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4 UNITED STATES DISTRICT COURT
5 NORTHERN DISTRICT OF CALIFORNIA

6 AAT BIOQUEST, INC.,

7 Plaintiff,

8 v.

9 TEXAS FLUORESCENCE
10 LABORATORIES, INC.,

11 Defendant.

Case No. [14-cv-03909-DMR](#)

**ORDER GRANTING PLAINTIFF'S
MOTION FOR SUMMARY JUDGMENT
AND DENYING DEFENDANT'S
MOTION FOR SUMMARY JUDGMENT**

Dkt. No. 27, 35

12
13 Before the court are cross-motions for summary judgment pursuant to Federal Rule of
14 Civil Procedure 56 filed by Plaintiff AAT Bioquest Inc. ("AAT") and Defendant Texas
15 Fluorescence Laboratories ("TEFLabs").¹ The court conducted a hearing on March 3, 2015. After
16 full consideration of the parties' submissions and oral argument, for the reasons stated below, the
17 PMSJ is granted, and the DMSJ is denied.

18 **I. FACTS**

19 **A. The Complaint**

20 AAT brings this complaint against TEFLabs for infringement of AAT's United States
21 Patent No. 8,779,165 ("the '165 Patent"), entitled "Fluorescent Ion Indicators² and Their
22 Applications." Compl. [Docket No. 1] at ¶¶ 12-18. AAT sells a fluorescent calcium ion indicator
23 called Fluo-8 AM, which is an embodiment of Claim 1 of the '165 Patent.

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25 ¹ The parties' cross-motions were sequentially briefed: TEFLabs filed its motion first [Docket No.
26 27, "DMSJ"], then AAT filed its response and cross-motion [Docket No. 35, "PMSJ"]. TEFLabs
27 next filed a reply [Docket No. 36, "Reply"], followed by AAT's surreply [Docket No. 39,
"Surreply"].

28 ² Fluorescent ion indicators are also referred to as "fluo calcium ion indicators" or "fluo indicators."

1 TEFLabs concedes that it makes and sells a fluorescent calcium ion indicator with the
2 same structure as Fluo-8 AM, which it calls “Fluo-2 MA AM.” TEFLabs also admits that Fluo-2
3 MA AM infringes Claim 1 of the ’165 Patent. Answer [Docket No. 11] at ¶ 7. However, it asserts
4 various defenses based on the invalidity of the patent and AAT’s alleged inequitable conduct. The
5 viability of these defenses is the sole issue in the parties’ motions.

6 **B. Overview of Technology**

7 The basic facts regarding the technology at issue are not disputed. The ’165 Patent
8 involves calcium ion indicators, which are chemical compounds that detect the presence and
9 quantity of calcium ion in a cell. Calcium ion indicators are also referred to generically as “dyes,”
10 because they attach to and highlight the target calcium ion. Calcium ion indicators typically
11 contain (1) a binding component³ that attaches (or “chelates”) to calcium ion and (2) a reporter
12 component that illuminates when the binding component binds to calcium ion, making it easier to
13 observe the presence of calcium ion. They may also include Acetoxymethyl (AM) ester groups
14 that assist the indicator in entering the cell, a process called “cell loading.” For Fluo calcium ion
15 indicators, the reporter component is based upon fluorescein, a fluorescent molecule. Calcium ion
16 indicators based on rhodamine reporters are called rhod calcium ion indicators.

17 Fluo indicators are used to detect calcium ion through a procedure called an “intracellular
18 calcium assay.” The following chemical processes occur during an assay: the fluo indicator is
19 added to living cells and internalized, or “loaded,” into the cells. Once inside the cells, the
20 indicator dye is hydrolyzed (i.e., the AM ester is cleaved off the indicator), which activates the
21 indicator to be capable of binding to calcium ion. The cells are then exposed to a substance that
22 causes the release of calcium ions from within storage sites inside the cell. Once the calcium ions
23 are released, they bind to the indicator. During this binding event, the properties of the indicator
24 are changed and it becomes fluorescent. A light source is used to illuminate the calcium ions; the
25 source most commonly used in the process relevant to this case is an argon laser emitting 488
26 nanometer wavelength light. A person can then measure the intensity of the fluorescence emitted
27 from the cells, which indicates the amount of calcium ion in the cells.

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³ The binding component in the indicators at issue is called the “BAPTA ion chelator.”

1 An indicator is more desirable if it loads quickly into a cell, followed by rapid cleaving of
2 the AM esters, because slow cleavage can cause the indicator to leak out of the cell. Another
3 desirable characteristic is a strong fluorescence signal; a weak or “quenched” signal may make it
4 more difficult to detect calcium ion. An indicator that can be loaded at a variety of temperatures is
5 more desirable than an indicator that may only be loaded at a specific temperature. Another
6 characteristic of an indicator is the “binding affinity” between the binding component of the
7 indicator and the targeted calcium ion.⁴ High-affinity binding results from greater molecular force
8 between the targeted molecule and the binding component, while low-affinity binding involves
9 less intermolecular force between the two.

10 **C. Relevant Prior Art and Patent History**

11 **1. Prior Art: Tsien Patent**

12 In 1991, a patent in the field of fluorescent calcium ion indicators was awarded to Roger
13 Tsien and Akwasi Minta (United States Patent No. 5,049,673, or the “Tsien Patent”). Dr. Minta
14 later founded TEFLabs in 1992.

15 **2. Patent-in-Suit: ’165 Patent**

16 AAT was founded in 2006 by Dr. Zhenjun Diwu. On April 13, 2007, Dr. Diwu and
17 several co-inventors filed United States Provisional Patent Application No. 60/923,452 (“the ’452
18 Application” or the “provisional patent application”). The ’452 Application was “directed to a
19 family of fluorescent dyes that are useful for preparing fluorescent metal ion indicators.”

20 On February 29, 2008, Dr. Diwu filed United States Patent Application No. 12/040,753
21 (“the ’753 Application”), the non-provisional continuation of the ’452 Application.

22 On March 11, 2011, Dr. Diwu filed a related patent application as United States Patent
23 Application No. 12/932,683 (“the ’683 Application”), which was a continuation of the then-
24 pending ’753 Application.⁵ The Applicants disclosed the Tsien Patent as prior art in an
25 Information Disclosure Statement dated June 20, 2011, during prosecution of the ’683
26 Application. Claim 33 of the ’683 Application, added by amendment on August 28, 2013,

27 ⁴ The targeted molecule is known generically as a “ligand.”

28 ⁵ The ’753 Application was eventually deemed abandoned on December 15, 2011.

1 discloses the subject matter of what eventually became Claim 1 of the '165 Patent.

2 The '683 Application was granted and issued as the '165 Patent on July 15, 2014.

3 **II. LEGAL STANDARD**

4 **A. General Summary Judgment Standard**

5 A court shall grant summary judgment “if . . . there is no genuine dispute as to any material
6 fact and the movant is entitled to judgment as a matter of law.” Fed. R. Civ. P. 56(a). The burden
7 of establishing the absence of a genuine issue of material fact lies with the moving party. *Celotex*
8 *Corp. v. Catrett*, 477 U.S. 317, 322-23 (1986), and the court must view the evidence in the light
9 most favorable to the non-movant. *See Anderson v. Liberty Lobby, Inc.*, 477 U.S. 242, 255 (1986)
10 (citation omitted). A genuine factual issue exists if, taking into account the burdens of production
11 and proof that would be required at trial, sufficient evidence favors the non-movant such that a
12 reasonable jury could return a verdict in that party’s favor. *Id.* at 248. The court may not weigh
13 the evidence, assess the credibility of witnesses, or resolve issues of fact. *See id.* at 249; *SCA*
14 *Hygiene Products Aktiebolag v. First Quality Baby Products, LLC*, 767 F.3d 1339, 1347 (Fed.
15 Cir. 2014) *vacated on other grounds*, 2014 WL 7460970 (Fed. Cir. Dec. 30, 2014) (“the district
16 court [is] not permitted to assess the credibility of . . . witnesses on summary judgment”).

17 To defeat summary judgment once the moving party has met its burden, the nonmoving
18 party may not simply rely on the pleadings, but must produce significant probative evidence, by
19 affidavit or as otherwise provided by Federal Rule of Civil Procedure 56, supporting the claim that
20 a genuine issue of material fact exists. *TW Elec. Serv., Inc. v. Pac. Elec. Contractors Ass’n*, 809
21 F.2d 626, 630 (9th Cir. 1987); *SCA Hygiene*, 767 F.3d at 1347 (party “may not rely solely on
22 pleadings and speculation to create a genuine issue of material fact; it must identify particular
23 evidence that creates such a dispute”). In other words, there must exist more than “a scintilla of
24 evidence” to support the non-moving party’s claims; conclusory assertions will not suffice.
25 *Anderson*, 477 U.S. at 252. Similarly, “[w]hen opposing parties tell two different stories, one of
26 which is blatantly contradicted by the record, so that no reasonable jury could believe it, a court
27 should not adopt that version of the facts” when ruling on the motion. *Scott v. Harris*, 550 U.S.
28 372, 380 (2007).

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Where, as here, the parties have filed cross-motions for summary judgment, “[e]ach motion must be considered on its own merits In fulfilling its duty to review each cross-motion separately, the court must review the evidence submitted in support of each cross-motion.” *Fair Hous. Council of Riverside Cnty., Inc. v. Riverside Two*, 249 F.3d 1132, 1136 (9th Cir. 2001). *See also Conceptus, Inc. v. Hologic, Inc.*, 771 F. Supp. 2d 1164, 1174 (N.D. Cal. 2010) (court considering simultaneous cross-motions for summary judgment in patent infringement action must consider evidentiary material identified and submitted in support of both motions before ruling on each of them) (citing *Fair Hous. Council*, 249 F.3d at 1134).

B. Standard for Summary Judgment of Invalidity

A patent is presumed valid. 35 U.S.C. § 282. This presumption can be rebutted, but the party challenging validity must meet the “high burden” of proving invalidity by “clear and convincing evidence.” *Sciele Pharma Inc. v. Lupin Ltd.*, 684 F.3d 1253, 1260 (Fed. Cir. 2012). *See also U.S. v. Telectronics, Inc.*, 857 F.2d 778, 785 (Fed. Cir. 1988) (“The burden of proving invalidity . . . rests with the challenger . . . [and] must be proven by facts supported by clear and convincing evidence.”). The burden is especially high when the party challenging validity relies on the same evidence that was before the patent examiner. *Tokai Corp. v. Easton Enterprises, Inc.*, 632 F.3d 1358, 1367 (Fed. Cir. 2011) (“When no prior art other than that which was considered by the PTO examiner is relied on by the attacker, he has the added burden of overcoming the deference that is due to a qualified government agency presumed to have properly done its job.”) (citations omitted).

“[A] moving party seeking to invalidate a patent at summary judgment must submit such clear and convincing evidence of invalidity so that no reasonable jury could find otherwise.” *Eli Lilly v. Barr Labs., Inc.*, 251 F.3d 955, 962 (Fed. Cir. 2001) (citing *Anderson*, 477 U.S. at 248). “Alternatively, a moving party seeking to have a patent held not invalid must show that the nonmoving party, who bears the burden of proof at trial, failed to produce clear and convincing evidence on an essential element of a defense upon which a reasonable jury could invalidate the patent.” *Id.*

III. DISCUSSION

TEFLabs concedes its infringement of the '165 Patent and moves for summary judgment on the basis of four defenses:

- (A) the '165 Patent is invalid because it fails to meet the written description requirement;
- (B) the '165 Patent is invalid because it fails to meet the enablement requirement;
- (C) the '165 Patent is invalid because it is obvious or anticipated; and
- (D) the '165 Patent is unenforceable because of AAT's inequitable conduct.

The court analyzes each defense below.

A. Written Description Requirement

TEFLabs asserts the '165 Patent is invalid because it fails to meet the written description requirement under 35 U.S.C. § 112 ¶ 1.⁶

1. AAT's Motion to Strike

As a preliminary matter, AAT moves to strike the portions of the DMSJ related to the written description defense because TEFLabs did not assert it in either its Answer or its Invalidity Contentions, and instead raises it for the first time here.

“This district has adopted Patent Local Rules that ‘require parties to state early in the litigation and with specificity their contentions with respect to infringement and invalidity.’” *Monolithic Power Sys., Inc. v. O2 Micro Int'l Ltd.*, No. C08-04567-CW, 2009 WL 3353306, at *2 (N.D. Cal. Oct. 16, 2009) (quoting *O2 Micro Int'l, Ltd. v. Monolithic Power Sys., Inc.*, 467 F.3d 1355, 1359 (Fed. Cir. 2006)). Patent L.R. 3-3(d) requires that a party provide “[a]ny grounds of invalidity based on . . . enablement or written description under 35 U.S.C. § 112(1).” “The rules are designed to require parties to crystallize their theories of the case early in the litigation and to adhere to those theories once they have been disclosed.” *Mediatek Inc. v. Freescale Semiconductor, Inc.*, No. 11-cv-5341 YGR, 2014 WL 690161, at *1 (N.D. Cal. Feb. 21, 2014). “Any invalidity theories not disclosed pursuant to Local Rule 3-3 are barred, accordingly, from presentation at trial (whether through expert opinion testimony or otherwise).” *Id.*

⁶ The Leahy-Smith America Invents Act, enacted in 2011, amended several parts of the Patent Act. The successor statute to 35 U.S.C. § 112 ¶ 1, which applies to patents filed after September 16, 2012, is codified at 35 U.S.C. § 112(a). Because the patent at issue was filed before September 16, 2012, the older version of the statutes apply.

1 Neither party supplied TEFLabs’s Invalidation Contentions, but AAT submitted TEFLabs’s
 2 response to an interrogatory asking TEFLabs to “identify all legal and factual grounds on which
 3 you contend that [the Asserted Claim] is unenforceable.” *See* Carter Decl. at H. TEFLabs’s
 4 response does not include a written description defense. Instead, the response identifies invalidity
 5 defenses based on anticipation, obviousness, enablement, and best mode. *Id.* at 14-18. TEFLabs
 6 now contends that the “best mode argument” that it disclosed in the interrogatory response was
 7 actually a written description argument. *See* Reply at 4 (TEFLabs “corrected” its invalidity
 8 argument “from best mode, which is a statutory requirement without teeth since 2013, to written
 9 description, which is still a basis for invalidity”).

10 The disclosure of a “best mode” invalidity theory does not equate to the disclosure of a
 11 “written description” theory. *See Univ. Of Rochester v. G.D. Searle & Co.*, 358 F.3d 916, 921-22
 12 (Fed. Cir. 2004) (“Although there is often significant overlap between the three requirements
 13 [written description, best mode, and enablement], they are nonetheless independent of each
 14 other.”). However, upon review, TEFLabs’s description of its best mode argument in its
 15 interrogatory response arguably suggests enough of a written description argument⁷ that the court
 16 declines to strike that theory on the technical grounds that it was not previously disclosed.

17 **2. Merits of Written Description Invalidation Defense**

18 The written description requirement is set forth in the first paragraph of 35 U.S.C. § 112.
 19 In pertinent part, Section 112 ¶ 1 provides that:

20 The specification shall contain a written description of the invention,
 21 and of the manner and process of making and using it, in such full,
 22 clear, concise, and exact terms as to enable any person skilled in the
 23 art to which it pertains, or with which it is most nearly connected, to
 24 make and use the same, and shall set forth the best mode
 25 contemplated by the inventor of carrying out his invention.

26 35 U.S.C. § 112 ¶ 1. *See also Novozymes A/S v. DuPont Nutrition Biosciences APS*, 723 F.3d
 27 1336, 1344 (Fed. Cir. 2013) *cert. denied*, 134 S. Ct. 1501 (2014).

28 To satisfy the written description requirement, “the description must clearly allow persons
 of ordinary skill in the art to recognize that the inventor invented what is claimed.” *Ariad Pharm.*,

⁷ *See, e.g.*, Carter Decl. at H at 18 (“The claimed compound was not *described* in any of the 34 examples”) (emphasis added).

1 *Inc. v. Eli Lilly & Co.*, 598 F.3d 1336, 1351 (Fed. Cir. 2010). In other words, “the test requires an
2 objective inquiry into the four corners of the specification from the perspective of a person of
3 ordinary skill in the art. Based on that inquiry, the specification must describe an invention
4 understandable to that skilled artisan and show that the inventor actually invented the invention
5 claimed.” *Id.* Because the specification is viewed from the perspective of one of skill in the art, a
6 patentee may rely on information that is “well-known in the art” for purposes of meeting the
7 written description requirement. *See Falko–Gunter Falkner v. Inglis*, 448 F.3d 1357, 1366-68
8 (Fed. Cir. 2006). “The written description inquiry presents an issue of fact.” *Novozymes*, 723
9 F.3d at 1344 (citations omitted). *See also Union Oil Co. of California v. Atl. Richfield Co.*, 208
10 F.3d 989, 1001 (Fed. Cir. 2000) (“[W]ritten description questions are intensely factual, and should
11 be dealt with on a case-by-case basis, without the application of wooden rules.”).

12 TEFLabs failed to present any evidence relevant to this legal standard. TEFLabs did not
13 address or provide evidence relating to: (1) the level of a person of ordinary skill in the art⁸; (2) the
14 nature of the invention claimed; (3) what a person of ordinary skill in the art would have
15 understood based on the disclosure within the four corners of the ‘165 Patent specification; or (4)
16 what that person would have understood based on what was well-known in the art. TEFLabs
17 presented only attorney argument to suggest that a person of ordinary skill in the art would not
18 have understood the claimed invention to be adequately described.

19 Indeed, TEFLabs’s briefs rely mostly on attorney argument, as they include virtually no
20 citations to legal authority, and only minimal citations to mostly undifferentiated masses of
21 evidence.⁹ It is not the court’s task to pan through these exhibits in an effort to discover

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23 ⁸ “Factors that may be considered in determining level of ordinary skill in the art include: (1) the
24 educational level of the inventor; (2) type of problems encountered in the art; (3) prior art
25 solutions to those problems; (4) rapidity with which innovations are made; (5) sophistication of
26 the technology; and (6) educational level of active workers in the field. These factors are not
27 exhaustive but are merely a guide to determining the level of ordinary skill in the art.” *Daiichi*
28 *Sankyo Co. v. Apotex, Inc.*, 501 F.3d 1254, 1256 (Fed. Cir. 2007).

⁹ TEFLabs attached over 1,300 pages of exhibits to its opening brief and 65 pages of exhibits to
its reply. The following sentence from the opening brief is typical of TEFLabs’s tendency to cite
to a mass of evidence in support of its attorney’s conclusory argument: “In the ‘683 Application,
AAT Bioquest submitted deceptive affidavits. 12932683_FILE_HISTORY.” DMSJ at 45. The
file history for the ‘683 Application is 427 pages long.

1 TEFLabs’s argument. *See Digital Reg of Texas, LLC v. Adobe Sys., Inc.*, No. C 12-1971 CW,
2 2014 WL 3883437, at *1 (N.D. Cal. Aug. 6, 2014) (denying defendant’s motion for
3 reconsideration of judge’s summary judgment order for plaintiff; “Adobe must explain its
4 summary judgment arguments and cannot rely on the Court to sift through the countless exhibits
5 to manufacture a summary judgment argument.”); *Lockformer Co. v. PPG Indus., Inc.*, No. 99-C-
6 6799, 2003 WL 1563703, at *2 n. 1 (N.D. Ill. Mar. 25, 2003) *aff’d*, 138 Fed. Appx. 314 (Fed. Cir.
7 2005) (“It is not the Court’s task to search through the record to find evidence that supports PPG’s
8 position. Accordingly, the Court will not consider PPG’s unsupported assertions and anticipated
9 testimony in ruling on this motion [for summary judgment]”).

10 TEFLabs’s failure to provide any evidence or argument regarding what constitutes
11 “persons of ordinary skill in the art” dooms its motion for summary judgment on the written
12 description requirement. *See Suffolk Technologies, LLC v. AOL Inc.*, 752 F.3d 1358, 1367 (Fed.
13 Cir. 2014) (granting summary judgment, and holding that “[w]ithout expert testimony” or other
14 “affirmative evidence,” “mere attorney argument” was insufficient to undermine credible
15 testimony from defendant’s expert). Without this evidence, the court cannot assess the most basic
16 elements of the written description requirement, i.e., whether the description “clearly allow[s]
17 persons of ordinary skill in the art to recognize that the inventor invented what is claimed.”

18 In contrast, AAT’s request for a judgment of “no invalidity” on the basis of written
19 description includes competent expert testimony. AAT’s expert Dr. Wayne Patton defines a
20 person of ordinary skill in the art of fluorescent ion indicators as “a senior graduate student,
21 postdoctoral fellow or practicing Master’s degree or Ph.D. level scientist trained in organic
22 chemistry who is familiar with the synthesis, properties and biological application of ion indicator
23 dyes and might be engaged in practical research at a university, research institute, government
24 laboratory or in industry.” *See Patton Decl.* at ¶ 25. The basis for this definition is the doctorate-
25 level education of ’165 Patent inventor Dr. Diwu; the types of problems and prior art solutions
26 encountered in the art; the relatively slow pace of innovation in the field; and the difficulty of
27 identifying new chemical structures with improved qualities out of billions of possibilities. *Id.* at
28 ¶ 26. Dr. Patton points to specific support in the relevant patent applications and concludes that

1 “[b]ased on [his] review, a person of ordinary skill in the art at the time of the invention would
2 have understood the claimed compound to be adequately supported in both the provisional and
3 utility applications.” Patton Surreply Decl. [Docket No. 39-1] at ¶ 9. Based on this evidence,
4 AAT concludes that a person of ordinary skill in the relevant art would have understood that the
5 ’165 Patent claimed Fluo-8 AM.

6 In response, rather than address Dr. Patton’s testimony, TEFLabs offers two reasons why
7 this court should ignore the “person of ordinary skill in the art” standard.¹⁰ First, TEFLabs
8 contends that “the test for written description is whether the structure for the compound of claim 1
9 was presented in the specification.” See Reply at 3. According to TEFLabs, any lay person can
10 determine that the written description requirement is not met here, because the structure for the
11 compound is not present in the specification. This is not the law. The standard is not whether a
12 specific chemical structure is set forth in the patent specification, but whether a person of ordinary
13 skill in the art could read the patent specification and recognize that the inventor claimed what was
14 invented. See *Union Oil*, 208 F.3d at 1001 (“The written description requirement does not require
15 identical descriptions of claimed compounds, but it requires enough disclosure in the patent to
16 show one of skill in this art that the inventor ‘invented what is claimed.’”; finding substantial
17 evidence of adequate written description in a patent for gasoline products in which the claims “do
18 not describe each gasoline product in terms of molecular structures or lists of ingredients” but
19 instead “specify the chemical properties of the gasolines”); *Ex Parte Sorenson*, 3 U.S.P.Q.2d 1462
20 (B.P.A.I. May 28, 1987) (an “appellant’s specification need not describe the claimed invention in
21 *ipsis verbis* to comply with the written description requirement”).

22 Second, TEFLabs makes a convoluted argument that “[n]o ‘person of ordinary skill’
23 approach is required in this case to determine that . . . the omission of the compound from the
24 provisional and two non-provisional applications was deliberate; that Plaintiff had canceled the

25 ¹⁰ At the hearing, TEFLabs cited *In re Ruschig*, 54 C.C.P.A. 1551 (1997) in support of its
26 argument that it was unnecessary to follow the “person of ordinary skill in the art” standard.
27 However, contrary to TEFLabs’s contention, the *Ruschig* court did apply that standard to
28 determine whether the chemical compound of the claim was sufficiently described therein. *Id.* at
1558-59 (“The issue here . . . is a question of fact: Is the compound of claim 13 described therein?
Does the specification convey clearly to those skilled in the art, to whom it is addressed, in any
way, the information that appellants invented that specific compound?”).

1 claim for the compound and admitted that it was not in the specification; that no written
2 description was ever provided for the claimed compound; that the structure format was changed;
3 that Plaintiff misrepresented [sic] structure 284G for the required written description; and that
4 claim 1 could not have overcome the previously asserted written description rejection without
5 misrepresentation and concealment.” Reply at 4. As best as the court can decipher, TEFLabs
6 argues as follows: Claim 26 of the ’683 Application claimed the same compound that is in Claim 1
7 of the ’165 Patent; the patent examiner rejected Claim 26 for inadequate written description; the
8 Applicants “admitted” to the examiner that the compound was not in Claim 26 and canceled Claim
9 26, thereby canceling the subject matter of Claim 26; but then the Applicants reintroduced the
10 same compound that was the subject of the rejected Claim 26 by “relying on a false structure” in
11 the newly-added Claim 33.

12 Setting aside the fundamental defect in TEFLabs’s argument—that no authority permits
13 TEFLabs to ignore the “person of ordinary skill” standard—this perplexing argument is also
14 flawed. It appears to be directed primarily toward TEFLabs’s inequitable conduct defense,
15 discussed below. To the extent that it is a written description defense, TEFLabs seems to be
16 saying that Claim 1 of the ’165 Patent does not meet the written description requirement because
17 Claim 26 of the ’683 Application did not meet the written description requirement, and the two
18 claims are the same. However, TEFLabs does not explain how the two claims are the same, or
19 why Claim 26 failed to meet the written description requirement in the first instance, nor does it
20 cite to any specific evidence to shed light on these conclusions.¹¹ Without this information, the
21 court cannot divine why TEFLabs believes Claim 1 fails to meet the written description
22 requirement. Defendant’s argument succumbs for that reason.¹²

23 ¹¹ TEFLabs cites only isolated “page numbers” within the *unpaginated* ’683 Application file, with
24 no further explanation or argument about why that portion of the application file supports
TEFLabs’s argument.

25 ¹² Even so, AAT has provided competent expert evidence explaining how each of TEFLabs’s
26 nested conjectures is incorrect. First, AAT’s expert found “no evidence [in the patent history] that
27 the Applicants represented in any way that the compound was unsupported by the specification or
28 that they cancelled the subject matter of claim 1 of the ’165 patent.” *See* Patton Surreply Decl. at
¶ 8. Instead, “Claims 22 and 24-26 were cancelled for simplicity, but the subject matter of the
issued claim remained encompassed by Claims 21 and 23.” *Id.* *See also* DMSJ at Ex. 12
(Amendment and Response to Accompany Request for Prioritized Examination dated August 28,

1 In sum, TEFLabs’s written description arguments are hopelessly flawed for multiple
2 reasons. Because TEFLabs has failed to provide clear and convincing evidence on any element of
3 its written description defense such that a reasonable jury could invalidate the patent, TEFLabs’s
4 motion for summary judgment of invalidity on the basis of written description is **denied**. AAT’s
5 motion for summary judgment of no invalidity on the basis of written description is **granted**. *See*
6 *Eli Lilly*, 251 F.3d at 962 (“[A] moving party seeking to have a patent held not invalid must show
7 that the nonmoving party, who bears the burden of proof at trial, failed to produce clear and
8 convincing evidence on an essential element of a defense upon which a reasonable jury could
9 invalidate the patent.”).

10 **B. Enablement Requirement**

11 A patent specification must enable a person of ordinary skill in the art to make and use the
12 invention. 35 U.S.C. § 112 ¶ 1. “This requirement is met when at the time of filing the
13 application one skilled in the art, having read the specification, could practice the invention
14 without undue experimentation.” *Cephalon, Inc. v. Watson Pharm., Inc.*, 707 F.3d 1330, 1336
15 (Fed. Cir. 2013) (citation omitted).

16 TEFLabs originally challenged the validity of the ’165 Patent as failing to meet the
17 enablement requirement. DMSJ at 11. TEFLabs then appeared to have abandoned this argument,
18 as it did not respond to AAT’s counter-arguments or even mention enablement in its Reply. *See*

19 2013) (“[Claim 26 is] canceled . . . Claim 33 specifies a compound, support for which is found in
20 Claim 23 and throughout the specification, see e.g. compound 284, example 14, page 83.”).
21 Second, AAT’s expert found no evidence that the examiner relied on or accepted a “false”
22 structure. *See* Patton Surreply Decl. at ¶ 9 (“TEFLabs alleges that the Applicants relied on a
23 ‘false’ structure (284G) to establish adequate written description for the Fluo-8 AM compound of
24 Claim [1] . . . I have reviewed and compared the disclosures made in the provisional and utility
25 patent applications and I disagree with TEFLabs’ conclusion that the compound was somehow
26 falsified . . . [T]here is no evidence or suggestion that the cited structures in the utility application
27 were hidden in any way. Rather the making and use of the structures were extensively described
28 and supported, and a person of ordinary skill in the art would have understood this. As such, there
can be no deception, because these structural changes were supported by the extensive synthetic
details which show one of ordinary skill how to make such compounds.”). Finally, AAT’s
evidence supports that to the extent there was a typographical error in the 284G structure depicted
in the published application, that error bears no relation to TEFLabs’s argument regarding the
adequacy of the patent’s written description. *See* Patton Decl. at ¶¶ 54-55 (clerical error in the
published application was immaterial to applicants’ reliance on the 284G structure; applicants
relied on 284G structure to show support for the H group at the K position of claim 33, and the
typographical error was unrelated to that substitution at position K; typo was also unrelated to the
addition of AM esters to the compound).

1 *Collins v. City of San Diego*, 841 F.2d 337, 339 (9th Cir. 1988) (“It is well established in this
2 Circuit that claims which are not addressed in the . . . brief are deemed abandoned.”) (citation
3 omitted). At the hearing, TEFLabs confirmed that it had dropped its enablement defense.

4 Nevertheless, AAT moves affirmatively for a summary judgment of no invalidity on the
5 basis of enablement. To that end, AAT has provided competent and uncontroverted expert
6 evidence explaining how different portions of the ’165 Patent specification would assist a person
7 to make and use the compound of Claim 1 without undue experimentation. First, Figures 6 and 7
8 of the specification disclose two methods of synthesis of a fluo indicator with AM esters and
9 indicate that the substituents may be varied. Then, the specification at columns 57-60 provides
10 synthesis details for arriving at a compound that differs from Fluo-8 AM by a single substituent
11 (i.e., the substituent at position K). Finally, example 14 of the specification provides detailed
12 disclosure for how to generate a similar molecule with that substituent that is the same as in Fluo-8
13 AM (i.e., with a hydrogen in place of a methyl group at position K). Patton Decl. at ¶¶ 31-32;
14 ’165 Patent at col. 77-80.

15 AAT has shown that TEFLabs failed to demonstrate by clear and convincing evidence that
16 the ’165 Patent does not meet the enablement requirement. AAT has also provided uncontradicted
17 evidence as to why the ’165 Patent does meet the enablement requirement. As such, AAT’s
18 motion for summary judgment of no invalidity on the basis of enablement is **granted**.

19 **C. Anticipation**

20 **1. Legal Standard**

21 If the claimed invention was “described in a printed publication” either before the date of
22 invention, or more than one year before the U.S. patent application was filed, then that prior art
23 anticipates the patent. 35 U.S.C. § 102(a)-(b). The anticipation inquiry proceeds on a claim-by-
24 claim basis. *Finisar Corp. v. DirecTV Grp., Inc.*, 523 F.3d 1323, 1334 (Fed. Cir. 2008). That is,
25 “[t]o anticipate a claim, a single prior art reference must expressly or inherently disclose each
26 claim limitation. But disclosure of each element is not quite enough—this court has long held that
27 anticipation requires the presence in a single prior art disclosure of all elements of a claimed
28 invention arranged as in the claim.” *Id.* (citations omitted).

2. **Differences Between the Invention Claimed in Tsien Patent and '165 Patent**

TEFLabs alleges that the Tsien Patent (through its disclosure of Fluo-2) discloses the compound of Claim 1 of the '165 Patent (i.e., Fluo-8 AM). The patent examiner expressly considered and rejected this argument. *See* Carter Decl. at Ex. C (Notice of Allowability of '165 Patent, dated May 15, 2014) at 3. Understanding the argument requires some background on the teachings of the Tsien Patent.

The Tsien Patent claims a genus of calcium indicators. It expressly discloses the chemical structures for five calcium ion indicators: the rhodamine calcium ion indicators Rhod-1 and Rhod-2, and the fluo calcium ion indicators Fluo-1, Fluo-2, and Fluo-3.¹³ Sometime after the Tsien Patent was issued, another fluo calcium ion indicator called “Fluo-4” was introduced to the market.¹⁴ According to AAT, by 2007, “the most well-known and most oft used fluo indicators for intracellular calcium detection were Fluo-3 and Fluo-4.” Patton Decl. at ¶ 26(j). As stated above, AAT sells a fluorescent calcium ion indicator called Fluo-8 AM,¹⁵ which is an embodiment of Claim 1 of the '165 Patent.

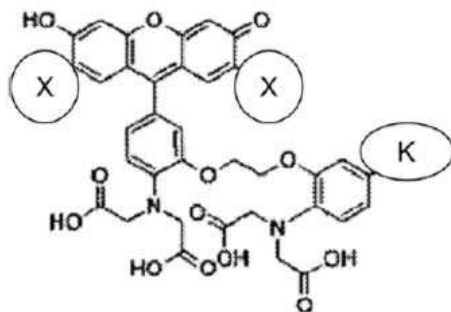
The compounds in the genus of indicators disclosed in the Tsien Patent share a generic formula. The base structure for the disclosed fluo acid compounds is below:

¹³ The three disclosed fluo indicators do not have AM esters (and are therefore sometimes referred to as Fluo-1 acid, Fluo-2 acid, or Fluo-3 acid). *Id.* at ¶ 34.

¹⁴ It is disputed whether Fluo-4 was claimed in the Tsien Patent or whether Fluo-4 is the embodiment of a patent owed by a different entity. *Compare* Minta Decl. [Docket No. 36-5] at ¶ 10 (“Molecular Probes made Fluo-4, a compound claimed in the Tsien Patent called Fluo-4, and paid royalties to UC Berkeley [the assignee of the Tsien Patent.]” *with* Patton Decl. at ¶ 26(l) (Fluo-4 was patented in 1996) *and* Surreply at 1-2 (“In 2001-2001, TEFLabs willfully infringed Molecular Probes’ U.S. Patent No. 6,162,931 (the '931 Patent) directed to Fluo-4 (the next generation fluo indicator after Tsien’s disclosures of Fluo-2 and Fluo-3). *Molecular Probes v. Tex. Fluorescence Labs, Inc.*, No. 3:02-cv-461 (N.D. Cal.) . . . [T]he case settled with TEFLabs taking a license to the '931 Patent in order to have the right to continue to sell the otherwise infringing Fluo-4 product.”). For purposes of this motion, it is irrelevant who invented Fluo-4; what matters is that Fluo-4 was popular in the market before Fluo-8 was introduced, which the parties do not dispute.

¹⁵ AAT named this compound Fluo-8 AM ostensibly because it was twice as bright as Fluo-4. Diwu Decl. at ¶ 11.

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The circled Xs and K represent sites at which variable substituents¹⁶ may be found. For example, the Fluo-2 molecule has hydrogen (H) at position X and a methyl group (CH₃) at position K, whereas the Fluo-3 molecule has chlorine (Cl) at position X and a methyl group at position K:

Molecule	X position	K position
Fluo-2	H	CH ₃
Fluo-3	Cl	CH ₃

By TEFLabs’s own admission, there are at least two differences between the structure of the Fluo-8 AM claimed in the ’165 Patent and the Fluo-2 compound disclosed in the Tsien Patent: (1) the Fluo-8 AM molecule has hydrogen at the K position; and (2) Fluo-8 AM has AM esters. See DMSJ at 39 (“The only two differences between [Fluo-8 AM] [and] the Fluo-2 example presented in the Tsien patent and the Minta paper are the acetoxymethyl (‘AM’) ester form and the ‘H’ versus ‘CH₃’ substituent at position ‘k’”); Reply at 6 (“There were only 2 differences between the compound of claim 1 and the Fluo-2 example in the Tsien patent - AM esters and the affinity substituent k=H rather than CH₃”). In fact, the patent examiner’s “no anticipation” finding was based on one of these admitted differences; namely, that Fluo-8 was not anticipated by Fluo-2 because the latter had a methyl group at the K position. See Carter Decl. at Ex. C at 2-3 (Fluo-8, as the elected compound in claim 33, “is not the . . . (AM) ester of the compound fluo-2 of Tsien.

¹⁶ A substituent is an atom or a group of atoms substituted in the place of a hydrogen atom on a parent chain of a hydrocarbon. Patton Decl. at ¶ 23(g).

1 The compound in canceled claim 34 is the AM ester of fluo-2. The compounds differ by a methyl
2 group. This compound is not anticipated.”).

3 TEFLabs’s admission alone is fatal to its motion for summary judgment, because an
4 anticipation defense requires TEFLabs to show that the Tsien Patent “expressly or inherently
5 disclose[s] . . . all elements of a claimed invention arranged as in the claim,” *Finisar Corp.*, 523
6 F.3d at 1334, but TEFLabs has not identified where in the Tsien Patent the two admitted
7 differences identified above were disclosed.

8 **3. Disclosure of Genus**

9 TEFLabs also contends that the Tsien Patent anticipated Fluo-8 because it claims a genus
10 of indicators, a species of which is Fluo-8. However, “[i]t is well established that the disclosure of
11 a genus in the prior art is not necessarily a disclosure of every species that is a member of that
12 genus.” *Atofina v. Great Lakes Chem. Corp.*, 441 F.3d 991, 999 (Fed. Cir. 2006). “There may be
13 many species encompassed within a genus that are not disclosed by a mere disclosure of the
14 genus. On the other hand, a very small genus can be a disclosure of each species within the
15 genus.” *Id.*

16 According to AAT’s expert, Claim 1 of the Tsien Patent claims a broad genus of calcium
17 indicators. Patton Decl. at ¶¶ 39-40. The genus shares a generic formula, but includes 11 sites,¹⁷
18 each of which has up to eight substituent options; the permutations of these substituents means the
19 Tsien Patent encompasses over 20 billion compounds with “dramatically different physical and
20 functional properties, with no guidance as to those with superior cell loading ability and
21 brightness.” *Id.* at ¶¶ 40-41. Of these possible permutations, the Tsien Patent discloses only five
22 specific embodiments. *Id.* at ¶ 42. Thus, AAT argues, one of ordinary skill would not be able to
23 envision the billions of other species encompassed within this vast genus, much less the particular
24 Fluo-8 AM species that is the compound of claim 1 of the ‘165 Patent. *Id.* at ¶ 45. *Accord Sanofi-*
25 *Synthelabo v. Apotex, Inc.*, 550 F.3d 1075, 1084 (Fed. Cir. 2008) (affirming district court’s
26 holding of no anticipation where “the description of the genus would not lead a person of ordinary
27 skill to a ‘small recognizable class with common properties’”) (citation omitted).

28 ¹⁷ The sites are designated as E1, E2, W, X, Q1, Q2, Y, Z, Z1, Z3, and Z4 in the claim.

1 TEFLabs responds that Patton’s estimate of the number of possible compounds claimed in
2 the Tsien Patent is exaggerated. According to TEFLabs, the practice in the industry is to keep the
3 substituents at certain sites fixed or limited to a smaller subset of options. *See* Reply at 5. Thus,
4 rather than containing billions of possibilities, the genus claim in the Tsien Patent was in reality
5 restricted to only “36 possible combinations of substituents.” *Id.* In support of this position,
6 TEFLabs cites to a table in the declaration of Dr. Minta ostensibly showing 18 fluo indicators that
7 were commercially available in 2006. *See* Minta Decl. at ¶ 18, Table 3 (referring to information in
8 DMSJ Ex. 7 (Simpson (2006) article entitled “Fluorescent Measurement of [Ca²⁺]”).

9 The underpinnings of TEFLabs’s argument are flawed. There is no suggestion that the 18
10 listed indicators were the only ones on the market, nor that the products available in 2006 were
11 exemplary of all fluo indicators that had been on the market since the issuance of the Tsien Patent.
12 Thus, on its face, TEFLabs’s cultivated list of selected products is insufficient to show that in
13 practice, the genus claim of the Tsien Patent encompassed only a limited number of species. Next,
14 AAT’s expert directs the court to examples of indicator products with changes at the variable
15 positions that TEFLabs claims were “fixed” in industry practice. *See* Patton Surreply Decl. at ¶ 15
16 (“I disagree with TEFLabs’ contention that there were only 36 possible combinations . . . at the
17 time of the invention in 2007 or that industry practice fixed 7 . . . of the 11 variable positions in the
18 Tsien claim. In fact, TEFLabs itself marketed products as early as 1998 with changes to at least
19 three of these variables It is my understanding that all of the variable positions of the Tsien
20 patent have been pursued by many chemists, including Dr. Minta himself.”). AAT also disputes
21 whether TEFLab’s list is accurate. AAT asserts that TEFLabs's list includes calcium indicators
22 (“Oregon Green” and “Calcium Green”) that are not fluo indicators. *See* Patton Surreply Decl. at
23 ¶ 18 (“[T]he Oregon Green BAPTA and Calcium Green indicators are not fluo indicators, so my
24 understanding that no one made a commercial intracellular calcium indicator with [hydrogen at the
25 K position] until Dr. Diwu and his co-inventors did . . . still stands. Moreover, nothing about these
26 different and distinct indicators would have prompted one of ordinary skill in the art to look at
27 them in order to solve the problems that existed with fluo calcium ion indicators at the time of the
28 invention in 2007.”).

1 In sum, AAT has shown that TEFLabs failed to demonstrate by clear and convincing
2 evidence that the '165 Patent was invalid because it was anticipated by prior art. For the above
3 reasons, TEFLabs's motion for summary judgment on anticipation is **denied**, and AAT's motion
4 for summary judgment of no anticipation is **granted**.

5 **D. Obviousness**

6 **1. Legal Standard**

7 The Patent Act forbids issuance of a patent when "the differences between the subject
8 matter sought to be patented and the prior art are such that the subject matter as a whole would
9 have been obvious at the time the invention was made to a person having ordinary skill in the art
10 to which said subject matter pertains." 35 U.S.C. § 103.

11 "Generally, a party seeking to invalidate a patent as obvious must demonstrate by clear and
12 convincing evidence that a skilled artisan would have had reason to combine the teaching of the
13 prior art references to achieve the claimed invention, and that the skilled artisan would have had a
14 reasonable expectation of success from doing so." *In re Cyclobenzaprine Hydrochloride*
15 *Extended-Release Capsule Patent Litig.*, 676 F.3d 1063, 1068-69 (Fed. Cir. 2012). "[W]hile an
16 analysis of any teaching, suggestion, or motivation to combine known elements is useful to an
17 obviousness analysis, the overall obviousness inquiry must be expansive and flexible." *Id.* at 1069
18 (citing *KSR Int'l Co. v. Teleflex, Inc.*, 550 U.S. 398, 419 (2007)).

19 Thus, to resolve the issue of obviousness, the court considers factual questions including:
20 (1) the scope and content of the prior art; (2) the level of ordinary skill in the pertinent art; and (3)
21 differences between the claimed invention and the prior art. *KSR*, 550 U.S. at 406. The court may
22 also consider "secondary considerations," such as "commercial success, long felt but unsolved
23 needs, [and] failure of others," to "give light to the circumstances surrounding the origin of the
24 subject matter sought to be patented." *Id.* (quoting *Graham v. John Deere Co.*, 383 U.S. 1, 17-18
25 (1966)). "Another indicia of non-obviousness of a product is the acclamations it receives when it
26 is released, and the copying that occurs," although "a showing of copying is only equivocal
27 evidence of non-obviousness in the absence of more compelling objective indicia of other
28 secondary considerations." *Ecolchem, Inc. v. S. California Edison Co.*, 227 F.3d 1361, 1380

1 (Fed. Cir. 2000). The secondary evidence of nonobviousness is “often the most probative and
2 determinative of the ultimate conclusion of obviousness or nonobviousness.” *Pro-Mold and Tool*
3 *Co., Inc. v. Great Lakes Plastics, Inc.*, 75 F.3d 1568, 1573 (Fed. Cir. 1996).

4 “Where a skilled artisan merely pursues ‘known options’ from a ‘finite number of
5 identified, predictable solutions,’ the resulting invention is obvious under Section 103.” *In re*
6 *Cyclobenzaprine Hydrochloride*, 676 F.3d at 1070 (*quoting KSR*, 550 U.S. at 421). “Where,
7 however, a defendant urges an obviousness finding by merely throwing metaphorical darts at a
8 board in hopes of arriving at a successful result, but the prior art gave either no indication of which
9 parameters were critical or no direction as to which of many possible choices is likely to be
10 successful, courts should reject hindsight claims of obviousness.” *Id.* (quotations omitted).

11 2. Examiner’s Determination of Non-Obviousness

12 “[A] party challenging [a presumptively valid patent] shoulders an enhanced burden if the
13 invalidity argument relies on the same prior art considered during [the patent] examination.”
14 *Tokai*, 632 F.3d at 1367.

15 The patent examiner, in granting the ’165 Patent, withdrew its previous finding of
16 obviousness on the basis of Fluo-8’s unexpected results.¹⁸ Carter Decl. at Ex. C at 3. In doing so,
17 the examiner looked at the same prior art (i.e., the Tsien Patent and the fluo indicators disclosed
18 therein) on which TEFLabs’s obviousness argument relies. Specifically, the examiner stated that
19 “H to Me [hydrogen to methyl group] analogs are generally *prima facie* obvious” but found “the
20 affidavit of inventor Diwu filed on August 28, 2013 . . . sufficient to rebut such a rejection on the
21 grounds of unexpected results.” *Id.* The examiner finding of non-obviousness focused on the fact
22 that Fluo-8 performed better than other previously-known fluo indicators disclosed in the Tsien
23 Patent. *See id.* (“Since the compound . . . is better than the Tsien compounds in various calcium
24 imaging assays . . . any possible 103(a) [obviousness] rejection over Tsien for this compound has

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26 ¹⁸ The doctrine of unexpected results is based on the principle that “that which would have been
27 surprising to a person of ordinary skill in a particular art would not have been obvious. The
28 principle applies most often to the less predictable fields, such as chemistry, where minor changes
in a product or process may yield substantially different results.” *In re Soni*, 54 F.3d 746, 750
(Fed. Cir. 1995) (reversing obviousness rejection where applicant presented evidence that
compound with increased molecular weight produced dramatically superior results).

1 been overcome.”). The patent examiner also noted that Fluo-2 was an unlikely lead compound.
2 Carter Decl. at Ex. C at 3 (“[T]he fluo-2 compound was described by Tsien as the poorest
3 performer in the patent specification . . . which gives little motivation to select this compound as
4 one for further manipulation, i.e. demethylation.”).

5 Of course, the fact that the examiner determined that Claim 1 was not obvious does not
6 alone resolve the question. The court thus turns to the parties’ arguments.

7 3. *Prima Facie Obviousness*

8 TEFLabs contends that “the compound of claim 1 is an obvious combination of prior art
9 elements.” DMSJ at 40. However, this argument fails to cite to any evidence regarding what a
10 person of ordinary skill in the art would have understood at the time of patent filing. This reason
11 alone is sufficient to deny TEFLabs’s motion for summary judgment. *Advanced Media Networks*
12 *LLC v. Row 44 Inc.*, No. CV 12-11018 GAF JCGX, 2014 WL 5623951, at *5 (C.D. Cal. Nov. 4,
13 2014) (“[B]ecause [defendants’] motion [for summary judgment] lacks the most basic information
14 regarding what a skilled artisan would have considered obvious at the time of the invention, the
15 argument cannot advance past that point and the Court therefore does not address the numerous
16 other flaws in the pending motion.”).

17 In any event, even if the court overlooks TEFLabs’s foundational failure, its obviousness
18 arguments are still inadequate. For patents that claim new chemical compounds, the question of
19 “*prima facie* obviousness . . . generally turns on the structural similarities and differences between
20 the claimed compound and the prior art compounds.” *Otsuka Pharm. Co., Ltd. v. Sandoz Inc.*, 678
21 F.3d 1280, 1291 (Fed. Cir. 2012) (citation omitted). Whether a new chemical compound would
22 have been *prima facie* obvious follows a two-part inquiry. *Id.* “First, the court determines
23 whether a chemist of ordinary skill would have selected the asserted prior art compounds as lead
24 compounds, or starting points, for further development efforts.” *Id.* Second, the court determines
25 “whether the prior art would have supplied one of ordinary skill in the art with a reason or
26 motivation to modify a lead compound to make the claimed compound with a reasonable
27 expectation of success.” *Id.* at 1292 (citation omitted).

28 With respect to the first step of the inquiry, a lead compound is “a compound in the prior

1 art that would be most promising to modify in order to improve upon its . . . activity and obtain a
2 compound with better activity.” *Id.* at 1291 (citation omitted). “[A] lead compound is a natural
3 choice for further development efforts.” *Id.* “In determining whether a chemist would have
4 selected a prior art compound as a lead, the analysis is guided by evidence of the compound’s
5 pertinent properties.” *Id.* at 1292 (citation omitted).

6 Such properties may include positive attributes such as activity and potency; adverse
7 effects such as toxicity, and other relevant characteristics in evidence. *See Eisai v. Dr. Reddy’s*
8 *Labs, Ltd.*, 533 F.3d 1353, 1358 (Fed. Cir. 2008) (considering a prior art compound’s lipophilicity
9 and low molecular weight); *Pfizer, Inc. v. Apotex, Inc.*, 480 F.3d 1348, 1363 (Fed. Cir. 2007)
10 (considering the “strength, solubility, and other known chemical characteristics” of a prior art
11 saltforming acid). “Absent a reason or motivation based on such prior art evidence, mere
12 structural similarity between a prior art compound and the claimed compound does not inform the
13 lead compound selection.” *Otusakai*, 678 F.3d at 1292 (some citations omitted).

14 TEFLabs simply asserts that “Fluo-2 is a ‘lead compound’ (along with Fluo-1 and Fluo-3)
15 and is a logical starting point for investigating competitive alternatives to Fluo-4.” Reply at 6.
16 This is conclusory attorney argument with no evidence cited to support it.

17 In contrast, AAT has provided expert evidence explaining why Fluo-2 is not a lead
18 compound. Specifically, the Tsien Patent teaches that an electron withdrawing group is necessary
19 at the X position for a fluo calcium indicator to be useful for intracellular detection; otherwise, the
20 fluorescence signal of the indicator would be quenched. Patton Decl. at ¶ 36. Fluo-3 has an
21 electron-withdrawing group (Cl) at the X position, whereas Fluo-2 has a non-electron-
22 withdrawing group (H), so the Tsien Patent taught that Fluo-3 was a more desirable compound
23 than Fluo-2. *Id.* at ¶ 35. *See also* Tsien Patent at cols. 12:34-37 (“In most applications, fluo-3 will
24 be generally preferable over fluo-1 and fluo-2 because of its lesser sensitivity to pH and its larger
25 fluorescence enhancement on binding [calcium ion]”); 21:55-56 (“[F]luorescence of fluo-2 was
26 almost completely quenched as the pH was titrated from pH 7.7 to 4.1 in the absence of [calcium
27 ion]”); and 21:65-22:1 (“[B]ecause protonation has such a powerful effect on the fluorescence and
28 is spectrally indistinguishable from a drop in [calcium ion], fluo-2 is too pH sensitive for general

1 use.”). According to AAT, in developing their inventions, Dr. Diwu and his colleagues discovered
2 that it was not necessary to have an electron withdrawing group to make a useful calcium
3 indicator. Diwu Decl. ¶ 14.

4 Against this evidence, TEFLabs responds with declarations from Dr. Minta and Dr. Joseph
5 Kao, an expert with a doctorate in physical chemistry who was on Dr. Tsien’s research team at UC
6 Berkeley. *See* Reply at 6-7; Minta Decl. at ¶ 42; Kao Decl. [Docket No. 36-3] at ¶ 7. Dr. Minta
7 contends that the hydrogen at position X in Fluo-2 is actually “mildly electron withdrawing”; Dr.
8 Patton responds that “[t]his is scientifically incorrect” and that hydrogen is non-electron-
9 withdrawing. Patton Surreply Decl. at ¶ 22. Dr. Kao takes issue with the examiner’s reference to
10 Tsien’s statements that Fluo-2 was the “poorest performer.” Dr. Kao notes that “Roger Tsien is
11 extremely modest and cautious about the utility of his molecules” and simply because he
12 counseled against using Fluo-2 for *general use* did not mean he discounted the utility of Fluo-2
13 entirely. Kao Decl. at ¶ 8.

14 These quibbles miss the forest for the trees. While TEFLabs is busy attacking AAT’s
15 evidence showing that Fluo-2 was *not* a lead compound, it provides no affirmative evidence that
16 Fluo-2 was a lead compound. This means that TEFLabs has failed to meet its burden on summary
17 judgment, and that AAT has met its burden.

18 **4. Objective Indicia of Nonobviousness**

19 Finally, AAT points the court to evidence regarding objective indicia of non-obviousness.
20 First, the fact that TEFLabs copied AAT’s invention supports a finding of non-obviousness.
21 *Specialty Composites v. Cabot Corp.*, 845 F.2d 981, 991 (Fed. Cir. 1988). TEFLabs admitted that
22 a prospective customer asked whether it manufactured Fluo-8 AM in early 2010, and thereafter
23 TEFLabs “purchased Fluo-8 from AAT” and analyzed it. TEFLabs began selling Fluo-8 AM in
24 the second Q2 2010. Second, failures by others to create an improved calcium indicator and the
25 resulting unmet need are also strong objective indicia of non-obviousness. Diwu Decl. at ¶¶ 7-8
26 (efforts at Molecular Probes, Molecular Devices, and AnaSpec from 1993-2006 to address the
27 weakness of the known fluo indicators such as Fluo-3 and Fluo-4 were unsuccessful “in part
28 because [we] followed the Tsien reference’s teaching that focused on compounds with electron-

1 withdrawing groups at . . . the X position”). Third, the immediate commercial success of Fluo-8
2 AM is another strong indicator of nonobviousness. *See* Diwu Decl. at ¶ 17 (AAT’s total Fluo-8
3 AM revenue increased more than 100% year over year from 2007 to 2009, effectively creating a
4 new market segment); ¶ 18 (in 2008 and 2009, before the patent issued, three of market leaders in
5 the high throughput screening space sought out AAT and licensed the patent application which
6 eventually issued as the ‘165 Patent under no threat of litigation); at ¶ 17 (by 2010, AAT’s yearly
7 Fluo-8 AM sales totaled more than a half million dollars); ¶ 19 (three leading high throughput
8 screening companies have purchased Fluo-8 AM to specifically replace Fluo-3 AM or Fluo-4
9 AM); ¶ 17 (AAT achieved these sales without any salesperson in the field). Finally, TEFLabs’s
10 sale of its admittedly infringing copy of Fluo-8 AM, and its own sales materials extolling the
11 superiority of Fluo-8 over its predecessors, are indicia of non-obviousness. *See generally* Carter
12 Decl. at Exs. I, J, K, L, M, N.

13 TEFLabs responds to this with weak evidence of its own. First, as proof that Fluo-8 was
14 not successful, TEFLabs cites to a declaration from Dr. Minta that Fluo-8 “represents less than
15 10% of the fluo indicator market.” Minta Decl. at ¶ 15. But this conclusion rests on the thinnest
16 of reeds, for Dr. Minta based it on his review of “UC Berkeley receipts that I can locate” for sales
17 of Fluo-3 and Fluo-4 between 2003 and 2007, and general revenue reports comparing AAT and
18 TEFLabs’s revenues between 2007 and 2014. This does little to shed light on Fluo-8’s market
19 share and how it has changed over time. Second, TEFLabs argues that it simply opted not to
20 commercialized Fluo-8, even though the Tsien Patent ostensibly covered it, because it opted to
21 pursue other business strategies. This is belied by TEFLabs’s marketing materials, which explain
22 that Fluo-8 was commercialized *after* Fluo-3 and Fluo-4 because TEFLabs did not appreciate its
23 brightness until then. Carter Decl. at Ex. L at 19.

24 In sum, TEFLabs has failed at the outset to make a case of obviousness by not providing
25 evidence regarding what a person of ordinary skill in the art would have considered to be obvious
26 at the time of the invention. In addition, TEFLabs has failed to submit evidence demonstrating
27 why Fluo-2 would have been selected as a lead compound. In contrast, AAT has presented
28 evidence of the objective indicia of non-obviousness, as well as evidence that Fluo-8 AM

1 produced unexpectedly superior results when compared with the other fluo indicators available in
2 the market at that time. For these reasons, TEFLabs’s motion for summary judgment of invalidity
3 on the basis of obviousness is **denied**, and AAT’s motion for summary judgment of no invalidity
4 on the basis of obviousness is **granted**.

5 **E. Inequitable Conduct**

6 **1. Legal Standard**

7 Where a patent applicant breaches the duty to prosecute a patent application with candor
8 and good faith, it may result in a finding of inequitable conduct. 37 C.F.R. § 1.56(a) (2004);
9 *Purdue Pharma L.P. v. Endo Pharm. Inc.*, 438 F.3d 1123, 1128 (Fed. Cir. 2006). Inequitable
10 conduct is an equitable defense to patent infringement that, if proved, bars enforcement of a
11 patent. *Therasense, Inc. v. Becton, Dickinson & Co.*, 649 F.3d 1276, 1285 (Fed. Cir. 2011).
12 “[T]he remedy for inequitable conduct is the ‘atomic bomb’ of patent law.” *Id.* at 1288. “Unlike
13 validity defenses, which are claim specific . . . inequitable conduct regarding any single claim
14 renders the entire patent unenforceable.” *Id.* “[B]ecause the penalty for inequitable conduct is so
15 severe . . . [t]he need to strictly enforce the burden of proof and elevated standard of proof in the
16 inequitable conduct context is paramount.” *Star Scientific, Inc. v. R.J. Reynolds Tobacco Co.*, 537
17 F.3d 1357, 1365 (Fed. Cir. 2008).

18 To prove inequitable conduct, the accused infringer must present “evidence that the
19 applicant (1) made an affirmative misrepresentation of material fact, failed to disclose material
20 information, or submitted false material information, and (2) intended to deceive the [PTO].” *Star*
21 *Scientific*, 537 F.3d at 1365 (“The burden of proving inequitable conduct lies with the accused
22 infringer.”). Intent and materiality are separate requirements. “A district court should not use a
23 ‘sliding scale,’ where a weak showing of intent may be found sufficient based on a strong showing
24 of materiality, and vice versa. Moreover, a district court may not infer intent solely from
25 materiality. Instead, a court must weigh the evidence of intent to deceive independent of its
26 analysis of materiality.” *Therasense*, 649 F.3d at 1290. “[A] threshold level of each element—
27 i.e., both materiality and intent to deceive—must be proven by clear and convincing evidence.
28 And even if this elevated evidentiary burden is met as to both elements, the district court must still

1 balance the equities to determine whether the applicant’s conduct before the PTO was egregious
2 enough to warrant holding the entire patent unenforceable.” *Star Scientific*, 537 F.3d at 1365
3 (citations omitted). Thus, “even if a threshold level of both materiality and intent to deceive are
4 proven by clear and convincing evidence, the court may still decline to render the patent
5 unenforceable.” *Id.*

6 The intent element requires the accused infringer to “prove that the patentee acted with the
7 specific intent to deceive the PTO. A finding that the misrepresentation or omission amounts to
8 gross negligence or negligence under a ‘should have known’ standard does not satisfy this intent
9 requirement.” *Therasense*, 649 F.3d at 1290 (citation omitted). “Because direct evidence of
10 deceptive intent is rare, a district court may infer intent from indirect and circumstantial evidence.”
11 *Id.* “However, to meet the clear and convincing evidence standard, the specific intent to deceive
12 must be the single most reasonable inference able to be drawn from the evidence.” *Id.* (citations
13 omitted). “Indeed, the evidence must be sufficient to require a finding of deceitful intent in the
14 light of all the circumstances.” *Id.* (emphasis in original, quotation omitted). When there are
15 multiple reasonable inferences that may be drawn, intent to deceive cannot be found. *Id.* at 1290-
16 91. In a case involving nondisclosure of information, “the accused infringer must prove by clear
17 and convincing evidence that the applicant knew of the reference, knew that it was material, and
18 made a deliberate decision to withhold it.” *Id.* at 1290.

19 “The materiality required to establish inequitable conduct is but-for materiality.”
20 *Therasense*, 649 F.3d at 1291. “When an applicant fails to disclose prior art to the PTO, that prior
21 art is but-for material if the PTO would not have allowed a claim had it been aware of the
22 undisclosed prior art.” *Id.* “Hence, in assessing the materiality of a withheld reference, the court
23 must determine whether the PTO would have allowed the claim if it had been aware of the
24 undisclosed reference.” *Id.* “In making this patentability determination, the court should apply
25 the preponderance of the evidence standard and give claims their broadest reasonable
26 construction.” *Id.* at 1291-92.

27 “Determining at summary judgment that a patent is unenforceable for inequitable conduct
28 is permissible, but uncommon.” *Digital Control, Inc. v. Charles Mach. Works*, 437 F.3d 1309,

1 1313 (Fed. Cir. 2006). When a party has failed to established inequitable conduct by clear and
2 convincing evidence, summary judgment is properly granted against that party. *See Astrazeneca*
3 *Pharms. LP v. Teva Pharms. USA, Inc.*, 583 F.3d 766, 777 (Fed. Cir. 2009) (affirming summary
4 judgment of no inequitable conduct, when the factual premises could not be established by clear
5 and convincing evidence).

6 **2. TEFLabs’s Basic Evidentiary Failures**

7 TEFLabs makes conclusory, scattershot allegations of instances of inequitable conduct
8 supported by a handful of impenetrable evidentiary citations.¹⁹ These unsupported attorney
9 arguments are insufficient to meet TEFLabs’s burden. *See Digital Control*, 437 F.3d at 1313 (“A
10 genuine issue of material fact [regarding inequitable conduct] is not raised by the submission of
11 merely conclusory statements or completely insupportable, specious, or conflicting explanations
12 or excuses.”) (citations omitted).

13 Notwithstanding this evidentiary failure, TEFLabs’s arguments are still insufficient to the
14 extent they accuse “AAT Bioquest” or “Plaintiff” of engaging in inequitable conduct, because they
15 fail to demonstrate the specifics of the inequitable conduct. *See Exergen Corp. v. Wal-Mart*
16 *Stores, Inc.*, 575 F.3d 1312, 1328 (Fed. Cir. 2009) (“[T]o plead the ‘circumstances’ of inequitable
17 conduct with the requisite ‘particularity’ under Rule 9(b), the pleading must identify the specific
18 who, what, when, where, and how of the material misrepresentation or omission committed before
19 the PTO.”).

20 **3. TEFLabs’s Arguments Regarding Inequitable Conduct**

21 **a. Opportunities for Design Around Tsien Patent**

22
23 ¹⁹ Those are mass citations to the approximately 1,000 pages in the patent files of the ’452
24 Application, the ’753 Application, the ’683 Application, and two other patent applications that
25 TEFLabs contends are related to the ’165 Patent. *See, e.g.*, DMSJ at 45 (“In Base Application
26 ’753, AATBioquest falsified three compounds (284, 304, and 306) from the compounds identified
27 in the provisional application; and submitted new compounds 365 and 366 while still claiming the
28 benefit of the provisional application filing date. The falsified and new compounds are
fundamental to the claims in subsequent applications. [12040753_FILE_HISTORY].”). *See also*
supra n. 9. The only evidentiary citation for TEFLabs’s inequitable conduct arguments in its
Reply brief is to the declaration of Dr. Minta at paragraph 52. Reply at 8. For reasons explained
below, this *de minimis* evidence fails to meet TEFLabs’s elevated evidentiary burden to
demonstrate inequitable conduct by clear and convincing evidence.

1 to present the appearance of a legitimate research effort” and “claim[ed] a large number dummy
2 substituents at positions j, m, n, V, R3, and R4 without presenting any example of substituents at
3 those positions in any of the four utility patent applications”). Again, TEFLabs cites no evidence
4 in support of this argument. AAT’s evidence supports that these positions are not “dummy”
5 positions, and AAT has “commercialized and is pursuing patent protection on additional
6 compounds with different molecules at these alleged ‘dummy’ positions.” Diwu Decl. at ¶ 20.

7 **d. Deceptive naming and failure to disclose Fluo-2**

8 TEFLabs contends that AAT committed inequitable conduct by “not disclosing elected
9 species compound was a Fluo-2 compound until the examiner cited Fluo-2” and “renaming Fluo-2
10 AM as Fluo-8H, but not disclosing Fluo-8H (and its affinity relationship with Fluo-8 in the
11 affidavits).” Reply at 9-10. The patent file shows otherwise. See Carter Decl. at Ex. Q (Diwu
12 affidavit to patent examiner in ’683 Application dated March 2, 2011, including structural
13 comparison of Fluo-8 AM with Fluo-2 compounds).

14 **e. Falsifying compounds**

15 TEFLabs argues that AAT “falsif[ied] compounds 304, 306, and 284 by removing
16 halogens in order to backdate the priority of Fluo-2 versions to the provisional application.”
17 Reply at 10. *See also* DMSJ at 23-24 (example compounds 304 and 306 from the provisional
18 application were “changed . . . from fluorinated compounds to the preferred Fluo-2 compounds in
19 an application filed years later”). There is no evidence to support this argument. There is hardly
20 *argument* to support this argument, as the court cannot determine from TEFLabs’s briefing how it
21 believes compounds 304, 306, and 284 had been changed, how those changes qualify as
22 “falsifications,” when those falsifications were presented to the patent examiner, or how those
23 falsifications are material or significant. Moreover there is no evidence showing that these alleged
24 falsifications were made with the requisite intent to deceive.

25 In sum, TEFLabs has failed to present clear and convincing evidence supporting a finding
26 of inequitable conduct. TEFLabs’s motion for summary judgment of inequitable conduct is
27 **denied**, and AAT’s motion for summary judgment of no inequitable conduct is **granted**.

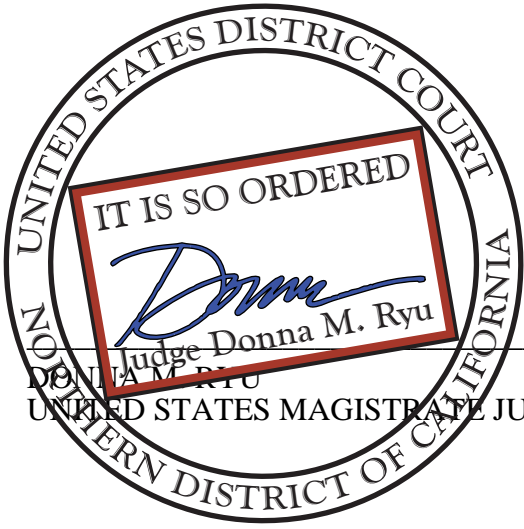
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IV. CONCLUSION

For the reasons stated above, AAT’s motion for summary judgment is **granted** and TEFLabs’s motion for summary judgment is **denied**. In light of this order, all pretrial and trial dates are vacated. The parties shall appear for a further case management conference on May 6, 2015, and shall file a joint updated case management conference statement by April 29, 2015.

IT IS SO ORDERED.

Dated: April 13, 2015



Judge Donna M. Ryu
UNITED STATES MAGISTRATE JUDGE