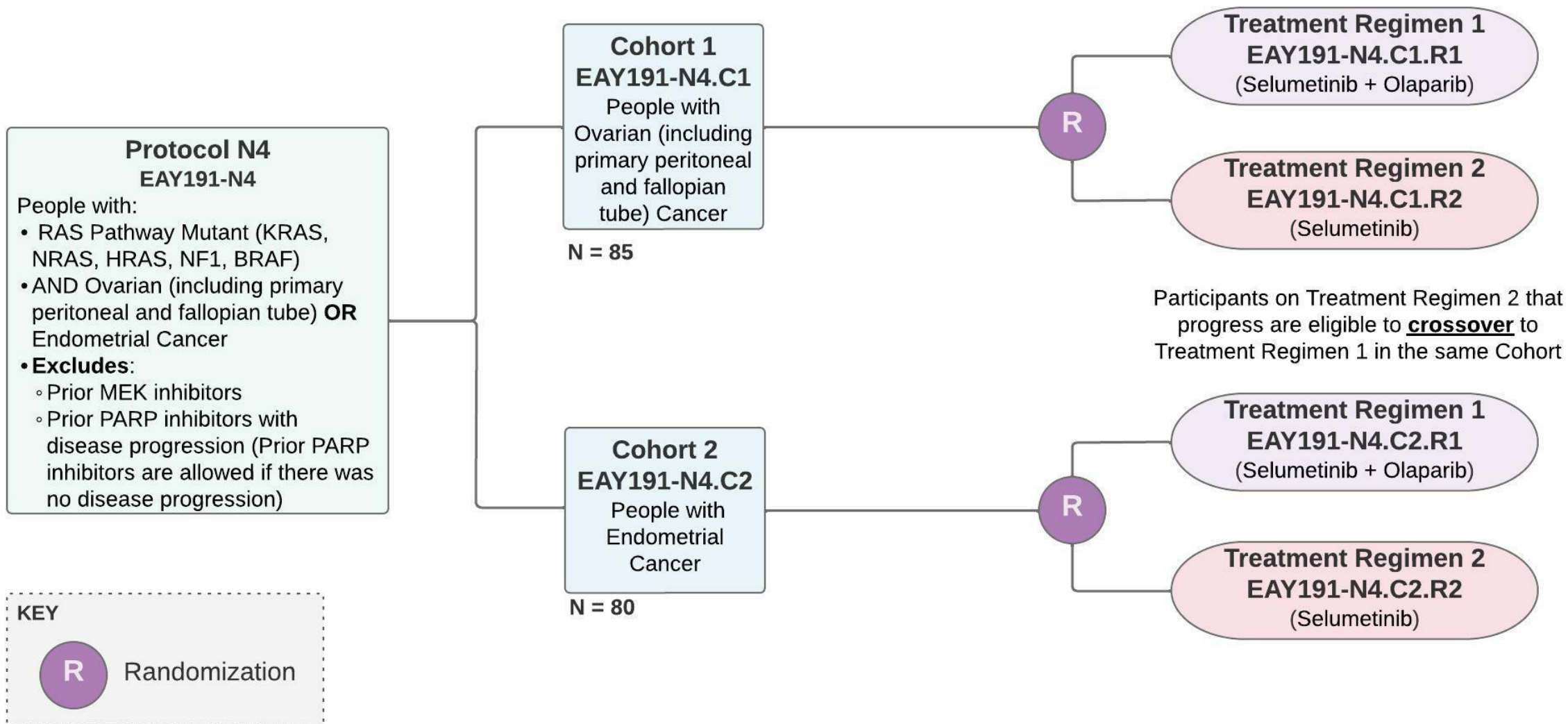
Study #	Document Number	Document Title	Group	Current Status	Next Steps
1	MYELOMATCH	Master Screening and Reassessment Protocol (MSRP) for the NCI MyeloMATCH Clinical Trials	SWOG	11/12/2021: Approval on Hold	CIRB Reviewed on 9/15, version with stips under CTEP review now.
2	MM1YA-S01	A Randomized Phase II Study Comparing Cytarabine + Daunorubicin (7 + 3) to (Daunorubicin and Cytarabine) Liposome, Cytarabine + Daunorubicin + Venetoclax, and Azacitidine + Venetoclax in Patients Aged 59 or Younger with High-Risk (Adverse) Acute Myeloid Leukemia; A MYELOMATCH Clinical Trial	SWOG	9/2/2022: Approval on Hold	CIRB Reviewed on 9/15, version with stips under CTEP review now.
3	MM1YA-CTG01	A Measurable Residual Disease (MRD) Driven, Phase II Study of Venetoclax Plus Chemotherapy for Newly Diagnosed Younger Patients with Intermediate Risk Acute Myeloid Leukemia: A Tier 1 MYELOMATCH Clinical Trial	CCTG	9/2/2022: Approval on Hold	CIRB Review Oct 6, responding to stips
6	MM10A-EA02 (Concept)	A Randomized Phase II Study of HMA-Based Therapies for the Treatment of Older Adults with Newly Diagnosed FLT3-Mutated Acute Myeloid Leukemia: A myeloMATCH Treatment Trial	ECOG-ACRIN	8/29/2022: Approved, no protocol yet	Waiting for protocol

MyeloMATCH Priority 2 List

Study #	Document Number	Document Title	Group	Current Status	Next Steps
4	MM2YA-EA01	Eradicating Measurable Residual Disease in Patients with Acute Myeloid Leukemia (AML) Prior to StEm Cell Transplantation (ERASE): A MyeloMATCH Treatment Trial	ECOG-ACRIN	1/21/2022: In Review	Reviewed by CIRB, responding to stips
5	MM1OA-S02	A Randomized Phase II Study of Targeted Agents with the Oral DNMT1 Inhibitor Cedazuridine-Decitabine (ASTX727) for the Treatment of Newly Diagnosed TP53-Mutated Acute Myeloid Leukemia in Older Patients: A myeloMATCH Treatment Trial	SWOG	10/17/2022: In Review	Scheduled for PRC 11/4
7	MM1MDS-EA03 (Concept)	ERADICATE MDS with TP53 Study: Efficacy of targeted Agents Combinations with Oral DNMT1 Inhibitor C-DEC (ASTX727) for Treatment of Higher Risk myelodysplastic Syndromes with TP53 Mutations: A Non-Comparative, Parallel, Multi-Arm Phase 2 Study	ECOG-ACRIN	8/12/2022: Approval on hold, waiting for drug commitment	W+A6:G8aiting for Astex commitment, Genentech has provided commitment on 10/20

Review Schemas

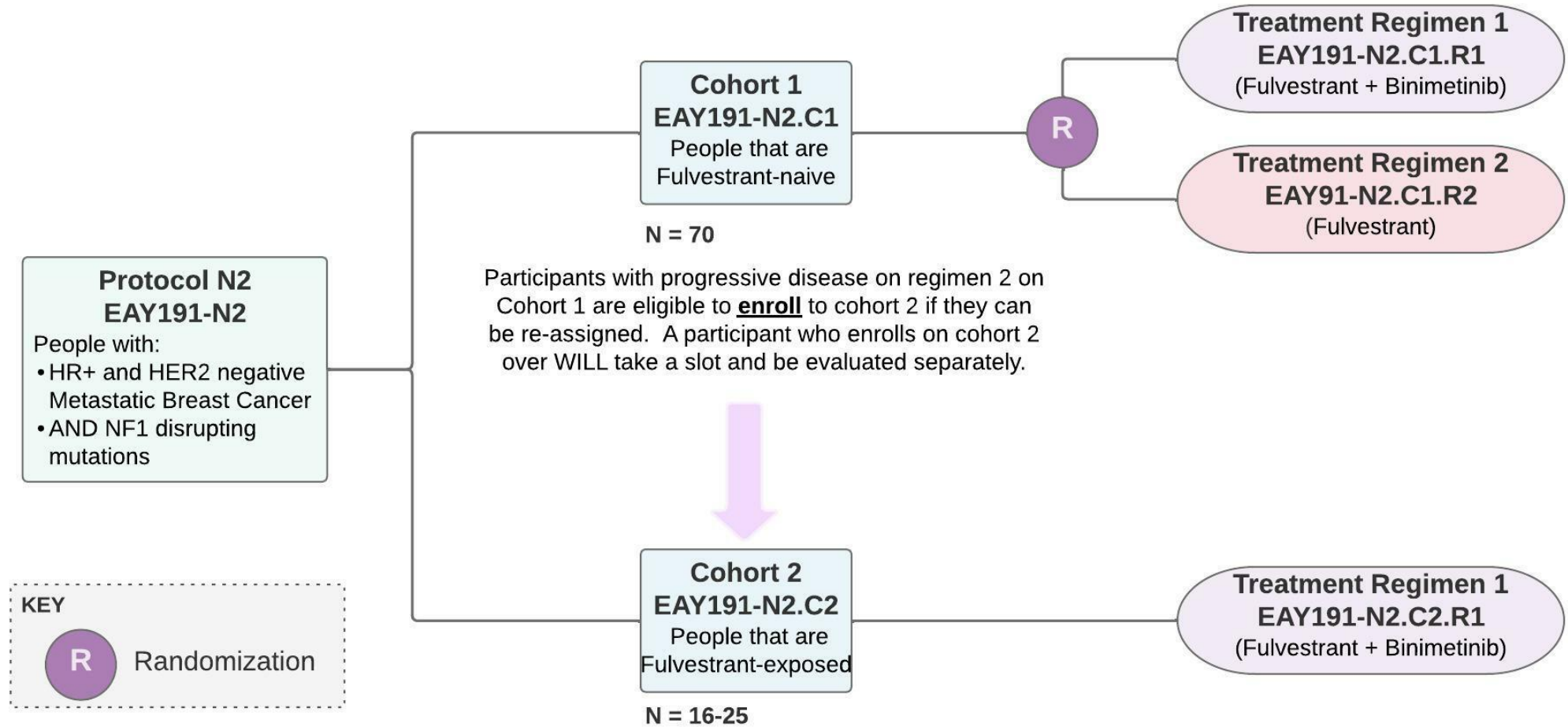
EAY191-N4



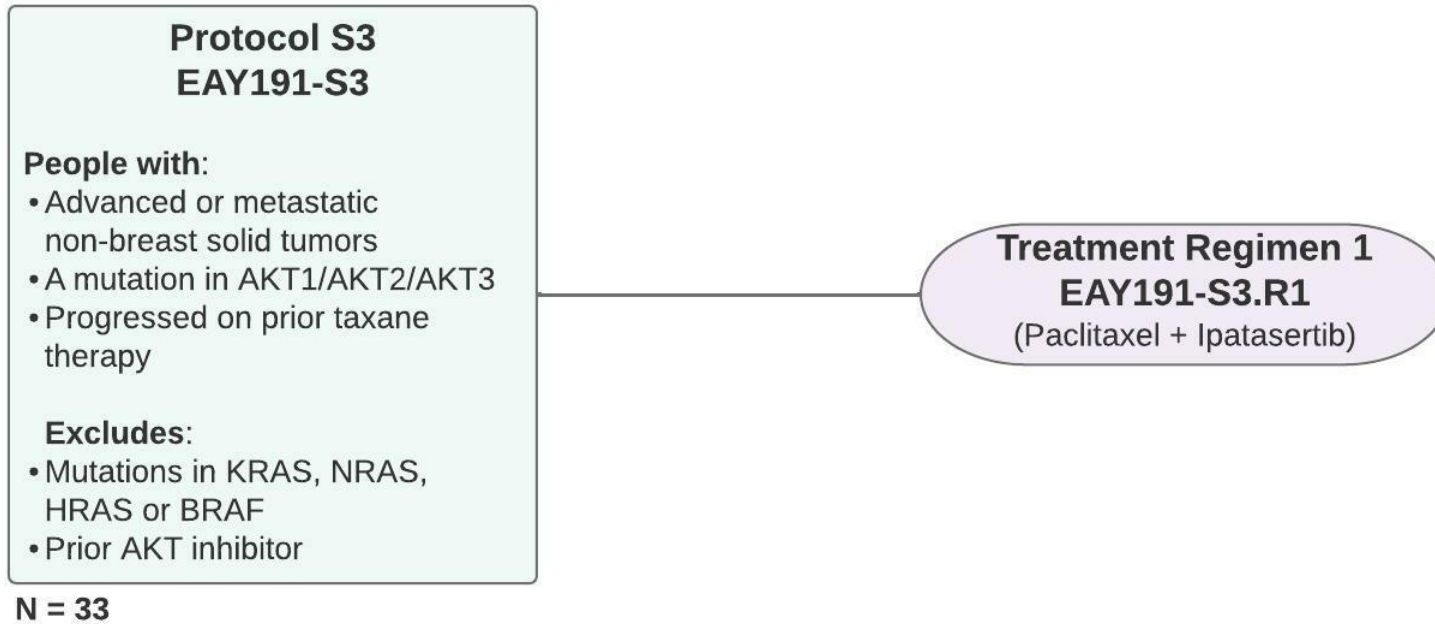
EAY191-E4



EAY191-N2- Draft



EAY191-S3



MMIYA-CTG01 Draft

