

**UNITED STATES  
SECURITIES AND EXCHANGE COMMISSION**  
Washington, D.C. 20549

**FORM 8-K**

**CURRENT REPORT**  
Pursuant to Section 13 or 15(d)  
of the Securities Exchange Act of 1934

Date of Report (Date of earliest event reported): January 13, 2020 (January 12, 2020)

**G1 THERAPEUTICS, INC.**

(Exact name of registrant as specified in its charter)

**Delaware**  
(State or other jurisdiction  
of incorporation)

**001-38096**  
(Commission  
File Number)

**26-3648180**  
(IRS Employer  
Identification No.)

**700 Park Offices Drive**  
**Suite 200**  
**Research Triangle Park, NC**  
(Address of principal executive offices)

**27709**  
(zip code)

Registrant's telephone number, including area code: (919) 213-9835

Check the appropriate box below if the Form 8-K filing is intended to simultaneously satisfy the filing obligation of the registrant under any of the following provisions:

- Written communications pursuant to Rule 425 under the Securities Act (17 CFR 230.425)
- Soliciting material pursuant to Rule 14a-12 under the Exchange Act (17 CFR 240.14a-12)
- Pre-commencement communications pursuant to Rule 14d-2(b) under the Exchange Act (17 CFR 240.14d-2(b))
- Pre-commencement communications pursuant to Rule 13e-4(c) under the Exchange Act (17 CFR 240.13e-4(c))

Securities registered pursuant to Section 12(b) of the Act:

| Title of each class              | Trading<br>Symbol | Name of each exchange<br>on which registered |
|----------------------------------|-------------------|--|
| Common stock, \$0.0001 par value | GTHX              | The Nasdaq Stock Market                      |

Indicate by check mark whether the registrant is an emerging growth company as defined in Rule 405 of the Securities Act of 1933 (17 CFR §230.405) or Rule 12b-2 of the Securities Exchange Act of 1934 (17 CFR §240.12b-2).

Emerging Growth Company

If an emerging growth company, indicate by check mark if the registrant has elected not to use the extended transition period for complying with any new or revised financial accounting standards provided pursuant to Section 13(a) of the Exchange Act.

**Item 7.01 Regulation FD Disclosure**

Beginning on January 12, 2020, representatives of G1 Therapeutics, Inc. (the “Company”) will make presentations to certain investors and analysts. The Company’s presentation is attached to this Current Report on Form 8-K as Exhibit 99.1 (the “Investor Presentation”) and is incorporated herein by reference. It is also located on the Company’s website at [www.g1therapeutics.com](http://www.g1therapeutics.com) under “Investors.”

Pursuant to General Instruction B.2 of Current Report on Form 8-K, the information contained in, or incorporated into, Item 7.01, including the Investor Presentation attached hereto as Exhibit 99.1, is being furnished and shall not be deemed “filed” for the purposes of Section 18 of the Securities Exchange Act of 1934, as amended (the “Exchange Act”), or otherwise subject to the liabilities of that section, nor shall it be deemed incorporated by reference into any registration statement or other filing under the Securities Act of 1933, as amended, or the Exchange Act, except as shall be expressly set forth by specific reference to such filing.

**Item 9.01 Financial Statements and Exhibits.**

(d) Exhibits

| <b>Exhibit No.</b> | <b>Description</b>  |
|--------------------|---|
| 99.1               | <a href="#">Investor Presentation dated January 14, 2020</a>              |
| 104                | Cover Page Interactive Data File (embedded with the Inline XBRL document) |

**SIGNATURES**

Pursuant to the requirements of the Securities Exchange Act of 1934, the Registrant has duly caused this report to be signed on its behalf by the undersigned thereunto duly authorized.

**G1 THERAPEUTICS, INC.**

By: /s/ James Stillman Hanson  
James Stillman Hanson  
General Counsel

Date: January 13, 2020



# Next Generation Cancer Therapies

*38th Annual J.P. Morgan Healthcare Conference*

January 14, 2020

## Forward-looking statements

This presentation contains forward-looking statements within the meaning of the Private Securities Litigation Reform Act of 1995. Words such as "may," "will," "expect," "plan," "anticipate," "estimate," "intend" and similar expressions (as well as other words or expressions referencing future events, conditions or circumstances) are intended to identify forward-looking statements. Forward-looking statements in this presentation include, but are not limited to, the therapeutic potential of trilaciclib, rintodestrant (G1T48), and lerociclib, the expected timing of data availability from ongoing clinical trials, the expected timing of initiation of future clinical trials, the expected timing for the completion of marketing applications in the U.S. and Europe for trilaciclib in SCLC, and the ability to add additional indications for trilaciclib, and are based on the Company's expectations and assumptions as of the date of this presentation. Each of these forward-looking statements involves risks and uncertainties. Factors that may cause the Company's actual results to differ from those expressed or implied in the forward-looking statements in this presentation are discussed in the Company's filings with the U.S. Securities and Exchange Commission, including the "Risk Factors" sections contained therein and include, but are not limited to, the Company's ability to complete clinical trials for, obtain approvals for and commercialize any of its product candidates; the Company's initial success in ongoing clinical trials may not be indicative of results obtained when these trials are completed or in later stage trials; the inherent uncertainties associated with developing new products or technologies and operating as a development-stage company; the Company's development of a CDK4/6 inhibitor to reduce chemotherapy-induced myelosuppression is novel, unproven and rapidly evolving and may never lead to a marketable product; and market conditions, including competition. Except as required by law, the Company assumes no obligation to update any forward-looking statements contained herein to reflect any change in expectations, even as new information becomes available.



## Next-generation cancer therapies

### Trilaciclib

First-in-class  
myelopreservation  
therapy

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### Rintodestrant

(G1T48)

Potential best-in-class  
oral SERD

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### Lerociclib

Differentiated  
oral CDK4/6  
inhibitor

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**Committed to improving lives  
and outcomes of people living  
with cancer**



# Current chemotherapy landscape

**~1M**  
U.S. patients  
**receive**  
chemotherapy  
**annually**



Chemotherapy  
remains the cornerstone  
**of treatment**  
for most cancers



## Myelosuppression

is currently an unavoidable consequence of chemo that impacts patient safety, QoL and costs to the HC system

|  |  |   |
|--|--|---|
| Neutropenia and anemia                         | Risk of infection: G-CSF use, associated bone pain | RBC transfusions and ESA rescue         |
| Impaired anti-tumor immunity                   |  | Fatigue                                 |
| Hospitalizations and unscheduled office visits | Chemotherapy dose delays and reductions            | Risk of bleeding: platelet transfusions |



# Patient experience of myelosuppression: burdensome and far-reaching

# 89%

**OF CANCER PATIENTS**  
with myelosuppression rate it as having  
a moderate to major impact on their life\*

*"...the overall fatigue was the worst. It stole my energy and joy for both life and family. It made me want to quit chemo numerous times."*

*"I don't feel like doing ANYTHING some days. It's like depression but completely physical. Of course, everyone's trying to be supportive. And I have my own obligations, but I feel like a burden."*

*"...it so happened I had a father dying in the hospital and I was strictly forbidden from entering a hospital (except my own)."*



\*Sterling IRB-reviewed online survey in 4Q19 of 301 patients treated with chemotherapy within past 12 months who experienced myelosuppression; respondents: 51% breast cancer; 33% lung cancer; 16% colorectal cancer; manuscript in preparation

**What if we can improve the chemo experience for people living with cancer?**



Our solution:  
**Trilaciclib**

**First-in-class**  
myelopreservation therapy  
that has the potential to make  
chemotherapy safer, improve  
the patient experience, and in  
some settings, help patients  
live longer

## Our solution: trilaciclib



30-minute IV infusion prior to chemo; **given first time and every time** chemo is administered



**Preserves** bone marrow and immune system function from damage by chemo



**Protects** patients from the dangerous side effects of myelosuppression



In some settings, may help **patients live longer**



Can be **incorporated into multiple chemo regimens**, including I/O + chemo

**FDA  
Breakthrough  
Therapy  
Designation for  
SCLC**



\*As observed in clinical trials to date

## Body of evidence in SCLC: three randomized trials

**Trilaciclib reduces chemotherapy-related toxicity and need for rescue interventions**

- ✓ Significant improvement in patient experience, notably **less fatigue**
- ✓ **Less neutropenia and anemia**
- ✓ **Reduced G-CSF usage and transfusions**

Three randomized, placebo-controlled, double-blind trials in:  
1<sup>st</sup>-line SCLC (+ etop/carbo), 1<sup>st</sup>-line SCLC (+ etop/carbo/Tecentriq), 2<sup>nd</sup>/3<sup>rd</sup>-line SCLC (+ topotecan)

# Improved treatment experience in SCLC

## Patient survey findings\* (patients not enrolled in trilaciclib trials)

- 88% of SCLC respondents reported that myelosuppression **had moderate to major impact on their life**
- Of those, 63% noted fatigue as their biggest myelosuppressive issue
- Of those who noted fatigue, average rating of 8.4 on 10-point scale of how bothersome fatigue was

\*Sterling IRB-reviewed online survey in 4Q19



## Patient Reported Outcomes data (n=235) (pooled from three randomized, placebo-controlled, double-blind trials)

| Subscale   | <<Trilaciclib Better   Placebo Better>> | Hazard Ratio (95% CI) |
|--|---|-----------------------|
| Fatigue  |   | 0.56 (0.37, 0.85)     |
| Functional Well-being                              |   | 0.44 (0.28, 0.70)     |
| Physical Well-being                                |   | 0.62 (0.39, 0.97)     |
| Anemia – Trial Outcome Index                       |   | 0.53 (0.34, 0.83)     |
| Functional Assessment of Cancer Treatment - Anemia |   | 0.46 (0.29, 0.72)     |

Updated from data presented at MASCC 2019

# Significant multi-lineage myelopreservation benefits support improved patient experience

|                        |   | PLACEBO +<br>CHEMO | TRILA +<br>CHEMO |                   |
|------------------------|---|--------------------|------------------|-------------------|
|                        | <b>Patients (intent-to-treat population)</b>                              | <b>119</b>         | <b>123</b>       | <b>P-VALUE*</b>   |
| <b>Neutrophils</b>     | Mean duration (days) of severe neutropenia in cycle 1 (SD)                | 4 (5.1)            | 0 (1.8)          | <b>&lt;0.0001</b> |
|                        | Occurrence of severe neutropenia  | 63 (52.9%)         | 14 (11.4%)       | <b>&lt;0.0001</b> |
|                        | Occurrence of G-CSF administration  | 67 (56.3%)         | 35 (28.5%)       | <b>&lt;0.0001</b> |
|                        | Incidence of G-CSF administration (event rate per 100 cycles)             | 40.6               | 16.4             | <b>&lt;0.0001</b> |
| <b>Red Blood Cells</b> | Occurrence of Grade 3/4 anemia  | 38 (31.9%)         | 25 (20.3%)       | <b>0.0279</b>     |
|                        | Occurrence of ESA administration  | 14 (11.8)          | 4 (3.3%)         | <b>0.0254</b>     |
|                        | Occurrence of RBC transfusions on/after 5 weeks                           | 31 (26.1%)         | 18 (14.6%)       | <b>0.0252</b>     |
|                        | Incidence of RBC transfusions on/after 5 weeks (event rate per 100 weeks) | 3.1                | 1.5              | <b>0.0027</b>     |
| <b>Platelets</b>       | Occurrence of Grade 3/4 thrombocytopenia                                  | 43 (36.1%)         | 24 (19.5%)       | <b>0.0067</b>     |



\*2-sided p-value; pooled data from three randomized, placebo-controlled, double-blind SCLC trials, updated from data presented at MASCC 2019

# Preparing to bring trilaciclib to market in SCLC

## U.S.

**Rolling NDA submission initiated 4Q19;  
expect to complete submission 2Q20**

Building strong, functional capabilities in U.S.



## Ex-U.S.

Planning to submit MAA in 2H20

**Evaluating partnership opportunities to commercialize trilaciclib ex-U.S.**



## Pursuing additional indications: breast cancer

### Preliminary survival benefit observed in mTNBC randomized trial



- ✓ Patients able to tolerate more cycles of chemo, without increased toxicity
- ✓ Reduced rate of RBC transfusions
- ✓ Patient-reported outcomes data support improved patient experience
- ✓ Significant improvement in OS

# Preliminary overall survival benefit in mTNBC

|                           | Control (GC only)<br>(Group 1) | Trilaciclib + GC<br>(Group 2) | Trilaciclib + GC<br>(Group 3) | Trilaciclib + GC<br>(Group 2+3) |
|---------------------------|--------------------------------|-------------------------------|-------------------------------|---------------------------------|
| ITT population            | N = 34                         | N = 33                        | N = 35                        | N = 68                          |
| <b>Median OS (months)</b> | <b>12.6</b>                    | <b>20.1</b>                   | <b>17.8</b>                   | <b>20.1</b>                     |
| <b>HR</b>                 |                                | <b>0.33</b>                   | <b>0.34</b>                   | <b>0.36</b>                     |
| <b>p-value</b>            |                                | <b>0.028</b>                  | <b>0.0023</b>                 | <b>0.0015</b>                   |
| Median PFS (months)       | 5.7                            | 9.4                           | 7.3                           | 8.8                             |
| HR                        |                                | 0.60                          | 0.59                          | 0.59                            |
| p-value                   |                                | 0.13                          | 0.12                          | 0.063                           |

Hypothesis: trilaciclib's preservation of immune system function during chemo drives OS benefit



Group 1: GC only (Day 1/8) (n=34); Group 2: trilaciclib + GC (Day 1/8) (n=33); Group 3: trilaciclib only (Day 1/8), trilaciclib + GC (Day 2/9) (n=35)  
 ESMO 2019: Trilaciclib improves overall survival when given with gemcitabine/carboplatin (GC) in patients with metastatic triple negative breast cancer (mTNBC) in a randomized Phase 2 trial: Abstract #6255.  
 Results published concurrently in *The Lancet Oncology*

# Next step in breast cancer: I-SPY 2 neoadjuvant trial

## Goals of Phase 2 trial

- Evaluate trilaciclib in broadly-used chemo regimens (e.g. Taxol + AC)
- Evaluate trilaciclib in all breast cancer sub-types (ER+, HER2+, TNBC)
- Evaluate impact of trilaciclib on tumor immune microenvironment +/- PD-1
- Endpoints: biomarkers, efficacy and myelopreservation

Neoadjuvant breast cancer with high risk of recurrence → could be any HR or HER2 status (10 biomarker subtypes)

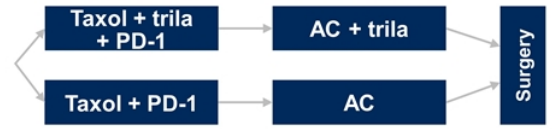


## Four-arm Bayesian design

### Chemotherapy + trilaciclib



### Chemotherapy + PD-1 + trilaciclib



AC = adriamycin and cyclophosphamide

Opportunity to improve patient outcomes  
across multiple indications

**~1 million\* patients in planned indications**

**68,000**

Small Cell  
Lung Cancer

**>350,000**

Adjuvant Breast  
Cancer

**>500,000**

Colorectal Cancer

**20,000**

Metastatic  
Triple-Negative  
Breast Cancer



\*Includes U.S., EU5 and Japan; Decision Resources Group estimate for chemo-treated patients in 2027; see Slide 31 for detail

## Next steps for trilaciclib across multiple indications

### SCLC

- Complete NDA submission in 2Q20
- PDUFA date assigned in 3Q20 (pending acceptance)
- MAA submission in 2H20

### Breast Cancer

- Initiate I-SPY2 trial in 2Q20
- Updated OS data from mTNBC trial in 2H20

### Colorectal Cancer

- Initiate Phase 3 trial in 4Q20

## Next-generation cancer therapies

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### Rintodestrant

(G1T48)

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### Lerociclib

Differentiated  
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## Improving options for ER+, HER2- breast cancer

- ER degradation shown to be most effective means of blocking estrogen signaling in ER+, HER2- breast cancer
- Only available SERD is fulvestrant – painful intramuscular injections

**Opportunity to improve options in first-line and adjuvant settings with oral SERD**

# Rintodestrant Phase 1 data: well tolerated with proof-of-concept anti-tumor activity

**Well tolerated;** favorable safety profile observed at all dose levels

**No dose-limiting toxicities observed;** maximum tolerated dose not reached

**AEs mostly Grade 1,** no bradycardia or cytopenias

<sup>18</sup>F-FES PET scans:  
**ER occupancy  $\geq$  80%** in doses  $\geq$  600 mg

Preliminary evidence of **anti-tumor activity** in heavily pre-treated population



ESMO 2019: Dose-escalation study of G1T48, an oral selective estrogen receptor degrader (SERD), in postmenopausal women with ER+/HER2- locally advanced or metastatic breast cancer (ABC); Abstract #3587



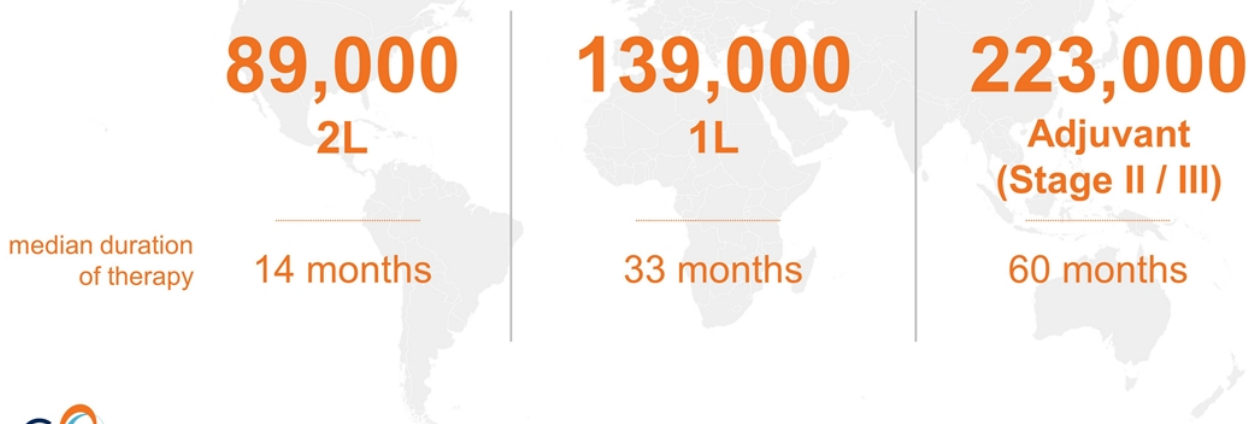
## Assessing the potential of rintodestrant

**Phase 1/2a program  
w/ ~100 patients  
enrolled by YE20:  
additional data 2H20**

- ~65 patients enrolled in Phase 1/2a trial to **identify dose** (expansion cohorts of 600 mg and 1,000 mg monotherapy)
- Collecting PD data from biopsies to **confirm activity in the tumor**
- **Additional arm** to include another ~40 patients evaluating **rintodestrant + palbociclib combination**

# Significant potential to improve ER+ breast cancer treatment

>450,000 2L, 1L, and adjuvant ER+ BC patients globally



\*Source: secondary epi sources, Decision Resources Group 2027 estimates for U.S., EU5 and Japan  
\*\*Duration estimates based on similar trial results in the same or similar patient populations as planned trials

## Next-generation cancer therapies

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oral SERD

---

### Lerociclib

Differentiated  
oral CDK4/6  
inhibitor

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# Lerociclib: differentiated oral CDK4/6 inhibitor

|            | DOSE-LIMITING NEUTROPENIA | MONITORING REQUIREMENT        | DOSING HOLIDAY | QT PROLONGATION | DILI | GRADE 3/4 DIARRHEA | VTE |
|------------|---------------------------|-------------------------------|----------------|-----------------|------|--------------------|-----|
| Ibrance®   | X                         | X                             | X              | -               | -    | -                  | -   |
| Kisqali®   | X                         | X                             | X              | X               | X    | -                  | -   |
| Verzenio®  | X                         | X                             | -              | -               | X    | X                  | X   |
| lerociclib | -                         | Potential for less monitoring | -              | -               | -    | -                  | -   |

Differentiated PK and tolerability profile

Continuous dosing (no holiday) with fewer dose-limiting toxicities

Potential for less CBC monitoring, reducing patient & physician burden

## Clinical overview: improved safety and tolerability profile

**Phase 1b/2 trial:** 110 patients lero + fulvestrant (similar entry criteria to PALOMA 3)

Low rates of Grade 4 neutropenia **without a drug holiday**

**Low rates of stomatitis and alopecia** across all dose levels

**65.2% clinical benefit rate;** median progression-free survival of 15 months (immature)

**Updated data 2H20** on 150mg and 200mg BID cohorts, enabling Phase 3 dose selection



SABCS 2019: Dose escalation and expansion study of lerociclib (G1T38), an oral CDK4/6 inhibitor, dosed with no drug holiday in combination with fulvestrant in patients with HR+/HER2- advanced breast cancer; Abstract #P1-19-17

## Significant progress in 2019, setting up catalysts in 2020

### Trilaciclib

- ✓ Breakthrough Therapy designation for SCLC
- ✓ Began rolling NDA submission for SCLC
- ✓ Preliminary OS improvement in TNBC

### Rintodestrant (G1T48)

- ✓ Phase 1 data demonstrated POC and potential best-in-class profile
- ✓ Completed Phase 2a monotherapy enrollment

### Lerociclib

- ✓ Demonstrated POC and differentiation from existing CDK4/6i
- ✓ Completed enrollment of BC and lung trials

## 2020: near-term clinical and regulatory milestones

| Therapy               | Indication                    | 1H20  | 2H20                              |
|-----------------------|-------------------------------|---|-----------------------------------|
| Trilaciclib           | Small cell lung cancer        | Complete NDA submission                               | PDUFA date assigned; MAA filing   |
|                       | Breast cancer                 | Initiate I-SPY 2 trial                                | OS update from Phase 2 TNBC trial |
|                       | Metastatic colorectal cancer  | FDA pre-Phase 3 meeting                               | Initiate Phase 3 trial            |
| Rintodestrant (G1T48) | ER+, HER2- BC                 | Initiate Phase 2 expansion w/ palbociclib combination | Data update                       |
| Lerociclib            | ER+, HER2- BC (+ fulvestrant) |   | Data update                       |

# G1 Therapeutics: improving outcomes in cancer treatment



**Trilaciclib: near-term commercial opportunity to improve outcomes for patients receiving chemo**



**Rintodestrant: potential best-in-class oral SERD**



**Global rights to all compounds provides multiple options for value-creating partnerships**



**Well funded with cash runway into 2H21**



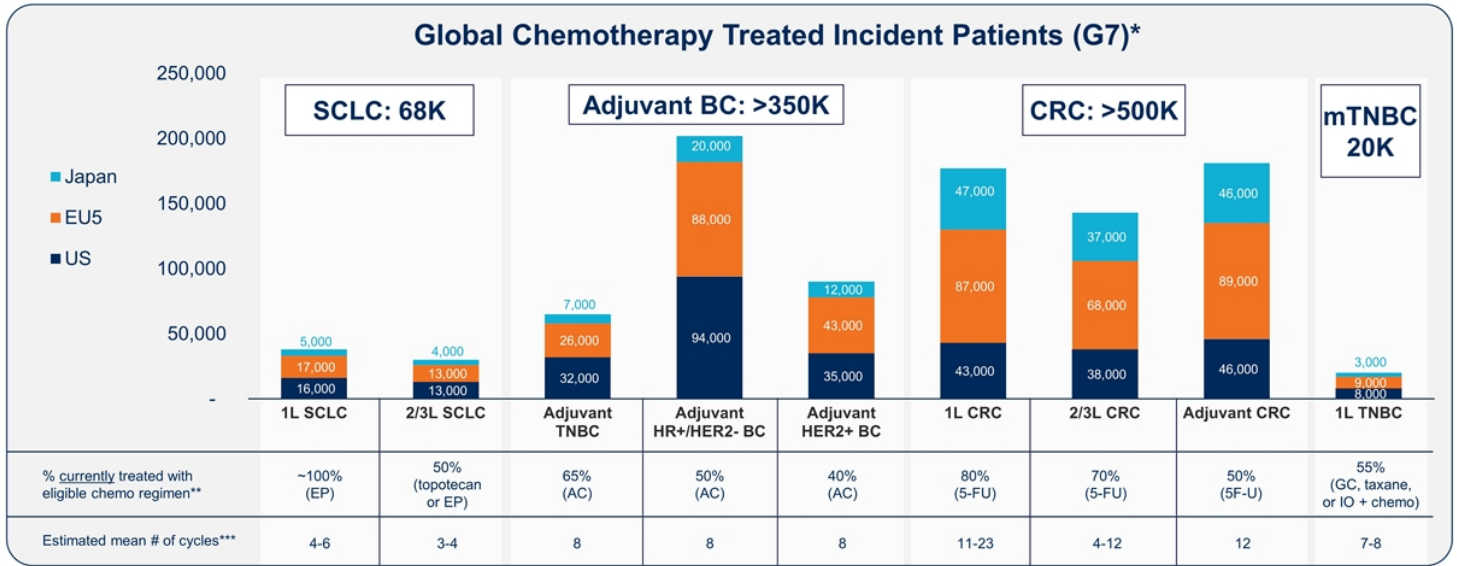


# Appendix



# Potential to improve outcomes across tumor types

Trilaciclib used first time, and every time, patient receives chemotherapy



\*Source: Secondary epi sources, 2027 estimates

\*\*EP refers to any regimen that includes etoposide + platinum; GC refers to gemcitabine/carboplatin; AC refers to any regimen that includes Adriamycin and cyclophosphamide; 5-FU refers to any regimen that includes fluorouracil (e.g., FOLFOLX). In addition to CRC, pancreatic cancer, gastroesophageal cancer and squamous cell carcinoma of the head and neck (SCCHN) are also treated with 5-FU regimens (% currently treated with 5-FU regimens varies by tumor type and region)

\*\*\*Source: SCLC and TNBC: G1 Therapeutics' completed trials; CRC and Adjuvant BC: number of cycles for eligible chemo regimens from Decision Resources Group Treatment Landscape and Forecast Assumptions 2018 Reports (CRC and BC)